Current Practice for Addressing Measurement Error in Epidemiologic Studies

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Outline

- Background
- Motivating Example
- Literature survey methodology
- Literature survey results
- Conclusions

STRATOS TG4: Measurement Error and Misclassification

MEMBERSHIP

- Laurence Freedman, Gertner/IMS, Co-Chair
- Victor Kipnis, NCI, Co-Chair
- Raymond Carroll, Texas A&M U
- Veronika Deffner, Munich, LMU
- Kevin Dodd, NCI
- Paul Gustafson, U. British Columbia
- Ruth Keogh, London School of Hygiene
- Helmut Kuechenhoff, Munich, LMU
- Pamela Shaw, U. Pennsylvania
- Janet Tooze, Wake Forest School of Medicine

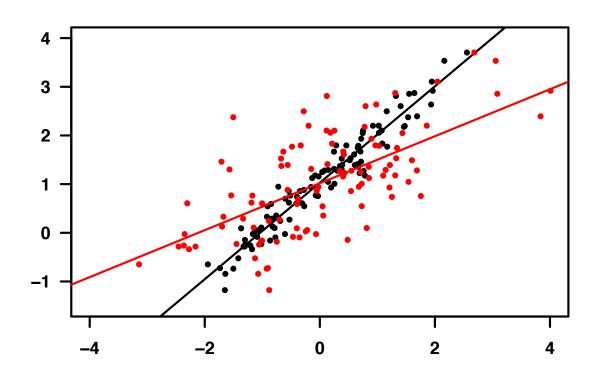
TG4 Projects

- 1. Literature Survey for how measurement error is addressed in 4 types of epidemiological studies
- 2. Guidance paper for nutritional epidemiologists
- 3. Guidance paper for biostatisticians

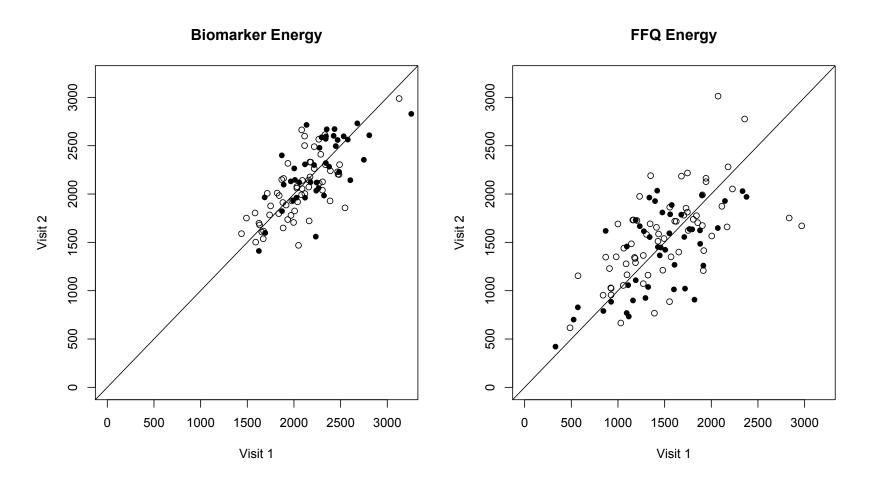
Background

- There have been many statistical advances to address in measurement error in the past few decades
- TG4 was interested in assessing the current practice for acknowledging and addressing measurement error in epidemiologic/observational studies
 - Want to identify knowledge gaps and opportunities for improvement
- We conducted a literature survey focused on types of epidemiologic studies with exposures that are well known to be subject to measurement error

The Problem with Covariate Measurement Error...



Nutritional Epidemiology Example: Measuring Energy Intake



Energy Intake vs Body Mass Index

Neuhouser et al AJE 2008

APPENDIX TABLE. Estimates of energy intake (kcal/day) obtained by self-reported food frequency questionnaire, a biomarker (total energy expenditure), and a calibrated food frequency questionnaire, according to body mass index category, Women's Health Initiative Nutritional Biomarkers Study, 2004–2005*

Body mass index†	Self-reported FFQ‡		Total energy expenditure		Calibrated FFQ	
category	Geometric mean	IQR‡	Geometric mean	IQR	Geometric mean	IQR
Nomal (<25.0)	1,407	1,157-1,759	1,894	1,714-2,083	1,912	1,853-1,980
Overweight (25.0-29.9)	1,462	1,196-1,837	2,043	1,904-2,232	2,028	1,962-2,103
Obese (≥30)	1,454	1,161–1,897	2,213	2,034-2,415	2,247	2,156–2,338

^{*} Note that the difference between FFQ energy intake (self-report) and total energy expenditure (biomarker) increases as body mass index increases. The biomarker-calibrated estimates, for the same women, correct for the measurement error using the model shown in table 4.

[†] Weight (kg)/height (m)².

[‡] FFQ, food frequency questionnaire; IQR, interquartile range (25th-75th percentiles).

Regression Calibration: A simple approach to adjust for ME

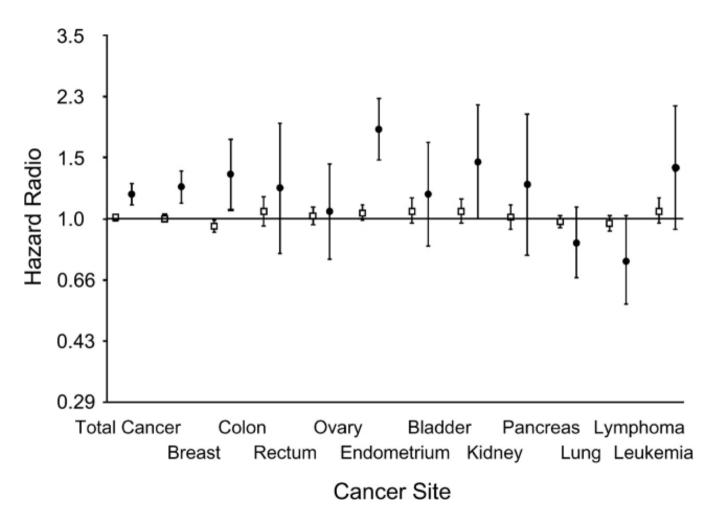
Prentice Biometrika 1982

- Suppose true intake: X
- Error-prone measure: Q
- Objective biomarker: W = X + u
- Predicted X = E(W|Q,Z) = E(X|Q,Z)

$$= a_1 + a_2 Q + a_3 Z + a_4 Z Q$$

Regression calibration: Regress outcome Y on predicted intake, other covariates Z

HR for Uncalibrated vs Calibrated Energy Intake Prentice, Shaw et al AJE 2009



Survey Areas

Each of four topic areas had its own literature search

- Nutritional intake cohort studies (Pamela Shaw/Ruth Keogh)
- Dietary intake population surveys (Kevin Dodd)
- Physical activity cohort studies (Janet Tooze)
- Air pollution cohort studies (Veronika Deffner/Helmut Kuechenhoff)

Overall Approach

- Focused on error-prone variable as exposure in analysis
- For cohort studies, literature search done in two stages
 - Search A: Survey recent articles to assess how often articles acknowledged and/or addressed measurement error
 - Search B: Survey recent articles that adjusted for measurement error to describe methods in current practice
- Questionnaires filled out for each reviewed article
- Excluded reviews, cross-sectional studies, case-control studies and meta-analyses
- Each topic area conducted a quality control review
 - 20% re-reviewed by independent reviewer

Nutritional Epidemiology Cohort Studies: Survey Methodology

- Date Range A: Feb 2014-Jun 2015; B:Jan 2001-Jul 2015
- Limited search to three common diseases with dietary risk factors: cancer, heart disease and diabetes
 - Restricted date range to find about 50 articles from Search A and 30 articles from Search B
- Search B: added (measurement error OR misclassification to Search A
 - Not many articles, so did additional key word searches including: (measurement error OR misclassification) AND nutritional epidemiology

Dietary Intake Population Studies: Survey Methodology

- Date range: Jan 2012 May 2015
- Term "Measurement error" not typically referred to in dietary intake surveys
 - Understood as variance around usual intake
 - Conducted Search A only

Physical Activity Cohort Studies: Survey Methodology

- Date range: Jan 2012 Sep 2015
- Search A: Very broad search terms: N=8760 from search; randomly selected N=610; N=51 from abstract review
- SEARCH B: Added "measurement error" OR misreport* OR misclassif* OR bias OR attenuat* OR calibrat*
 - N=610 from search; N=86 from abstract review

Air Pollution Cohort Studies: Survey Methodology

- Date range: Jan 2012 Dec 2014
- Search A broad search within "Web of Science":
 - Search B Additional keywords: "measurement error",
 "measurement uncertainty", misclassif*, attenuat*
 - A: 4595 hits, B: 386 hits
- After abstract review: A: 431 hits, B: 32 hits
- Random selection: Search A: 50/Search B:25

Number of Articles Reviewed*

	Search A	Search B
Nutritional Epidemiology cohort studies	51	27
Dietary Intake Population Survey	67	N/A
Physical Activity cohort studies	30	40
Air Pollution cohort studies	50	25

^{*} Number in table excludes articles that were identified by search terms but upon closer examination did not meet inclusion criteria

Search A Survey Results

Mention ME as potential as potential problem n(%) 48 (94%) 17 (57%) 53/67 (79%) 20 (40%) Used a method to adjust for ME N (%) 5 (10%) 0 (0%) 19/67 (28%) 3 (6%) ME N (%) Any Primary exposure Primary exposure Exclusively 21/30 (70%) 21/30 (70%) Statistic of main interest N (%) HR 45 (88%) HR 11 (37%) Mean 51 (76%) OR/RR 9(30%) Median 28(42%) RR 2 (4%) GLM 5 (17%) %-tiles 21(31%) Slope 5(10%) Other 5 (17%) Quality 31(46%)		Nutritional Epi Cohort N= 51	Phys activity Cohort N=30	Diet Intake Survey N=67	Pollution Cohort N=50
to adjust for 5 (10%) 0 (0%) 19/67 (28%) 3 (6%) ME N (%) % categorizing Any Primary exposure 50/51(98%) exposure Exclusively 21/30 (70%) 27/51 (53%) Statistic of HR 45 (88%) HR 11 (37%) Mean 51 (76%) main interest OR 3 (6%) OR/RR 9(30%) Median 28(42%) N (%) RR 2 (4%) GLM 5 (17%) %-tiles 21(31%)	as potential	48 (94%)	17 (57%)	53/67 (79%)	20 (40%)
% categorizing exposure Any 50/51(98%) exposure Exclusively 21/30 (70%) 21/30 (70%) Statistic of main interest N (%) HR 45 (88%) HR 11 (37%) Mean 51 (76%) OR/RR 9(30%) Median 28(42%) GLM 5 (17%) %-tiles 21(31%)	to adjust for ME	5 (10%)	0 (0%)	19/67 (28%)	3 (6%)
Statistic of main interest HR 45 (88%) HR 11 (37%) Mean 51 (76%) OR 3 (6%) OR/RR 9(30%) Median 28(42%) RR 2 (4%) GLM 5 (17%) %-tiles 21(31%)	% categorizing	50/51(98%) Exclusively	exposure		
	main interest	HR 45 (88%) OR 3 (6%) RR 2 (4%)	OR/RR 9(30%) GLM 5 (17%)	Median 28(42%) %-tiles 21(31%)	

Methods to Address Measurement Error

Nutritional Epi Cohort N= 27*	Phys Activity Cohort N=40	Dietary Intake Pop. Survey N=67	Pollution Cohort N = 25
Regression Calib. 26 (96%) SIMEX 1 (4%) Other 1 (4%)	Regression Calib. 1(50%) Other 1 (50%)		4 (80%) Instr Variables
Search A: None 90%	Search A: None 95%	Search A: None 72%	Search A: None 94%

- Number excludes articles that were identified by search terms but upon review did not use a method to correct for error.
- Row percents do not add to 100% due to use of multiple methods.

Other Observations from Diet an Physical Activity Cohort Surveys

- Common in the cohort studies to have multiple covariates with error: eg diet + physical activity, smoking, and/or alcohol intake
 - Many adjust for both diet+ PA, only 1 article adjusted for error in both physical activity (Zhang et al, AJE 2014)
 - Errors in smoking/alcohol not addressed
- Most categorized the continuous exposures
 - Impacts of categorizing an exposure subject to error are ignored
 - Common belief: categorization will lower impact of measurement error in the analysis
- Most people who mentioned error as a problem made an incomplete/incorrect claim
 - Many only mentioned attenuation in found associations
 - Some claimed no bias in associations since prospective subject recall
 - Some claimed no bias since instrument was validated

Other observations from Dietary Intake Population Surveys

- Most studies (80%) used 24HR as primary instrument
 - 31/53 used only 1 24HR, rest had repeats on at least a subsample
 - 8/31 (26%) reported percentiles subject to bias
- 16/31 papers with 1 24HR mentioned that usual intake or adjustment for within-person variation was needed
- 8/11 (73%) of papers using multiple 24HRs to report medians/percentiles, used a complex method (NCI/MSM)

Other Observations from the Air Pollution Cohort Survey

- Statements about the measurement error are often vague
 - The origin of the measurement error is often not clearly specified
 - The size and the impact of the measurement error is often not stated
- Measurement error is often mentioned but rarely addressed in detail or corrected
 - The majority of the studies use daily and spatially aggregated data
 - The often prevailing Berkson error (through temporal and spatial aggregation) is not or only insufficiently described and its implications are not discussed
 - Errors originating from staying in different microenvironments are often neglected or only poorly considered
- Many different exposure measures are analyzed separately or jointly; a homogeneous procedure is lacking

Conclusions

- In cohort studies: measurement error acknowledged, but implications not fully understood and commonly not addressed in statistical analysis
 - Very few used methods to adjust for measurement error
 - For PA studies, little motivation to adjust for error since the naïve associations are generally aligned with a priori hypotheses
 - Many studies had multiple variables measured w/error
- In dietary intake population surveys: minority corrected for measurement error
 - Majority of those that did apply a correction method were taking advantage of software (e.g. NCI method)
- Regression calibration most common method to address measurement error in diet and PA studies

More work is needed....

- Identify the various sources of measurement error
- Disseminate ideas of measurement error correction
 - Discussion of software in guidance documents, tutorials in clinical journals, talks at epi and clinical conferences
- Correct misconceptions, such as:
 - Random error won't cause bias in associations
 - Attenuation is the only possible direction of bias
 - Categorization reduces the effect of measurement error
 - Validated questionnaires don't have bias
 - Software is not available

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Regression Calibration

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