# Mplus Workshop, Part 1: Highlights from Muthén, Muthén \& Asparouhov (2016) Regression And Mediation Analysis Using Mplus 

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## Mplus Workshop: Overview of the Day

- 8:30-12:00: Highlights from the new book (Mplus Version 7.4)
- First morning block (Bengt, $11 / 2$ hours $^{1}$ ): Regression analysis
- Second morning block (Bengt, 1 1/2 hours): Mediation analysis
- Lunch: 12-1:30
- 1:30-6:30 (or longer): Time-series analysis (forthcoming Mplus Version 8)
- First afternoon block (Ellen, 1 1/2 hours): Introductory time-series analysis
- Second afternoon block (Ellen, 1 1/2 hours): Examples
- Third afternoon block (Tihomir, $11 / 2$ hours): Time-series implementation in Mplus Version 8

[^0]
## The Mplus User's Guide has Gotten a Companion

## Mplus

Statistical Analysis With Latent Variables
User's Guide

Regression And Mediation
Analysis Using Mplus

- 1. Linear regression analysis
- 2. Mediation analysis
- 3. Special topics in mediation analysis
- 4. Causal inference for mediation
- 5. Categorical dependent variable
- 6. Count dependent variable
- 7. Censored variable
- 8. Mediation with non-cont's variables
- 9. Bayesian analysis
- 10. Missing data

Table of Contents will be shown at www.statmodel.com. 500 pages. Lots of inputs and outputs. Paperback. All inputs and outputs will be posted. Most data sets will be posted. Perhaps assignments.

## Overview of the Morning: Highlights from the Book

- First morning block ( $11 / 2$ hours). Regression Analysis:
- Linear regression with an interaction
- Heteroscedasticity modeling
- Censored variable modeling: Tobit, censored-inflated, Heckman, and two-part analysis
- Bayes: Advantages over ML. Missing data on covariates
- Second morning block ( $11 / 2$ hours). Mediation Analysis:
- Moderated mediation with continuous mediator and outcome
- Monte Carlo simulation of moderated mediation
- Sensitivity analysis
- Mediation analysis using counterfactually-defined indirect and direct causal effects:
- Binary outcome
- Count outcome
- Two-part outcome

Note: The highlights skew toward the more advanced parts of the book to match the claim "Analyses you probably didnt know that you could do in Mplus".

## Example: Linear Regression with an Interaction



> Randomized field experiment in the Baltimore public schools where a classroom-based intervention aimed at reducing aggressive-disruptive behavior among elementary school students was carried out (Kellam et al., 2008)

- tx is a binary intervention variable
- agg1 is pre-intervention Grade 1 aggressive behavior score and agg5 the score in Grade 5
- txagg 1 is a treatment-baseline interaction (tx $\times$ agg1)


## Example: Linear Regression with an Interaction

$$
\begin{align*}
\operatorname{agg}_{i} & =\beta_{0}+\beta_{1} t x_{i}+\beta_{2} \operatorname{agg}_{i}+\beta_{3} \operatorname{txagg} 1_{i}+\varepsilon_{i} .  \tag{1}\\
\operatorname{agg}_{i} & =\beta_{0}+\beta_{1} \operatorname{tx}_{i}+\beta_{2} \operatorname{agg}_{i}+\beta_{3} \operatorname{tx}_{i} \operatorname{agg} 1_{i}+\varepsilon_{i}  \tag{2}\\
& =\beta_{0}+\beta_{2} \operatorname{agg}_{1}+\left(\beta_{1}+\beta_{3} \text { agg }_{i}\right) t x_{i}+\varepsilon_{i} . \tag{3}
\end{align*}
$$

The expression $\beta_{1}+\beta_{3} a g g 1$ is referred to as the moderator function or, when evaluated at a specific agg1 value, the simple slope. This means that agg 1 moderates the $\beta_{1}$ effect of tx on agg 5 by the term $\beta_{3}$ agg 1.

```
VARIABLE: USEVARIABLES = agg5 agg1 tx txagg1;
    USEOBSERVATIONS = gender EQ 1 AND (desgn11s EQ 1 OR
    desgn11s EQ 2 OR desgn11s EQ 3 OR desgn11s EQ 4);
DEFINE:
ANALYSIS: ESTIMATOR = MLR;
MODEL:
    IF (desgn11s EQ 4) THEN tx=1;
    IF (desgn11s EQ 1 OR desgn11s EQ 2 OR desgn11s EQ 3) THEN
    tx=0;
    agg5 = sctaa15s;
    agg1 = sctaa11f;
    CENTER agg1(GRANDMEAN);
    txagg1 = tx*agg1;
agg5 ON
tx (b1)
agg1 (b2)
txagg1 (b3);
MODEL CONSTRAINT:
    NEW(modlo mod0 modhi);
    modlo = b1+b3*(-1.06);
    mod0 = b1;
    modhi = b1+b3*1.06;
OUTPUT: SAMPSTAT PATTERNS STANDARDIZED RESIDUAL TECH4;
PLOT: TYPE = PLOT3;
```


## Example: Linear Regression with an Interaction

Table : Results for regression with a randomized intervention using treatment-baseline interaction ( $n=250$ )

|  | Estimate | S.E. | Est./S.E. | Two-Tailed P -Value |
| :---: | :---: | :---: | :---: | :---: |
| agg5 ON |  |  |  |  |
| tx | -0.285 | 0.124 | -2.307 | 0.021 |
| agg1 | 0.500 | 0.076 | 6.543 | 0.000 |
| txagg 1 | -0.066 | 0.130 | -0.511 | 0.609 |
| Intercepts |  |  |  |  |
| agg5 | 2.483 | 0.077 | 32.238 | 0.000 |
| Residual variances |  |  |  |  |
| agg5 | 0.952 | 0.090 | 10.612 | 0.000 |
| New/additional parameters |  |  |  |  |
| modlo | -0.215 | 0.177 | -1.211 | 0.226 |
| mod0 | -0.285 | 0.124 | -2.307 | 0.021 |
| modhi | -0.355 | 0.192 | -1.849 | 0.064 |

MODEL: agg5 ON
tx (b1)
agg1 (b2)
txagg 1 (b3);
MODEL CONSTRAINT:
LOOP(x,-1,1,0.1);
PLOT(effect);
effect $=\mathrm{b} 1+\mathrm{b} 3^{*} \mathrm{x}$;


## Heteroscedasticity Modeling: Example: LSAY Math Data $(n=2,019)$

Figure : Linear regression residuals for math10 plotted against math7


## Heteroscedasticity Modeling: (1) Using MODEL CONSTRAINT

The linear regression model assumes homoscedastic residual variances,

$$
\begin{align*}
y_{i} & =\beta_{0}+\beta_{1} x_{i}+\varepsilon_{i},  \tag{4}\\
V\left(\varepsilon_{i} \mid x_{i}\right) & =V\left(\varepsilon_{i}\right)=V(\varepsilon) . \tag{5}
\end{align*}
$$

An exponential function may instead be used for the residual variance,

$$
\begin{equation*}
V\left(\varepsilon_{i} \mid x_{i}\right)=e^{a+b x_{i}} \tag{6}
\end{equation*}
$$

where $a$ and $b$ are parameters to be estimated. If $b=0, V\left(\varepsilon_{i} \mid x_{i}\right)=e^{a}$ which means that the residual variance is not a function of $x$ so that homoscedasticity holds. If $b>0$, the residual variance increases as a function of $x$ and if $b<0$, the residual variance decreases as a function of $x$.

## Input for Heteroscedasticity Modeling

```
TITLE:
DATA:
VARIABLE:
DEFINE:
ANALYSIS:
MODEL: math10 ON math7 mothed male;
math10 (resvar);
MODEL CONSTRAINT:
    NEW(a b);
    resvar = EXP(a+b*math7);
OUTPUT: TECH8 SAMPSTAT
    CINTERVAL(BOOTSTRAP);
PLOT: TYPE = PLOT3;
```


## LL and BIC for Heteroscedasticity Modeling

Table : Loglikelihood and BIC for heteroscedasticity modeling of LSAY math data

|  | \#par's | $\operatorname{logL}$ | BIC |
| :--- | :---: | :---: | :---: |
| Regular <br> regression | 5 | -6972 | 13982 |
| Heteroscedasticity <br> regression | 6 | -6885 | 13816 |

# Non-Symmetric Bootstrap Confidence Intervals for Heteroscedasticity Modeling of the LSAY Math Data 

|  | Lower 2.5\% | Lower 5\% | Estimate | Upper 5\% | Upper 2.5\% |
| :--- | :---: | :---: | :---: | :---: | :---: |
| math10 ON |  |  |  |  |  |
| math7 | 0.981 | 0.986 | 1.017 | 1.050 | 1.058 |
| mothed | 0.529 | 0.579 | 0.872 | 1.148 | 1.192 |
| male | 0.115 | 0.215 | 0.822 | 1.426 | 1.514 |
| Intercepts |  |  |  |  |  |
| math10 | 6.664 | 7.061 | 8.759 | 10.444 | 10.804 |
| Residual |  |  |  |  |  |
| Variances |  |  |  |  |  |
| math10 | 999.000 | 999.000 | 999.000 | 999.000 | 999.000 |
| New/Additional |  |  |  |  |  |
| Parameters | 6.020 | 6.086 | 6.379 | 6.704 | 6.761 |
| a | -0.051 | -0.050 | -0.043 | -0.037 | -0.036 |
| b |  |  |  |  |  |

- Assuming homoscedasticity: Non-significant effect of male, $95 \% \mathrm{CI}$ is $[-0.167,1.336]$
- Allowing heteroscedasticity: Significant effect of male, $95 \%$ CI is $[0.115,1.514]$


## Heteroscedasticity Modeling: (2) Using Random Coefficients

$$
\begin{align*}
y_{i} & =\beta_{0}+\beta_{1 i} x_{i}+\beta_{2} z_{i}+\varepsilon_{i},  \tag{7}\\
\beta_{1 i} & =\beta_{1}+\beta_{3} z_{i}+\delta_{i} . \tag{8}
\end{align*}
$$

The residuals $\varepsilon$ and $\delta$ are allowed to covary. The model can be compared to regular regression with an interaction between the covariates $x$ and $z$ by inserting (8) into (7),

$$
\begin{equation*}
y_{i}=\beta_{0}+\beta_{1} x_{i}+\beta_{3} x_{i} z_{i}+\delta_{i} x_{i}+\beta_{2} z_{i}+\varepsilon_{i} . \tag{9}
\end{equation*}
$$

The random coefficient model allows for a heteroscedastic residual variance. Whereas in regular regression the residual variance is assumed to be the same for all individuals, $V(y \mid x, z)=V(\varepsilon)$, the residual variance for the random coefficient model varies with $x$. The conditional variance of $y$ in (9) is

$$
\begin{equation*}
V\left(y_{i} \mid x_{i}, z_{i}\right)=V\left(\delta_{i}\right) x_{i}^{2}+2 \operatorname{Cov}\left(\delta_{i}, \varepsilon_{i}\right) x_{i}+V\left(\varepsilon_{i}\right) \tag{10}
\end{equation*}
$$

```
ANALYSIS: TYPE = RANDOM;
MODEL: s |math10 ON math7;
    s WITH math10 (cov);
    math10 (resvary);
    s (vbeta);
OUTPUT: TECH1 SAMPSTAT STDYX RESIDUAL CINTERVAL;
PLOT: TYPE = PLOT3;
MODEL CONSTRAINT:
    PLOT (vygivenx);
    LOOP(x,25,90,1);
    vygivenx = vbeta*x*x + 2* cov*x + resvary;
```

- Better BIC than homoscedastic model



## Censored Variable Modeling

## $30 \%$ floor effect:


$59 \%$ floor effect:


- Censored-normal (Tobit)
- Censored-inflated
- Sample selection (Heckman)
- Two-part


## Censored-Normal (Tobit) Regression




$$
\begin{gather*}
y_{i}^{*}=\beta_{0}+\beta_{1} x_{i}+\varepsilon_{i}  \tag{11}\\
y_{i}= \begin{cases}0 & \text { if } y_{i}^{*} \leq 0 \\
y^{*} & \text { if } y_{i}^{*}>0\end{cases}
\end{gather*}
$$

Binary (probit) $: P\left(y_{i}>0 \mid x_{i}\right)=1-\Phi\left[\frac{0-\beta_{0}-\beta_{1} x_{i}}{\sqrt{V(\varepsilon)}}\right]=\Phi\left[\frac{\beta_{0}+\beta_{1} x_{i}}{\sqrt{V(\varepsilon)}}\right]$,

Continuous, positive : $E\left(y_{i} \mid y_{i}>0, x_{i}\right)=\beta_{0}+\beta_{1} x_{i}+\sqrt{V(\varepsilon)} \frac{\phi\left(z_{i}\right)}{\Phi\left(z_{i}\right)}$,

## Censored-Inflated Regression

- Latent class 0: subjects for whom only $y=0$ is observed
- Latent class 1: subjects following a censored-normal (tobit) model

Assume a logistic regression that describes the probability of being in class 0 ,

$$
\begin{equation*}
\operatorname{logit}\left(\pi_{i}\right)=\gamma_{0}+\gamma_{1} x_{i} \tag{14}
\end{equation*}
$$

For subjects in class 1 the usual censored-normal model of (15) is assumed with

$$
\begin{equation*}
y_{i}^{*}=\beta_{0}+\beta_{1} x_{i}+\varepsilon_{i} . \tag{15}
\end{equation*}
$$

Two ways $y=0$ is observed (mixture at zero).

## Sample Selection (Heckman) Regression

Consider the linear regression for the continuous latent response variable $y^{*}$,

$$
\begin{equation*}
y_{i}^{*}=\beta_{0}+\beta_{1} x_{i}+\varepsilon_{i} \tag{16}
\end{equation*}
$$

where the latent response variable $y_{i}^{*}$ is observed as $y_{i}=y_{i}^{*}$ when a binary variable $u_{i}=1$ and remains latent, that is, missing if $u_{i}=0$. A probit regression is specified for $u$,

$$
\begin{equation*}
u_{i}^{*}=\gamma_{1} x_{i}+\delta_{i} \tag{17}
\end{equation*}
$$

where the categories of the binary observed variable $u_{i}$ are determined by $u^{*}$ falling below or exceeding a threshold parameter $\tau$,

$$
u_{i}= \begin{cases}0 & \text { if } u_{i}^{*} \leq \tau \\ 1 & \text { if } u_{i}^{*}>\tau\end{cases}
$$

A key feature is that the residuals $\varepsilon$ and $\delta$ are assumed to be correlated and have a bivariate normal distribution with the usual probit standardization $V(\boldsymbol{\delta})=1$.

With censoring from below at zero and using probit regression with the event of $u=1$ referring to a positive outcome, the two-part model is expressed as

$$
\begin{align*}
\operatorname{probit}\left(\pi_{i}\right) & =\gamma_{0}+\gamma_{1} x_{i}  \tag{18}\\
\log y_{i \mid u_{i}=1} & =\beta_{0}+\beta_{1} x_{i}+\varepsilon_{i} \tag{19}
\end{align*}
$$

where $\pi_{i}=P\left(u_{i}=1 \mid x_{i}\right)$ and $\varepsilon_{i} \sim N(0, V(\varepsilon))$. Logistic regression can be used as an alternative to the probit regression in (18).
Maximum-likelihood estimation of the two-part model gives the same estimates as if the binary and the continuous parts were estimated separately using maximum-likelihood. Expressing (18) in terms of a latent response variable regression with a normal residual, the two residuals can be correlated but the correlation does not enter into the likelihood and is not estimated.

## Comparison of Censored-Inflated, Heckman, and Two-Part

- Like the censored-inflated and Heckman models, the two-part model has different regression equations for the two parts
- Unlike the censored-inflated model, the two-part model does not have a mixture at zero, nor does Heckman
- Unlike the Heckman model, the two-part model does not estimate a residual correlation between the two parts
- Duan et al. (1983) pointed to two advantages of the two-part model over Heckman:
- Applied to medical care expenses, it is preferable to the Heckman model because the censoring point of zero expense does not represent missing data but rather a real, observed value
- A bivariate normality assumption for the residuals is not needed


## Example: Comparing Methods on Heavy Drinking Data



NLSY Data on
Heavy Drinking
( $n=1,152$ )

- Dependent variable: frequency of heavy drinking measured by the question:
- "How often have you had 6 or more drinks on one occasion during the last 30 days?"
- Never (0); once (1); 2 or 3 times (2); 4 or 5 times (3); 6 or 7 times (4); 8 or 9 times (5); and 10 or more times (6)
- Covariates: gender, ethnicity, early onset of regular drinking (es), family history of problem drinking, and high school dropout.

```
    USEVARIABLES \(=\) hd84 male black hisp es fh 123 hsdrp;
    CENSORED = hd84 (B);
ANALYSIS: ESTIMATOR \(=\) MLR;
MODEL: hd84 ON male black hisp es fh123 hsdrp;
```

USEVARIABLES $=$ hd84 male black hisp es fh123 hsdrp;
CENSORED = hd84 (BI);
ANALYSIS: ESTIMATOR = MLR;
MODEL: hd84 ON male black hisp es fh123 hsdrp;
hd84\#1 ON male black hisp es fh123 hsdrp;

## DATA TWOPART

The DATA TWOPART command is used to create a binary and a continuous variable from a continuous variable with a floor effect. A cutpoint of zero is used as the default. Following are the rules used to create the two variables:
(1) If the value of the original variable is missing, both the new binary and the new continuous variable values are missing
(2) If the value of the original variable is greater than the cutpoint value, the new binary variable value is one and the new continuous variable value is the $\log$ of the original variable as the default

- If the value of the original variable is less than or equal to the cutpoint value, the new binary variable value is zero and the new continuous variable value is missing


## Input for Heckman and Two-Part



## Loglikelihood and BIC for Four Models for Frequency of Heavy Drinking

The Heckman and two-part models use $\log (y)$ so $\log \mathrm{L}$ and BIC values cannot be compared to those of tobit and censored-inflated:

| Model | $\log \mathrm{L}$ | \# parameters | BIC |
| :--- | :---: | :---: | :---: |
| Censored-normal (tobit) | -1530.512 | 8 | 3117 |
| Censored-inflated | -1499.409 | 15 | 3105 |
| Sample selection (Heckman) | -1088.182 | 16 | 2289 |
| Two-part | -1088.400 | 15 | 2283 |


| Parameter | Estimate | S.E. | Est./S.E. | Two-Tailed <br> P-Value |
| :--- | :---: | :---: | :---: | :---: |
| hd84 ON |  |  |  |  |
| male | 2.106 | 0.210 | 10.038 | 0.000 |
| black | -2.157 | 0.258 | -8.359 | 0.000 |
| hisp | -1.059 | 0.298 | -3.555 | 0.000 |
| es | 0.716 | 0.286 | 2.503 | 0.012 |
| fh123 | 0.615 | 0.317 | 1.938 | 0.053 |
| hsdrp | 0.240 | 0.265 | 0.908 | 0.364 |
| Intercepts |  |  |  |  |
| hd84 | -1.258 | 0.211 | -5.961 | 0.000 |
| Residual variances |  |  |  |  |
| hd84 | 8.678 | 0.559 | 15.525 | 0.000 |


|  |  |  |  | Two-Tailed <br> Parameter |
| :--- | ---: | ---: | ---: | :--- |
|  | Estimate | S.E. | Est./S.E. | P-Value |
| hd84 ON |  |  |  |  |
| male | 0.957 | 0.236 | 4.057 | 0.000 |
| black | -1.150 | 0.282 | -4.073 | 0.000 |
| hisp | -0.405 | 0.320 | -1.264 | 0.206 |
| es | 0.585 | 0.276 | 2.120 | 0.034 |
| fh123 | -0.031 | 0.329 | -0.095 | 0.924 |
| hsdrp | 0.390 | 0.263 | 1.487 | 0.137 |
| hd84\#1 ON |  |  |  |  |
| male | -1.025 | 0.166 | -6.157 | 0.000 |
| black | 0.962 | 0.208 | 4.621 | 0.000 |
| hisp | 0.570 | 0.215 | 2.651 | 0.008 |
| es | -0.204 | 0.198 | -1.032 | 0.302 |
| fh123 | -0.512 | 0.273 | -1.876 | 0.061 |
| hsdrp | 0.040 | 0.188 | 0.213 | 0.831 |
| Intercepts |  |  |  |  |
| hd84\#1 | 0.412 | 0.145 | 2.848 | 0.004 |
| hd84 | 1.567 | 0.189 | 8.290 | 0.000 |

## Comparisons of Results

- Heckman versus Two-part:
- Very similar $\log$ L/BIC and results (the Heckman probit coefficients need to be divided by $\sqrt{2}$ due to adding the factor)
- The Heckman residual correlation is significant
- Censored-inflated versus Two-part:
- Similar results (reverse signs for the binary part)
- LogL and BIC not comparable but limited model fit comparison can be made using MODEL CONSTRAINT:

Table : Estimated probability of zero heavy drinking and mean of heavy drinking for a subset of males who have zero values on the covariates black, hisp, es, fh123, and hsdrp

|  | Probability | Mean |
| :--- | :---: | :---: |
| Sample values | 0.441 | 1.538 |
| Censored-inflated estimates | 0.402 | 1.547 |
| Two-part estimates | 0.403 | 1.671 |



- Assignment: As an alternative, an ordinal approach may be good for these data given
(1) the limited number of response categories
(2) the slight ceiling effect for category 6,10 or more times so that the assumption of a log normal distribution can be questioned:
- Declare the positive part as categorical using the CATEGORICAL option of the VARIABLE command
- Use TRANSFORM = NONE in the DATA TWOPART command to avoid the log transformation
- Six key advantages of Bayesian analysis over frequentist analysis using maximum likelihood estimation:
(1) More can be learned about parameter estimates and model fit
(2) Small-sample performance is better and large-sample theory is not needed
(3) Parameter priors can better reflect results of previous studies
(4) Analyses are in some cases less computationally demanding, for example, when maximum-likelihood requires high-dimensional numerical integration
(5) In cases where maximum-likelihood computations are prohibitive, Bayes with non-informative priors can be viewed as a computing algorithm that would give essentially the same results as maximum-likelihood if maximum-likelihood estimation were computationally feasible
(6) New types of models can be analyzed where the maximum-likelihood approach is not practical

Figure : Prior, likelihood, and posterior for a parameter


- Priors:
- Non-informative priors (diffuse priors): Large variance (default in Mplus)
- A large variance reflects large uncertainty in the parameter value. As the prior variance increases, the Bayesian estimate gets closer to the maximum-likelihood estimate
- Weakly informative priors: Used for technical assistance
- Informative priors:
- Informative priors reflect prior beliefs in likely parameter values
- These beliefs may come from substantive theory combined with previous studies of similar populations


## Convergence: Trace Plot for Two MCMC Chains. PSR



Potential scale reduction criterion (Gelman \& Rubin, 1992):

$$
\begin{equation*}
P S R=\sqrt{\frac{W+B}{W}} \tag{20}
\end{equation*}
$$

where $W$ represents the within-chain variation of a parameter and $B$ represents the between-chain variation of a parameter. A PSR value close to 1 means that the between-chain variation is small relative to the within-chain variation and is considered evidence of convergence.

## Convergence of the Bayes

 Markov Chain Monte Carlo (MCMC) AlgorithmFigure : Premature stoppage of Bayes MCMC iterations using the Potential Scale Reduction (PSR) criterion


## Trace and Autocorrelation Plots Indicating Poor Mixing




# Bayes Posterior Distribution Similar to ML Bootstrap Distribution: Credibility versus Confidence Intervals 




- Frequentists often object to Bayes using informative priors
- But they already do use such priors in many cases in unrealistic ways (e.g. factor loadings fixed exactly at zero)
- Bayes can let informative priors reflect prior studies
- Bayes can let informative priors identify models that are unidentified by ML which is useful for model modification (BSEM)
- The credibility interval for the posterior distribution is narrower with informative priors


## Speed Of Bayes In Mplus

Wang \& Preacher (2014). Moderated mediation analysis using Bayesian methods. Structural Equation Modeling.

- Comparison of ML (with bootstrap) and Bayes: Similar statistical performance
- Comparison of Bayes using BUGS versus Mplus: Mplus is 15 times faster
- Reason for Bayes being faster in Mplus:
- Mplus uses Fortran (fastest computational environment)
- Mplus uses parallel computing so each chain is computed separately
- Mplus uses the largest updating blocks possible - complicated to program but gives the best mixing quality
- Mplus uses sufficient statistics
- Mplus Bayes considerably easier to use

Regressing $y$ On $x$ : Bringing $x$ 's Into The Model

ML estimation maximizes the log likelihood for the bivariate distribution of $y$ and $x$ expressed as,

$$
\log L=\sum_{i} \log \left[y_{i}, x_{i}\right]=\sum_{i=1}^{n_{1}} \log \left[y_{i} \mid x_{i}\right]+\sum_{i=1}^{n_{1}+n_{2}} \log \left[x_{i}\right]+\sum_{i=n_{1}+n_{2}+1}^{n_{1}+n_{2}+n_{3}} \log \left[y_{i}\right] .
$$

Figure : Missing data patterns. White areas represent missing data


## Example: Monte Carlo Simulation Study

- Linear regression with $40 \%$ missing on $x_{1}-x_{4}$; no missing on $y$
- $x_{3}$ and $x_{4}$ s are binary split $86 / 16$
- MAR holds as a function of the covariate $z$ with no missing
- $n=200$
- Comparison of Bayes and ML



## Bayes Treating Binary X's As Binary

| DATA: | FILE = MARn200replist.dat; |
| :--- | :--- |
|  | TYPE = MONTECARLO; |
| VARIABLE: | NAMES = y x1-x4 z; |
|  | USEVARIABLES = y x1-z; |
|  | CATEGORICAL = x3-x4; |
| DEFINE: | IF(z gt .25)THEN x1=_MISSING; |
|  | IF(z gt .25)THEN x2=_MISSING; |
|  | IF(-z gt .25)THEN x3=_MISSING; |
|  | IF(-z gt .25)THEN x4=_MISSING; |
| ANALYSIS: | ESTIMATOR = BAYES; |
|  | PROCESSORS $=2 ;$ |
|  | BITERATIONS $=(10000) ;$ |
|  | MEDIATOR $=\mathbf{O B S E R V E D ; ~}$ |
| MODEL: | y ON x1-z*.5; |
|  | y*1; |
|  | x1-z WITH x1-z; |

## ML Versus Bayes Treating Binary X's As Binary

- Attempting to estimate the same model using ML leads to much heavier computations due to the need for numerical integration over several dimensions
- Already in this simple model ML requires three dimensions of integration, two for the $x_{3}, x_{4}$ covariates and one for a factor capturing the association between $x_{3}$ and $x_{4}$.
- Bayes uses a multivariate probit model that generates correlated latent response variables underlying the binary $x$ 's - no need for numerical integration


## Bayes’ Advantage Over ML: Missing Data with a Binary Outcome

Figure : Mediation model for a binary outcome of dropping out of high school ( $\mathrm{n}=2898$ )


## Bayes With Missing Data On The Mediator

|  | CATEGORICAL = hsdrop; |
| :--- | :--- |
| ANALYSIS: | ESTIMATOR = BAYES; <br>  <br>  <br> PROCESSORS $=2 ;$ |
| MODEL: | BITERATIONS = (20000); |
| hsdrop ON math10 female-math7; |  |
| MODEL INDIRECT: | math10 ON female-math7; |
| OUTPUT: | hsdrop IND math10 math7(61.01 50.88); |
| PLOT: | SAMPSTAT PATTERNS TECH1 TECH8 CINTERVAL; |
|  | TYPE = PLOT3; |

Indirect and direct effects computed in probability scale using counterfactually-based causal effects.

## Bayesian Posterior Distribution Of Indirect Effect For High School Dropout



## Missing On The Mediator: ML Versus Bayes

ML estimates are almost identical to Bayes, but:

- ML needs Monte Carlo integration with 250 points because the mediator is a partially latent variable due to missing data
- ML needs bootstrapping (1,000 draws) to capture CIs for the non-normal indirect effect
- ML takes 21 minutes
- Bayes takes 21 seconds
- Bayes posterior distribution for the indirect effect is based on 20,000 draws as compared to 1,000 bootstraps for ML


# Missing On The Mediator And The Covariates Treating All Covariates As Normal: ML Versus Bayes 

- ML requires integration over 10 dimensions
- ML needs 2,500 Monte Carlo integration points for sufficient precision
- ML takes 6 hours with 1,000 bootstraps
- Bayes takes less than a minute
- Bayes posterior based on 20,000 draws as compared to 1,000 bootstraps for ML


# Missing On The Mediator And The Covariates Treating Binary Covariates As Binary: ML Versus Bayes 

6 covariates are binary.

- ML requires $10+15=35$ dimensions of integration: intractable
- Bayes takes 3 minutes for 20,000 draws


## Mediation Analysis

Figure : A basic mediation model with an exposure variable $x$, a control variable $c$, a mediator $m$, and an outcome $y$


## Moderated Mediation Analysis: Case 1 (xz)

Figure : Case 1 moderated mediation of $y$ on $x, m$ on $x$, both moderated by $z$


$$
\begin{align*}
\text { Indirect } & : \beta_{1}\left(\gamma_{1}+\gamma_{3} z\right)\left(x_{1}-x_{0}\right),  \tag{21}\\
\text { Direct } & :\left(\beta_{2}+\beta_{4} z\right)\left(x_{1}-x_{0}\right) . \tag{22}
\end{align*}
$$

## Moderated Mediation Analysis: Case $2(m z)$

Figure : Case 2 moderated mediation of $y$ on $m$ moderated by $z$


$$
\begin{align*}
& \text { Indirect: }\left(\beta_{1}+\beta_{4} z\right) \gamma_{1}\left(x_{1}-x_{0}\right),  \tag{23}\\
& \text { Direct }: \beta_{2}\left(x_{1}-x_{0}\right) . \tag{24}
\end{align*}
$$

## Moderated Mediation Analysis: Case 3 ( $m x$ )

Figure : Case 3 moderated mediation of $y$ on $m$ moderated by $x$


$$
\begin{align*}
& \text { Indirect }:\left(\beta_{1}+\beta_{3} x_{1}\right) \gamma_{1}\left(x_{1}-x_{0}\right),  \tag{25}\\
& \text { Direct }:\left(\beta_{2}+\beta_{3}\left(\gamma_{0}+\gamma_{1} x_{0}+\gamma_{2} c\right)\right)\left(x_{1}-x_{0}\right) . \tag{26}
\end{align*}
$$

## Example: Case 2 Moderated Mediation for Work Team Performance (Hayes, 2013; $n=60$ )

Figure : Case $2(m z)$ moderated mediation for work team behavior. The exposure variable is dysfunc (continuous). The interaction variable $m z$ is the product of the mediator variable negtone and the moderator variable negexp


```
TITLE:
DATA:
VARIABLE:
DEFINE:
ANALYSIS:
MODEL:
MODEL INDIRECT:
perform MOD negtone negexp(-.4,.6,.1)
mz dysfunc(0.4038 0.035);
OUTPUT: SAMPSTAT CINTERVAL(BOOTSTRAP);
PLOT: TYPE = PLOT3;
```

- The moderator variable negexp has $20^{t h}$ and $80^{\text {th }}$ percentiles -0.4 and 0.6 , respectively
- The exposure variable dysfunc has mean 0.4038 and standard deviation 0.369 so that $x_{1}-x_{0}=0.4038-0.035=0.369$. In other words, 0.035 is one standard deviation below the mean


## Indirect Effect Plot for Work Team Behavior Example

Figure : Indirect effect and bootstrap confidence interval for case 2 ( mz ) moderated mediation for work team behavior. The moderator variable is negexp and the indirect effect is labeled Total natural IE


## Ignore Chi-Square Test of Model Fit When Interaction Involves the Mediator

An alternative specification used in Preacher et al. (2007) avoids the two degrees of freedom that arise because of the two left-out arrows in the model. This saturates the model by allowing covariances between the moderator variable and the mediator residual and between the moderator-exposure interaction variable and the mediator residual. To accomplish this, the MODEL specification adds a line using WITH:

## MODEL:

perform ON negtone dysfunc negexp mz; negtone ON dysfunc; negexp mz WITH negtone dysfunc;

## Example: Case 3 Moderated Mediation



The effects of $x$ on $y$ are

$$
\begin{align*}
\text { Indirect } & :\left(\beta_{1}+\beta_{3} x_{1}\right) \gamma_{1}\left(x_{1}-x_{0}\right),  \tag{27}\\
\text { Direct } & :\left(\beta_{2}+\beta_{3}\left(\gamma_{0}+\gamma_{1} x_{0}\right)\right)\left(x_{1}-x_{0}\right) . \tag{28}
\end{align*}
$$

Quoting VanderWeele (2015, p. 46):
"An investigator might be tempted to only include such exposure-mediator interactions in the model if the interaction is statistically significant. - This approach is problematic. It is problematic because power to detect interaction tends to be very low unless the sample size is very large. - such exposure-mediator interaction may be important in capturing the dynamics of mediation... - A better approach - - is perhaps to include them by default and only exclude them if they do not seem to change the estimates of the direct and indirect effects very much."

```
TITLE:
DATA:
VARIABLE:
DEFINE:
ANALYSIS:
MODEL:
    x moderation of y regressed on m
FILE = xmVx4s1n200rep6.dat;
NAMES = y m x;
USEVARIABLES = y m x mx;
mx = m*x;
ESTIMATOR = ML;
BOOTSTRAP = 10000;
y ON m x mx;
m ON x;
MODEL INDIRECT:
y MOD m mx x(7 5);
OUTPUT:
SAMPSTAT CINTERVAL(BOOTSTRAP);
PLOT:
TYPE = PLOT3;
```


## Monte Carlo Study of Moderated Mediation



The model used for data generation is

$$
\begin{align*}
y_{i} & =\beta_{0}+\beta_{1} m_{i}+\beta_{2} x_{i}+\beta_{3} z_{i}+\varepsilon_{y i},  \tag{29}\\
m_{i} & =\gamma_{0}+\gamma_{1 i} x_{i}+\gamma_{2} z_{i}+\varepsilon_{m i},  \tag{30}\\
\gamma_{1 i} & =\gamma_{1}+\gamma_{3} z_{i}, \tag{31}
\end{align*}
$$

where $\gamma_{1 i}$ is a random slope. Inserting (31) in (30) shows that the random slope formulation is equivalent to adding an interaction term $x z$ as a covariate in the regression of $m$.

```
TITLE: Simulating Z moderation of X to M using a random slope, saving the
data for external Monte Carlo analysis
MONTECARLO:
NAMES = y m x z;
NOBS = 400;
NREPS = 500;
REPSAVE = ALL;
SAVE = xzrep*.dat;
CUTPOINTS = x(0);
MODEL POPULATION:
    x-z@1; [x-z@0];
    x WITHz@0.5;
    y ON m*.5 x*.2 z*.1; y*.5; [y*0];
    gamma1 |mON x;
    [gamma1*.3];
    gamma1 ON z*.2;
    gamma1@0;
    m ON z*.3; m*1; [m*0];
ANALYSIS:
TYPE = RANDOM;
MODEL:
y ON m*.5 (b)
x*.2 z*.1;
y*.5; [y*0];
gamma1 | m ON x;
[gamma1*.3] (gamma1);
gamma1 ON z*.2 (gamma3);
gamma1@0;
m ON z*.3; m*1; [m*0];
MODEL CONSTRAINT:
NEW(indavg*. }15\mathrm{ indlow*. }05\mathrm{ indhigh*.25);
indavg = b*gamma1;
indlow = b* (gamma1-gamma3);
indhigh = b*(gamma1+gamma3);
```


## Results for Monte Carlo Simulation of $z$ Moderation of $m$ Regressed on $x$ using $n=400$ and 500 Replications

|  | Population | Average | Std. Dev. | S.E. <br> Average | M.S.E. | 95\% Cover | \% Sig Coeff |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| gammal ON |  |  |  |  |  |  |  |
| z | 0.200 | 0.2010 | 0.0775 | 0.0771 | 0.0060 | 0.950 | 0.744 |
| y ON |  |  |  |  |  |  |  |
| m | 0.500 | 0.5007 | 0.0524 | 0.0494 | 0.0027 | 0.922 | 1.000 |
| x | 0.200 | 0.2056 | 0.0783 | 0.0784 | 0.0061 | 0.938 | 0.754 |
| z | 0.100 | 0.0963 | 0.0470 | 0.0433 | 0.0022 | 0.926 | 0.604 |
| m ON |  |  |  |  |  |  |  |
| z | 0.300 | 0.2999 | 0.0531 | 0.0545 | 0.0028 | 0.964 | 1.000 |
| Intercepts |  |  |  |  |  |  |  |
| y | 0.000 | -0.0017 | 0.0527 | 0.0522 | 0.0028 | 0.934 | 0.066 |
| m | 0.000 | -0.0008 | 0.0543 | 0.0545 | 0.0029 | 0.946 | 0.054 |
| gamma1 | 0.300 | 0.3010 | 0.0776 | 0.0770 | 0.0060 | 0.962 | 0.978 |
| Residual |  |  |  |  |  |  |  |
| y | 0.500 | 0.4938 | 0.0341 | 0.0347 | 0.0012 | 0.928 | 1.000 |
| m | 0.500 | 0.4940 | 0.0331 | 0.0346 | 0.0011 | 0.950 | 1.000 |
| gamma1 | 0.000 | 0.0000 | 0.0000 | 0.0000 | 0.0000 | 1.000 | 0.000 |
| New/Additional |  |  |  |  |  |  |  |
| Parameters indavg | 0.150 | 0.1505 | 0.0417 | 0.0416 | 0.0017 | 0.956 | 0.974 |
| indlow | 0.050 | 0.0497 | 0.0546 | 0.0548 | 0.0030 | 0.958 | 0.138 |
| indhigh | 0.250 | 0.2514 | 0.0628 | 0.0603 | 0.0039 | 0.928 | 0.988 |

## Sensitivity Analysis

Figure : Mediator-outcome confounding 1


Figure : Mediator-outcome confounding 2



A moderated mediation model of sex discrimination in the work place. The interaction variable $x z$ is the product of the exposure variable protest and the moderator variable sexism ( $n=129$ )

- Variables:
- Protest: binary exposure variable (2 randomized scenarios of female attorney taking action or not)
- Sexism: Moderator variable
- Respappr: Mediator - perceived appropriateness of response)
- Liking: Outcome - how well the subject likes the female attorney


# Results for Combined Moderated Mediation for Sex Discrimination 

|  | Estimate | S.E. | Est./S.E. | Two-Tailed P -Value |
| :---: | :---: | :---: | :---: | :---: |
| liking ON |  |  |  |  |
| respappr | 0.098 | 0.533 | 0.184 | 0.854 |
| protest | -3.119 | 1.750 | -1.782 | 0.075 |
| sexism | -0.462 | 0.502 | -0.919 | 0.358 |
| mx | 0.112 | 0.157 | 0.715 | 0.475 |
| mz | 0.039 | 0.100 | 0.392 | 0.695 |
| xz | 0.500 | 0.341 | 1.466 | 0.143 |
| respappr ON |  |  |  |  |
| protest | -2.687 | 1.738 | -1.546 | 0.122 |
| sexism | -0.529 | 0.320 | -1.654 | 0.098 |
| xz | 0.810 | 0.346 | 2.343 | 0.019 |
| Intercepts |  |  |  |  |
| liking | 6.510 | 2.623 | 2.482 | 0.013 |
| respappr | 6.567 | 1.596 | 4.114 | 0.000 |
| Residual Variances |  |  |  |  |
| liking | 0.779 | 0.135 | 5.767 | 0.000 |
| respappr | 1.269 | 0.156 | 8.121 | 0.000 |

Figure : Loop plot of indirect effect and confidence interval for combined moderated mediation case of sex discrimination. The moderator is labeled $z$ in MODEL CONSTRAINT and corresponds to the sexism variable


Table : Input for moderated mediation for sex discrimination data

```
TITLE: Hayes PROTEST moderation of X ->M, X->Y
DATA: FILE = protest.txt;
VARIABLE: NAMES = sexism liking respappr protest;
    USEVARIABLES = liking respappr protest sexism xz;
    xz = protest*sexism;
    ESTIMATOR = ML;
    BOOTSTRAP = 1000;
MODEL: liking ON respappr (beta1)
    protest (beta2)
    sexism
    xz (beta4);
    respappr ON protest (gamma1)
    sexism (gamma2)
    xz (gamma3);
MODEL INDIRECT:
    liking MOD respappr sexism(4,6,1) xz protest;
OUTPUT: SAMPSTAT STANDARDIZED
    CINTERVAL(BOOTSTRAP);
PLOT: TYPE = PLOT3 SENSITIVITY;
```

Figure : Sensitivity plot for the indirect effect and its confidence interval at the sexism mean of 5 in a study of sex discrimination in the workplace. The x -axis represents the residual correlation $\rho$ and the y -axis represents the indirect effect


## Counterfactually-Defined Causal Effects:

 Potential Outcomes, Counterfactuals, and Causal Effects|  |  | Potential Outcomes |  |  |
| :---: | :---: | :---: | :---: | :---: |
| i | $X_{i}$ | $Y_{i}\left(X_{i}=1\right)$ | $Y_{i}\left(X_{i}=0\right)$ | Causal effect <br> $Y_{i}\left(X_{i}=1\right)-Y_{i}\left(X_{i}=0\right)$ |
| 1 | 1 | 11 | 9 | 2 |
| 2 | 1 | 14 | 10 | 4 |
| 3 | 0 | 8 | $\boxed{5}$ | 3 |
| 4 | 1 | 9 | 8 | 1 |
| 5 | 0 | 18 | $\boxed{9}$ | 6 |
| 6 | 0 | 11 | 12 | 1 |
| True average <br> Observed average | 11.33 | 9 | 2.83 |  |

## Counterfactually-Defined Causal Effects: Robins, Pearl, VanderWeele, Imai

- Counterfactuals and potential outcomes:
- Chapter 4: continuous mediator and continuous outcome
- Chapter 8: continuous mediator and binary outcome, binary mediator and continuous or binary outcome, count outcome, two-part outcome
- Counterfactually-defined causal indirect and direct effects:
- Strict assumptions including no mediator-outcome confounding
- $X=$ exposure variable, $M=$ mediator, $Y=$ outcome
- Total effect: $E[Y(1, M(1))]-E[Y(0, M(0))]$, treatment group mean of $Y$ minus control group mean of $Y$
- The Total Natural Indirect Effect (TNIE) $=E[Y(1, M(1))]-E[Y(1, M(0))]$ where 1 and 0 represent treatment and control for the exposure variable
- What does it mean?
- Explanations in words and formulas


## Indirect Effect $T N I E=E[Y(1, M(1))]-E[Y(1, M(0))]$

- In words:
- $E[Y(1, M(1))]$ is the mean of the outcome when subjects get the treatment $(X=1)$ and $M$ varies as it would under the treatment condition $(X=1)$ - this is the treatment group mean
- $E[Y(1, M(0))]$ is the mean of the outcome when subjects get the treatment ( $X=1$ ) but $M$ varies as it would under the control condition $(X=0)$ - this is a counterfactual
- In formulas:
- To get an effect of $X$ on $Y$ we need to integrate out $M$
- $M$ has two different distributions $f(M \mid X): M(0)$ for $X=0$ and $M(1)$ for $X=1$. For example:
- $E[Y(1, M(0))]=\int_{-\infty}^{+\infty} E[Y \mid X=1, M=m] \times f(M \mid X=0) \partial M$
- In some cases, this integral is simple - integration does not need to be involved: (1) Continuous $M$, continuous $Y$, (2) Continuous $M$, binary $Y$ with probit
- In some cases, the integration is needed: (1) Continuous $M$, binary $Y$ with logistic (numerical integration needed), (2) Count $Y$, (3) $\log (Y)$
- Continuous $M$ and $Y$ :

$$
\begin{align*}
Y_{i} & =\beta_{0}+\beta_{1} M_{i}+\beta_{2} X_{i}+\varepsilon_{y i},  \tag{32}\\
M_{i} & =\gamma_{0}+\gamma_{1} X_{i}+\varepsilon_{m i} . \tag{33}
\end{align*}
$$

Inserting (33) in (32) and integrating over $M$,

$$
\begin{align*}
E\left[Y\left(x_{1}, M\left(x_{0}\right)\right)\right] & =\beta_{0}+\beta_{2} x_{1}+ \\
& +\beta_{1} \int_{-\infty}^{+\infty} M f\left(M ; \gamma_{0}+\gamma_{1} x_{0}, \sigma^{2}\right) \partial M, \\
& =\beta_{0}+\beta_{2} x_{1}+\beta_{1}\left(\gamma_{0}+\gamma_{1} x_{0}\right) . \tag{34}
\end{align*}
$$

Conditioning on $X=x_{1}$ in (32) and $X=x_{0}$ in (33) and inserting the mediator expression in the outcome expression, the expected value is the same:

$$
\begin{equation*}
=\beta_{0}+\beta_{2} x_{1}+\beta_{1}\left(\gamma_{0}+\gamma_{1} x_{0}\right) \tag{35}
\end{equation*}
$$

- TNIE for continuous $M$ and $Y$ :

$$
\begin{align*}
& E\left[Y\left(x_{1}, M\left(x_{1}\right)\right)\right]-E\left[Y\left(x_{1}, M\left(x_{0}\right)\right)\right]  \tag{36}\\
& =\beta_{0}+\beta_{2} x_{1}+\beta_{1}\left(\gamma_{0}+\gamma_{1} x_{1}\right)  \tag{37}\\
& -\left(\beta_{0}+\beta_{2} x_{1}+\beta_{1}\left(\gamma_{0}+\gamma_{1} x_{0}\right)\right)  \tag{38}\\
& =\beta_{1} \gamma_{1}\left(x_{1}-x_{0}\right) . \tag{39}
\end{align*}
$$

- Note 1: Often $x_{1}-x_{0}=1$ such as with a one-unit change or treatment/control.
- Note 2: $\beta_{0}, \gamma_{0}, \beta_{2}$ cancel out. The indirect effect is a product of 2 slopes. This is not the case for binary $Y$


## Now We Know How To Do TNIE for Binary $Y$

$$
\begin{align*}
& Y_{i}^{*}=\beta_{0}+\beta_{1} M_{i}+\beta_{2} X_{i}+\varepsilon_{y i}  \tag{40}\\
& M_{i}=\gamma_{0}+\gamma_{1} X_{i}+\varepsilon_{m i} . \tag{41}
\end{align*}
$$

Conditioning on $X=x_{1}$ and $X=x_{0}$, for $Y^{*}$ and $M$, respectively, and inserting $M$ into $Y$,

$$
\begin{align*}
E\left(Y^{*} \mid X\right) & =\beta_{0}+\beta_{1} \gamma_{0}+\beta_{1} \gamma_{1} x_{0}+\beta_{2} x_{1},  \tag{42}\\
V\left(Y^{*} \mid X\right) & =V\left(\beta_{1} \varepsilon_{m}+\varepsilon_{y}\right)=\beta_{1}^{2} \sigma_{m}^{2}+c .  \tag{43}\\
P(Y=1 \mid X) & =\Phi\left[E\left(Y^{*} \mid X\right) / \sqrt{V\left(Y^{*} \mid X\right)}\right],  \tag{44}\\
\text { TNIE } & =\Phi[1,1]-\Phi[1,0], \tag{45}
\end{align*}
$$

where $\Phi[1,1]$ uses $\beta_{0}+\beta_{1} \gamma_{0}+\beta_{1} \gamma_{1} x_{1}+\beta_{2} x_{1}$ in $E\left(Y^{*} \mid X\right)$ and $\Phi[1,0]$ uses $\beta_{0}+\beta_{1} \gamma_{0}+\beta_{1} \gamma_{1} x_{0}+\beta_{2} x_{1}$. All 6 parameters involved.

## Effects Expressed on an Odds Ratio Scale for a Binary Outcome: Probit Model

The total natural indirect effect odds ratio for a binary exposure can be expressed as

$$
\begin{align*}
\operatorname{TNIE}(O R) & =\frac{P\left(Y_{x_{1} M_{x_{1}}}=1\right) /\left(1-P\left(Y_{x_{1} M_{x_{1}}}=1\right)\right.}{P\left(Y_{x_{1} M_{x_{0}}}=1\right) /\left(1-P\left(Y_{x_{1} M_{x_{0}}}=1\right)\right)} \\
& =\frac{\Phi[\operatorname{probit}(1,1)] /(1-\Phi[\operatorname{probit}(1,1)])}{\Phi[\operatorname{probit}(1,0)] /(1-\Phi[\operatorname{probit}(1,0)])} \tag{46}
\end{align*}
$$

Odds Ratio Effects Assuming a Rare Binary Outcome:

## Logistic Model

VanderWeele and Vansteelandt (2010) show that with logistic regression the TNIE odds ratio is approximately equal to

$$
\begin{equation*}
\operatorname{TNIE}(O R) \approx e^{\beta_{1} \gamma_{1}+\beta_{3} \gamma_{1}} \tag{47}
\end{equation*}
$$

that is, the indirect effect odds ratio uses the same formula as the indirect effect with a continuous outcome, but exponentiated. When the treatment variable is continuous, the indirect effect odds ratio of (47) is modified as

$$
\begin{equation*}
\operatorname{TNIE}(O R)=e^{\left(\beta_{1} \gamma_{1}+\beta_{3} \gamma_{1} x_{1}\right)\left(x_{1}-x_{0}\right)} \tag{48}
\end{equation*}
$$

for a change from $x_{0}$ to $x_{1}$. For example, $x_{0}$ may represent the mean of the treatment and $x_{1}$ may represent the mean plus one standard deviation, so that $x_{1}-x_{0}$ corresponds to one standard deviation for the continuous treatment variable.

Drug intervention program for students in Grade 6 and Grade 7 in Kansas City schools ( $n=864$ ). MacKinnon et al. (2007), Clinical Trials.

- Schools were randomly assigned to the treatment or control group (the multilevel aspect of the data is ignored)
- The mediator is the intention to use cigarettes in the following 2-month period which was measured about six months after baseline
- The outcome is cigarette use or not in the previous month which was measured at follow-up
- Cigarette use is observed for $18 \%$ of the sample
- The total effect can be computed without doing a mediation analysis as the difference between the proportion of smokers in the treatment group and the proportion of smokers in the control group
- This results in an estimate of the total effect as the difference in the probabilities of $0.148-0.224=-0.076$
- The corresponding estimate of the total effect odds ratio is

$$
\begin{equation*}
T E(O R)=\frac{0.148 /(1-0.148)}{0.224 /(1-0.224)}=0.602 \tag{49}
\end{equation*}
$$

- Both estimates indicate a lowering of the smoking probability due to treatment

Table : Input for smoking data using probit

| TITLE: | Clinical Trials data from MacKinnon et al. (2007) |
| :--- | :--- |
| DATA: | FILE = smoking.txt; |
| VARIABLE: | NAMES = intent tx ciguse; |
|  | USEVARIABLES = tx ciguse intent; |
|  | CATEGORICAL = ciguse; |
| ANALYSIS: | ESTIMATOR = ML; |
|  | LINK = PROBIT; |
|  | BOOTSTRAP = 10000; |
|  | ciguse ON intent tx; |
| MODEL: | intent ON tx; |
|  |  |
| MODEL INDIRECT: | ciguse IND intent tx; |
|  | TECH1 TECH8 SAMPSTAT |
| OUTPUT: | CINTERVAL(BOOTSTRAP); |
|  | TYPE = PLOT3; |

Table : Bootstrap confidence intervals for smoking data effects using probit regression for the outcome cigarette

| Confidence intervals of total, indirect, and direct effects based <br> on counterfactuals (causally-defined effects) |  |  |  |  |  |
| :--- | :---: | :---: | :---: | :---: | :---: |
|  | Lower 2.5\% | Lower 5\% | Estimate | Upper 5\% | Upper 2.5\% |
| Effects from tx to ciguse |  |  |  |  |  |
| Tot natural IE | -0.040 | -0.036 | -0.022 | -0.008 | -0.006 |
| Pure natural DE | -0.104 | -0.095 | -0.050 | -0.005 | 0.004 |
| Total effect | -0.128 | -0.119 | -0.072 | -0.026 | -0.017 |
|  | Odds ratios for binary Y |  |  |  |  |
| Tot natural IE | 0.757 | 0.772 | 0.853 | 0.939 | 0.958 |
| Pure natural DE | 0.520 | 0.551 | 0.731 | 0.969 | 1.025 |
| Total effect | 0.433 | 0.461 | 0.624 | 0.841 | 0.896 |

## Effects for Smoking Data Using Probit

- The total natural indirect effect (TNIE) in probability metric is estimated as -0.022 and is significant because the $95 \%$ confidence interval does not cover zero: $[-0.040,-0.006]$
- The indirect effect odds ratio is estimated as 0.853 and is significant because the $95 \%$ confidence interval does not cover one: [0.757, 0.958]
- The direct effect in probability metric is estimated as -0.050 and is not significant. The direct effect odds ratio of 0.731 is not significant
- The total effect in probability metric of -0.072 is significant
- The total effect can be compared to the proportion of cigarette users in the control group of 0.224 . This shows a drop of $34 \%$ due to treatment

Table : Input for smoking data using logistic regression for the cigarette use outcome

```
TITLE: Clinical Trials data from MacKinnon et al. (2007)
DATA: FILE = smoking.txt;
VARIABLE: NAMES = intent tx ciguse;
    USEVARIABLES = tx ciguse intent;
    CATEGORICAL = ciguse;
ANALYSIS: ESTIMATOR = ML;
    LINK = LOGIT;
    BOOTSTRAP = 10000;
MODEL: ciguse ON intent (beta1)
    tx (beta2);
    intent ON tx (gamma);
MODEL INDIRECT:
    ciguse IND intent tx;
MODEL CONSTRAINT:
    NEW(indirect direct);
    indirect = EXP(beta1*gamma);
    direct = EXP(beta2);
OUTPUT: TECH1 TECH8 SAMPSTAT
    CINTERVAL(BOOTSTRAP);
PLOT: TYPE = PLOT3;
```

- Not assuming a rare outcome (using MODEL INDIRECT): TNIE $(\mathrm{OR})=0.858$, $\operatorname{TNDE}(\mathrm{OR})=0.716$
- Assuming a rare outcome (using MODEL CONSTRAINT): $\operatorname{TNIE}(\mathrm{OR})=0.843, \operatorname{TNDE}(\mathrm{OR})=0.686$
- The rare outcome results indicate stronger effects with estimates farther from one
- The rare outcome assumption may not be suitable here with $18 \%$ smoking prevalence
- Probit and logistic give similar results


## Moderated Mediation with a Binary Outcome: Vaccination

- Hopfer (2012) analyzed data from a randomized control trial aimed at increasing the vaccination rate for the human papillomavirus (HPV) among college women ( $n=394$ )
- Subjects were randomized into three different intervention groups and a control group where the groups were presented with different forms of video with vaccine decision narratives
- The mediator measures intent to get vaccinated
- Control variables are HPV communication with parents (yes/no), age, sexually active (yes/no), and HPV knowledge
- Only the effects of the combined peer-expert intervention are considered ( $t \times 2$ )
- In this group, to which $25 \%$ of the sample was randomized, the vaccination rate is $22.2 \%$ whereas in the control group it is $12.0 \%$
- This gives an estimate of the total intervention effect in the probability metric of 0.10 and in the odds ratio metric of 2.70

Figure : Moderated mediation model for the HPV vaccination data using a logistic regression for the vaccination outcome


Table : Input for the model with intervention-mediator interaction for HPV vaccination data

```
VARIABLE:
    USEVARIABLES = intent4 tx1 tx2 tx 3 vacc hpvcomm age
    sxyes knowl mx;
    CATEGORICAL = vacc;
    MISSING = ALL (99);
DEFINE: mx = intent4*tx2;
    CENTER age knowl(GRANDMEAN);
ANALYSIS: ESTIMATOR = ML;
    BOOTSTRAP = 10000;
MODEL: vacc ON intent4 tx1 tx2 tx3 hpvcomm age sxyes knowl mx;
    intent4 ON tx1 tx2 tx3 hpvcomm age sxyes knowl;
MODEL INDIRECT:
    vacc MOD intent4 mx tx2;
OUTPUT: SAMPSTAT PATTERNS CINTERVAL(BOOTSTRAP)
    TECH1 TECH8;
PLOT: TYPE = PLOT3;
```

Table : Results for HPV vaccination data

|  |  |  |  |
| :--- | :--- | :--- | :--- |
| Estimate | S.E. | Est./S.E. | Two-Tailed <br> P-Value |


| vacc ON |  |  |  |  |
| :--- | ---: | ---: | ---: | ---: |
| intent4 | 1.303 | 0.262 | 4.974 | 0.000 |
| tx1 | 0.320 | 0.435 | 0.735 | 0.463 |
| tx2 | -1.180 | 2.271 | -0.520 | 0.603 |
| tx3 | -0.818 | 2.141 | -0.382 | 0.703 |
| hpvcomm | 0.242 | 0.350 | 0.693 | 0.488 |
| age | 0.194 | 0.084 | 2.311 | 0.021 |
| sxyes | 0.219 | 0.333 | 0.658 | 0.511 |
| knowl | -0.041 | 0.072 | -0.572 | 0.568 |
| mx | 0.494 | 0.660 | 0.749 | 0.454 |
| intent4 ON |  |  |  |  |
| tx1 | 0.149 | 0.106 | 1.400 | 0.161 |
| tx2 | 0.300 | 0.092 | 3.270 | 0.001 |
| tx3 | -0.066 | 0.141 | -0.465 | 0.642 |
| hpvcomm | 0.093 | 0.078 | 1.196 | 0.232 |
| age | -0.049 | 0.021 | -2.283 | 0.022 |
| sxyes | 0.044 | 0.078 | 0.573 | 0.567 |
| knowl | -0.003 | 0.017 | -0.160 | 0.873 |
| Intercepts |  |  |  |  |
| intent4 | 2.718 | 0.082 | 32.959 | 0.000 |
| Thresholds |  |  |  |  |
| vacc $\$ 1$ | 6.227 | 0.877 | 7.100 | 0.000 |
| Residual Variances |  |  |  |  |
| intent4 | 0.591 | 0.041 | 14.293 | 0.000 |

# Table : Bootstrap confidence intervals without and with intervention-mediator interaction for HPV vaccination data 

| Confidence intervals of total, indirect, and direct effects based on counterfactuals (causally-defined effects) |  |  |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: |
|  | Lower 2.5\% | Lower 5\% | Estimate | Upper 5\% | Upper 2.5\% |
| Without intervention-mediator interaction |  |  |  |  |  |
| Effects from TX2 to VACC |  |  |  |  |  |
| Tot natural IE | 0.016 | 0.020 | 0.048 | 0.083 | 0.092 |
| Pure natural DE | -0.019 | -0.010 | 0.041 | 0.098 | 0.111 |
| Total effect | 0.013 | 0.024 | 0.089 | 0.165 | 0.182 |
| Odds ratios for binary Y |  |  |  |  |  |
| Tot natural IE | 1.155 | 1.197 | 1.448 | 1.833 | 1.932 |
| Pure natural DE | 0.803 | 0.894 | 1.523 | 2.715 | 3.045 |
| Total effect | 1.137 | 1.283 | 2.205 | 4.115 | 4.665 |
| With intervention-mediator interaction |  |  |  |  |  |
| Effects from TX2 to VACC |  |  |  |  |  |
| Tot natural IE | 0.016 | 0.020 | 0.056 | 0.099 | 0.109 |
| Pure natural DE | -0.022 | -0.012 | 0.037 | 0.095 | 0.107 |
| Total effect | 0.016 | 0.028 | 0.093 | 0.169 | 0.186 |
| Odds ratios for binary Y |  |  |  |  |  |
| Tot natural IE | 1.147 | 1.200 | 1.541 | 2.096 | 2.238 |
| Pure natural DE | 0.773 | 0.865 | 1.467 | 2.662 | 2.964 |
| Total effect | 1.178 | 1.313 | 2.260 | 4.234 | 4.791 |

Figure : Bootstrap distribution for the total natural indirect effect estimate in probability metric for the model with intervention-mediator interaction for the HPV vaccination data


## Mediation with a Count Outcome: $Y$ is the Log Rate



$$
\begin{align*}
\log \mu_{i} & =\beta_{0}+\beta_{1} M_{i}+\beta_{2} X_{i}+\beta_{3} M X_{i}+\beta_{4} C_{i}  \tag{50}\\
M_{i} & =\gamma_{0}+\gamma_{1} X_{i}+\gamma_{2} C_{i}+\varepsilon_{m i} . \tag{51}
\end{align*}
$$

As before, the counterfactually-based causal effects consider terms such as

$$
\begin{align*}
E\left[Y\left(x_{1}, M\left(x_{0}\right)\right)\right] & =\int_{-\infty}^{\infty} E\left[Y \mid C=c, X=x_{1}, M=m\right]  \tag{52}\\
& \times f\left(M \mid C=c, X=x_{0}\right) \partial M \tag{53}
\end{align*}
$$

This needs to take into account that the rate (mean) is

$$
\begin{equation*}
E\left[Y \mid C=c, X=x_{1}, M=m\right]=e^{\beta_{0}+\beta_{1} m+\beta_{2} x_{1}+\beta_{3} m x_{1}+\beta_{4} c} \tag{54}
\end{equation*}
$$

## School Removal Count Outcome: Case 3 ( $m x$ ) Moderation



Randomized field experiment in Baltimore public schools with a classroom-based intervention aimed at reducing aggressive-disruptive behavior among elementary school students (Kellam et al., 2008). The analysis uses $n=250$ boys.

- The outcome variable remove is the number of times a student has been removed from school during grades 1-7
- tx is the binary exposure variable representing the intervention
- The Fall baseline aggression score is agg1 which was observed before the intervention started
- The mediator variable agg5 is the Grade 5 aggression score.
- An intervention-mediator interaction variable $m x$ is included to moderate the influence of the mediator on the outcome.

Table : Input for negative binomial model for school removal data

```
VARIABLE:
    USEVARIABLES = remove agg5 agg1 tx mx;
    IDVARIABLE = prcid;
    COUNT = remove(NB);
    USEOBSERVATIONS = gender EQ 1 AND (desgn11s EQ 1 OR
    desgn11s EQ 2 OR desgn11s EQ 3 OR desgn11s EQ 4);
DEFINE: IF(desgn11s EQ 4)THEN tx=1;
    IF(desgn11s EQ 1 OR desgn11s EQ 2 OR desgn11s EQ 3)THEN
    tx=0;
    remove = total17;
    agg1 = sctaa11f;
    agg5 = sctaa15s;
    CENTER agg1 agg5(GRANDMEAN);
    mx = agg5*tx;
ANALYSIS: ESTIMATOR = ML;
    BOOTSTRAP = 10000;
    PROCESSORS = 8;
MODEL: remove ON agg5 tx mx agg1;
    agg5 ON tx agg1;
MODEL INDIRECT:
    remove MOD agg5 mx tx;
OUTPUT: SAMPSTAT TECH1 TECH8 PATTERNS
    CINTERVAL(BOOTSTRAP);
PLOT: TYPE = PLOT3;
```


## Table : Bootstrap confidence intervals for effects for school removal data

| Confidence intervals of total, indirect, and direct effects based <br> on counterfactuals (causally-defined effects) |  |  |  |  |  |  |  |  |
| :--- | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  | Lower 2.5\% | Lower 5\% | Estimate | Upper 5\% | Upper 2.5\% |  |  |  |
| Effects from TX to REMOVE |  |  |  |  |  |  |  |  |
| Tot natural IE | -0.341 | -0.283 | -0.119 | -0.024 | -0.010 |  |  |  |
| Pure natural DE | -0.681 | -0.608 | -0.272 | 0.125 | 0.213 |  |  |  |
| Total effect | -0.794 | -0.722 | -0.391 | -0.032 | 0.034 |  |  |  |
|  | Other effects |  |  |  |  |  |  |  |
| Pure natural IE | -0.358 | -0.327 | -0.183 | -0.050 | -0.023 |  |  |  |
| Tot natural DE | -0.587 | -0.525 | -0.208 | 0.135 | 0.213 |  |  |  |
| Total effect | -0.794 | -0.722 | -0.391 | -0.032 | 0.034 |  |  |  |

Figure : Total natural indirect effect bootstrap distribution for school removal data


- The indirect effect estimate -0.119 is in a log rate metric for the count outcome of school removal and is hard to interpret
- One way to make the effect size understandable is to compute the probability of a zero count
- The intervention increases the probability of a zero school removals from 0.294 to 0.435


## Two-Part Mediation Modeling

- Example from Hayes (2013):
- $n=262$ small-business owners' economic stress (Pollack et al., 2011)
- The exposure variable is a continuous variable representing economic stress
- The mediator variable is a continuous variable representing depressed affect
- The outcome variable is a continuous variable representing thoughts about withdrawing from their entrepreneurship

The outcome variable withdraw has a $30 \%$ floor effect:


## Table : Input for two-part mediation modeling of economic stress data

```
TITLE: Hayes ESTRESS example, cont's X
DATA: FILE = estress.txt;
VARIABLE: NAMES = tenure estress affect withdraw sex age ese;
    USEVARIABLES = affect estress u y;
    CATEGORICAL = u;
DEFINE: withdraw = withdraw - 1;
DATA TWOPART:
    NAMES = withdraw;
    BINARY = u;
    CONTINUOUS = y;
    CUTPOINT = 0;
ANALYSIS: ESTIMATOR = ML;
    LINK = PROBIT;
    BOOTSTRAP = 1000;
MODEL: y ON affect (betal)
estress (beta2);
[y] (beta0);
y (v);
affect ON estress (gamma1);
[affect] (gamma0);
affect (sig);
u ON affect (kappal)
estress (kappa2);
[u$1] (kappa0);
MODEL INDIRECT:
u IND affect estress (6.04 4.62);
-table continues-
```


## Table : Input for two-part mediation modeling of economic stress data

```
MODEL CONSTRAINT:
NEW(xl x0 eyl ey0 muml mum0 ayl ay0 bymll byml0 bym01
bym00 eym11 eym10 eym01 eym00 tnie pnde total pnie beta3 sd pill
pil0 pi01 pi00);
beta3 = 0;
x1=6.04;
x0=4.62;
eyl=EXP(v/2)*EXP(beta0+beta2*x1);
ey0=EXP(v/2)*EXP(beta0+beta2*x0);
muml=gamma0+gammal*xl;
mum0=gamma0+gamma1*x0;
ayl=sig*(beta1+beta3*x1);
ay0=sig*(betal+beta3*x0);
bym11=(ayl/muml+1);
bym10=(ay1/mum0+1);
bym01=(ay0/mum1+1);
bym00=(ay0/mum0+1)
sd=SQRT(kappa1*kappa1*sig+1);
pil1=PHI((-kappa0+kappa2*xl+kappa1*bym11*
(gamma0+gammal*xl))/sd);
pi10=PHI((-kappa0+kappa2*x 1 +kappa1*bym10*
(gamma0+gammal*x0))/sd);
pi01=PHI((-kappa0+kappa2*x0+kappal*bym1 1*
(gamma0+gammal*x1))/sd);
pi00=PHI((-kappa0+kappa2*x0+kappa1*bym00*
(gamma0+gammal*x0))/sd);
eym11=EXP((bym11*bym11-1)*muml*muml/(2*sig));
eym10=EXP((bym10*bym10-1)*mum0*mum0/(2*sig));
eym01=EXP((bym01*bym01-1)*muml*muml/(2*sig));
eym00=EXP((bym00*bym00-1)*mum0*mum0/(2*sig));
tnie=pi11*ey1*eym11-pi10*ey1*eym10;
pnde=pi10*ey1*eym10-pi00*ey0*eym00;
total=pi11*ey1*eym11-pi00*ey0*eym00;
pnie=pi01*ey0*eym01-pi00*ey0*eym00;
PLOT:
OUTPUT: SAMPSTAT TECH1 TECH8
CINTERVAL(BOOTSTRAP);
```


## Table : Bootstrap confidence intervals for four mediation models

| Confidence intervals for effects |  |  |  |  |  |
| :--- | :---: | :---: | :---: | :---: | :---: |
| Lower 2.5\% |  |  |  |  |  |
| Lower 5\% | Estimate | Upper 5\% | Upper 2.5\% |  |  |
| (1) Two-part: overall effects for continuous part of the outcome |  |  |  |  |  |
| TNIE | 0.104 | 0.121 | 0.203 | 0.293 | 0.311 |
| PNDE | -0.304 | -0.276 | -0.145 | -0.011 | 0.019 |
| TE | -0.124 | -0.089 | 0.058 | 0.207 | 0.246 |
| (2) Two-part: effects for binary part of the outcome |  |  |  |  |  |
| TNIE | 0.036 | 0.041 | 0.071 | 0.103 | 0.108 |
| PNDE | -0.074 | -0.062 | -0.016 | 0.028 | 0.035 |
| TE | -0.006 | 0.005 | 0.055 | 0.098 | 0.105 |
| (3) Two-part: conditional effects for continuous part of the outcome |  |  |  |  |  |
| TNIE | 0.043 | 0.053 | 0.112 | 0.177 | 0.194 |
| PNDE | -0.322 | -0.299 | -0.160 | -0.008 | 0.023 |
| TE | -0.219 | -0.184 | -0.048 | 0.105 | 0.131 |
| (4) Regular: effects using log $y$ |  |  |  |  |  |
| TNIE | 0.098 | 0.108 | 0.182 | 0.267 | 0.284 |
| PNDE | -0.236 | -0.209 | -0.084 | 0.044 | 0.066 |
| TE | -0.072 | -0.045 | 0.099 | 0.243 | 0.269 |
| (5) Regular: effects using the original $y$ |  |  |  |  |  |
| TNIE | 0.103 | 0.117 | 0.189 | 0.266 | 0.282 |
| PNDE | -0.263 | -0.243 | -0.109 | 0.027 | 0.051 |
| TE | -0.116 | -0.069 | 0.080 | 0.220 | 0.245 |


[^0]:    ${ }^{1} 10-15$ minutes of questions and answers at the end of each block (hold your questions).

