

US EPA ARCHIVE DOCUMENT

# Methodological Issues

# Effect modification: limits of conventional approach

- Low statistical power
- Limited functional forms
- Difficulty interpreting 3+ way interactions
- Misspecified “main” effects
- “Statistical interaction formulations are inadequate to capture the ecology of human development”\* [page 924](#)

\* Source: Evans GW. Dev Psychol 2003, 39:924-33.

# Alternatives to interaction terms

- Cumulative risk model (Rutter 1983)
  - Sum discrete risk using standard threshold cutoffs
- Decision tree analysis (Breiman, 2001)
  - Assumes no explicit causal model, fully capture complex interactions
- Systems dynamics models (Galea, 2009)
  - Feed back, non-linearities
- Hierarchical (aka multi-level) models (Raudenbush & Bryk, 2002)
  - Nested data; cross-level interactions; random slopes to model risk heterogeneity
  - Model the social ecology of risk

# Example: Multilevel (hierarchical models) for differential vulnerability

- Hypothesis: The effect of air pollution ( $PM_{10}$ ) is exacerbated for residents of high crime neighborhoods due to prolonged exposure to psychosocial hazards
- Clustered data
  - 1000 individuals
  - 50 neighborhoods/communities
- Individually monitored  $PM_{10}$  exposure
  - High (e.g., 90<sup>th</sup> percentile) vs. not-high

# Model 1:

## Does risk vary by social ecology?

Level-1 model (individual-level)



$Y_{ij}$  is the systolic blood pressure of the  $i$ th person in the  $j$ th neighborhood where:

$r_{ij}$  = random error associated with  $i$ th person in  $j$ th neighborhood  
 $\sim N(0, \sigma^2)$

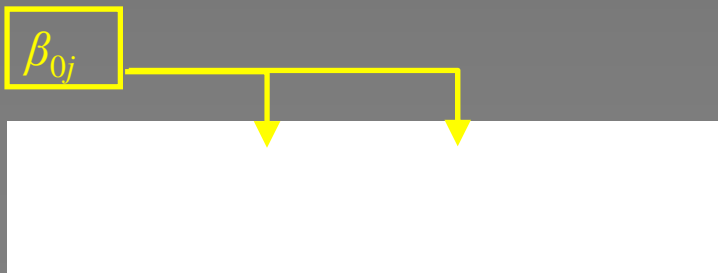
Level-2 model (neighborhoods)



$\beta_{0j}$  is the neighborhood-specific intercept where:

$\gamma_{00}$  = the overall mean SBP across all NBs.  
 $u_{0j}$  = a series of random deviations from the mean  $\sim N(0, \tau_{00})$

Multilevel model (mixed effects)

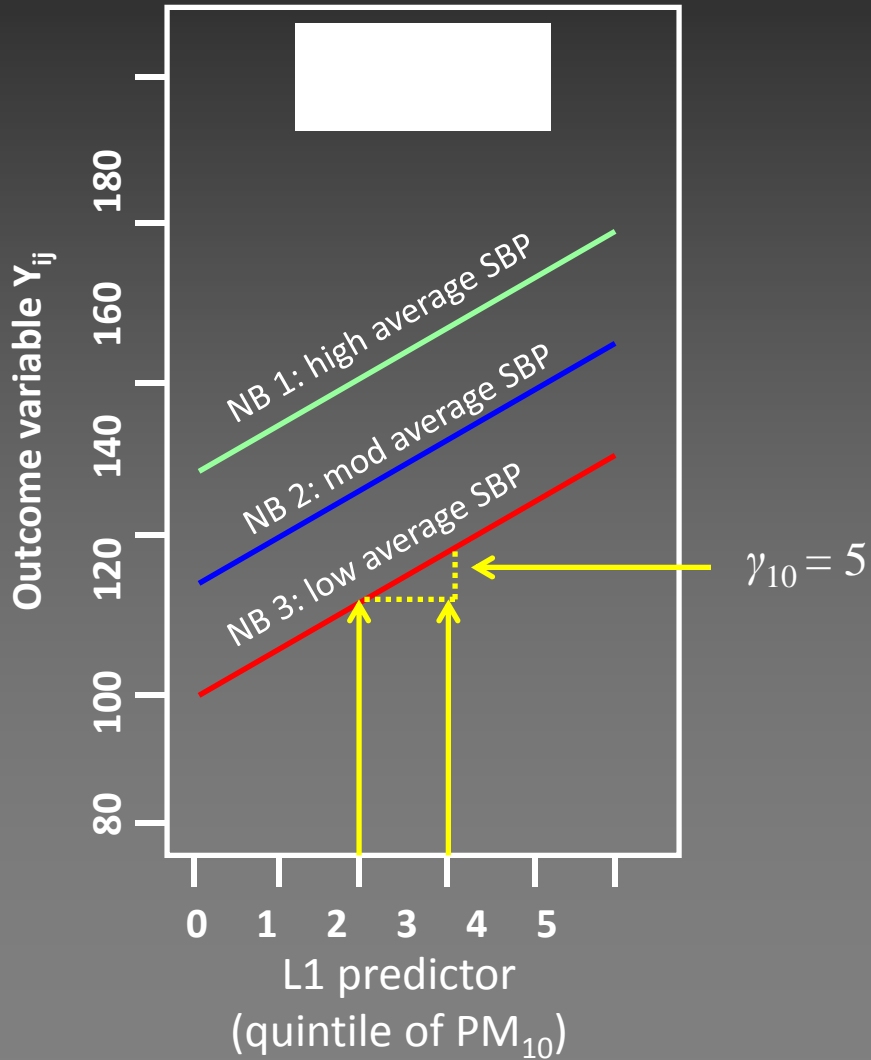


where:

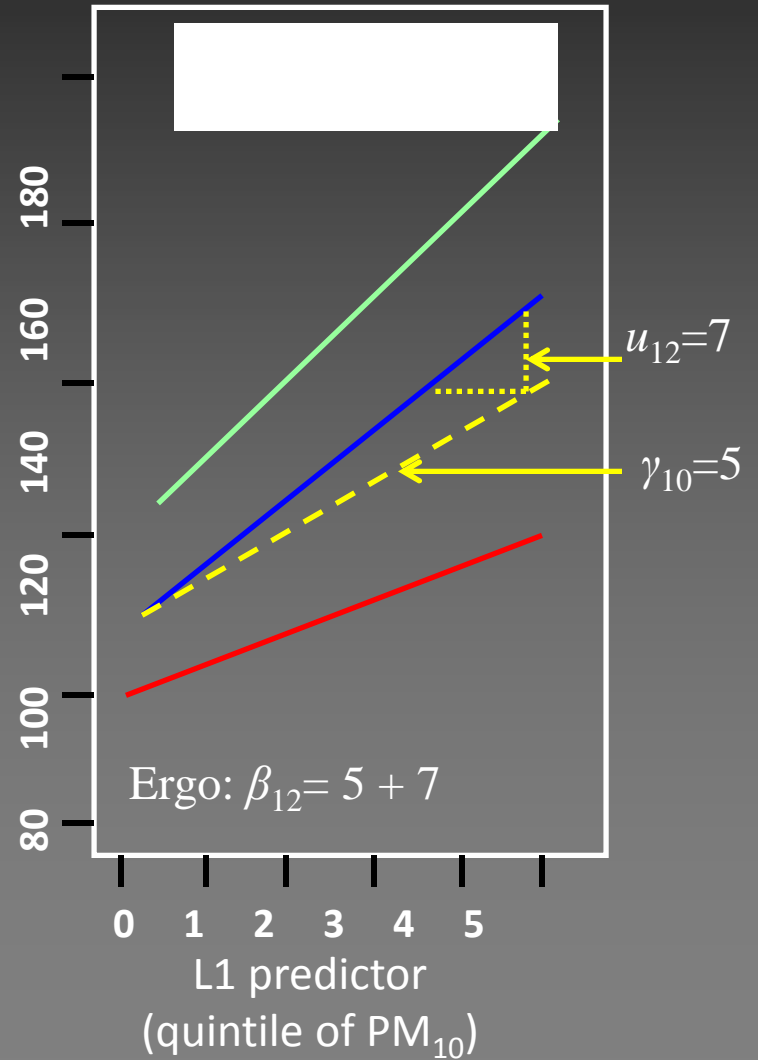
$$\text{cov}(r_{ij}, u_{0j}) = 0$$

# Visualizing random intercepts and slopes

Random intercepts:

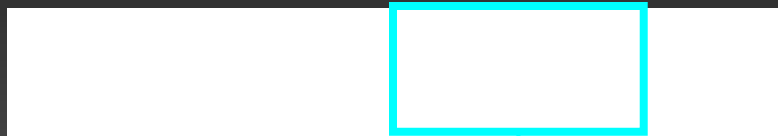


Random intercepts and slopes:



# Model 2: Modeling cross-level interactions

Level-1 model (individual-level)



$PM_{10ij}$  is a covariate coded 1 if  $i$ th person in  $j$ th neighborhood is exposed to high levels of PM10, 0 if not

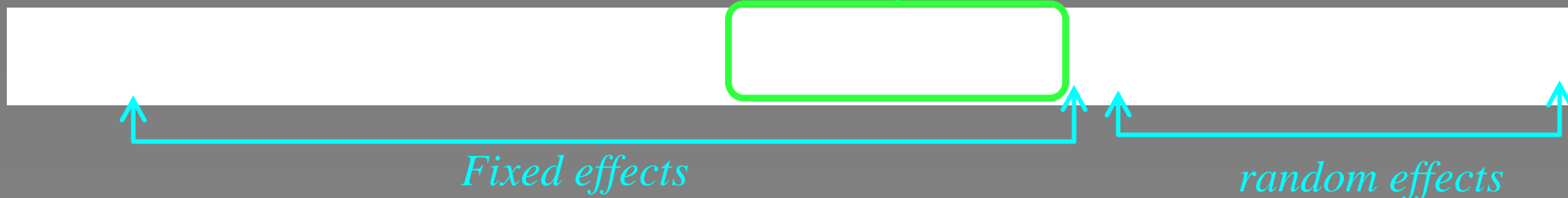
Level-2 model (neighborhood level)



$\gamma_{01}$  is the mean difference in SBP associated with a 1 standard deviation increase in crime rate in those not exposed to high PM<sub>10</sub>

$\gamma_{11}$  is extent to which the marginal change in SBP for those living in a high crime NB among exposed vs. non-exposed (implicit cross-level interaction)

Multilevel model:



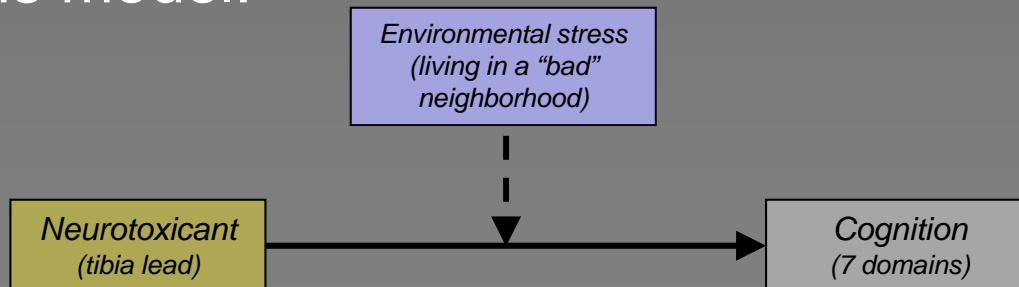
*Fixed effects*

*random effects*



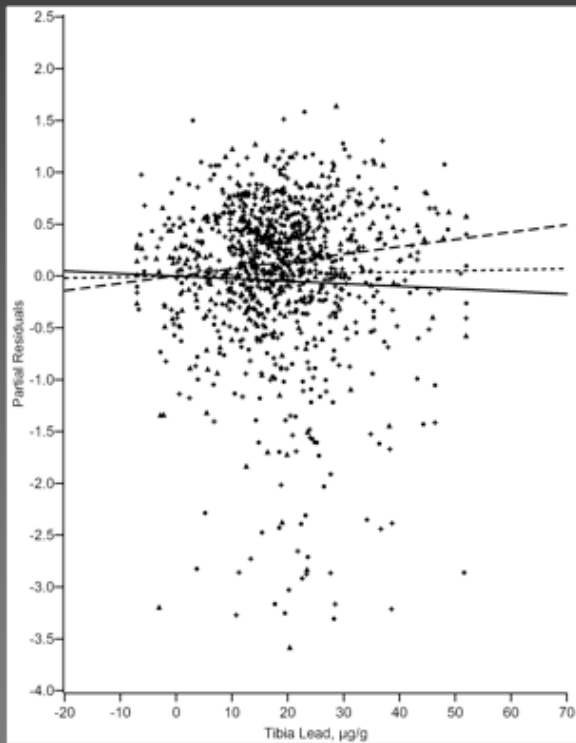
# Real world example: Environmental stress, lead and cognition

- Animal models show environmental stress worsens lead effect on brain/cognition
- The Baltimore Memory study
  1. 1140 community-dwelling adults aged 50-70 in 65 contiguous Baltimore neighborhoods
  2. Tibia lead measured using XRF spectroscopy
  3. Measure “environmental stress” (**toxicology term**) with a scale of neighborhood psychosocial hazards (**social epidemiology term**)
- Test this model:

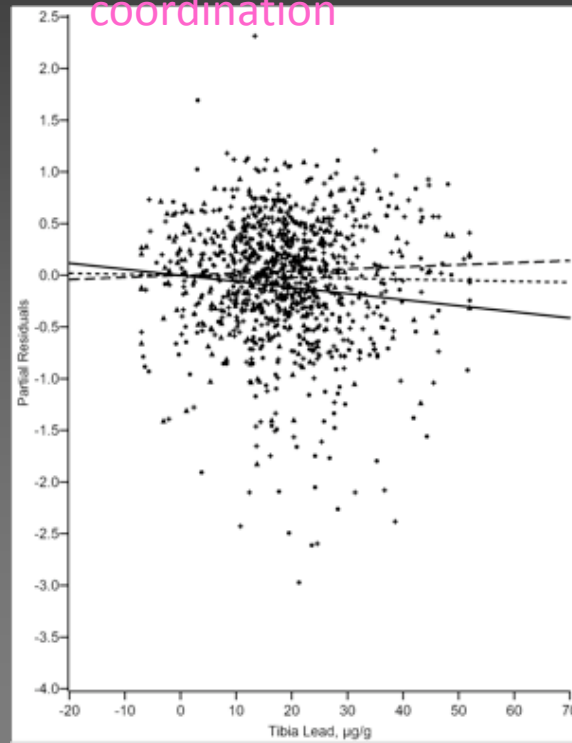


# Neighborhood psychosocial hazards exacerbate association of tibia lead on cognition

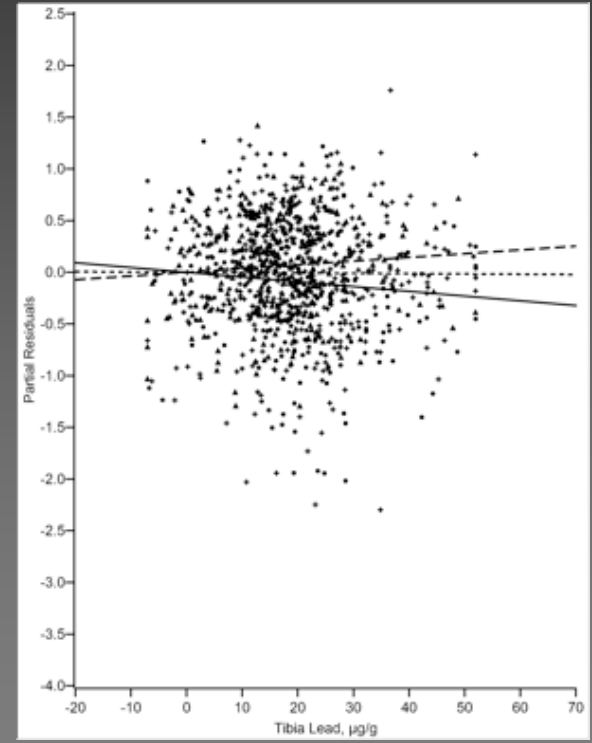
Processing speed



Eye-hand coordination



Executive ability



Source: Glass TA, et al. AJE 2009, 169:683-92.

Legend:

- Highest tertile NPH score = —————
- Middle tertile NPH score = .....
- Lowest tertile NPH score = - - - - -

# Living beyond our means: Why not Measure what we Want?

- Traditional Regression analysis models the mean response in the population
- The risk in the population may be high for a small subset
- Quantile regression:
  - directly estimate the effect on 95th percentile of risk, rather than on the mean risk
  - Modeling multiple quantiles estimates the change in the distribution