

**Graphically presenting associations of related
but distinct phenomena:
moving beyond p-values**

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Background: Increasing importance of multifactorial measures

- Exposures and/or outcomes increasingly multifactorial
 - -Omics studies: genes, proteins, metabolites
 - Nutrition: nutrients and/or foods
 - Social sciences: crimes
- Increases in computing power allow research to precede hypothesis formulation– ‘fishing expeditions’
- Multiple comparisons = increased risk of random false positives.

Distinguishing false from true positives: current options

Clinical judgment

→ Useful only if hypotheses already exist

Adjustment of p-value for significance

- Statistical correction: Bonferroni, Benjamini-Hochberg...
- Graphical visualization: Manhattan plots

→ Costs statistical power: increases false negatives

→ Does not address effect size

→ Does not address global null ('Is there a genetic effect?')

My method: Graph sample size against effect size (similar to a funnel plot)

Answers two questions:

→ Is there a main **effect?** → "Association" – maybe not causal

- Is the plot centered on 'no effect'?

→ Is there effect heterogeneity (subgroup effects)?

- Is the plot spread out / asymmetrical?
- Are extreme-valued points similar in some way?

Example: Risk of crime in LA: effects of lunar phase or victim sex

→ Is there a main effect?

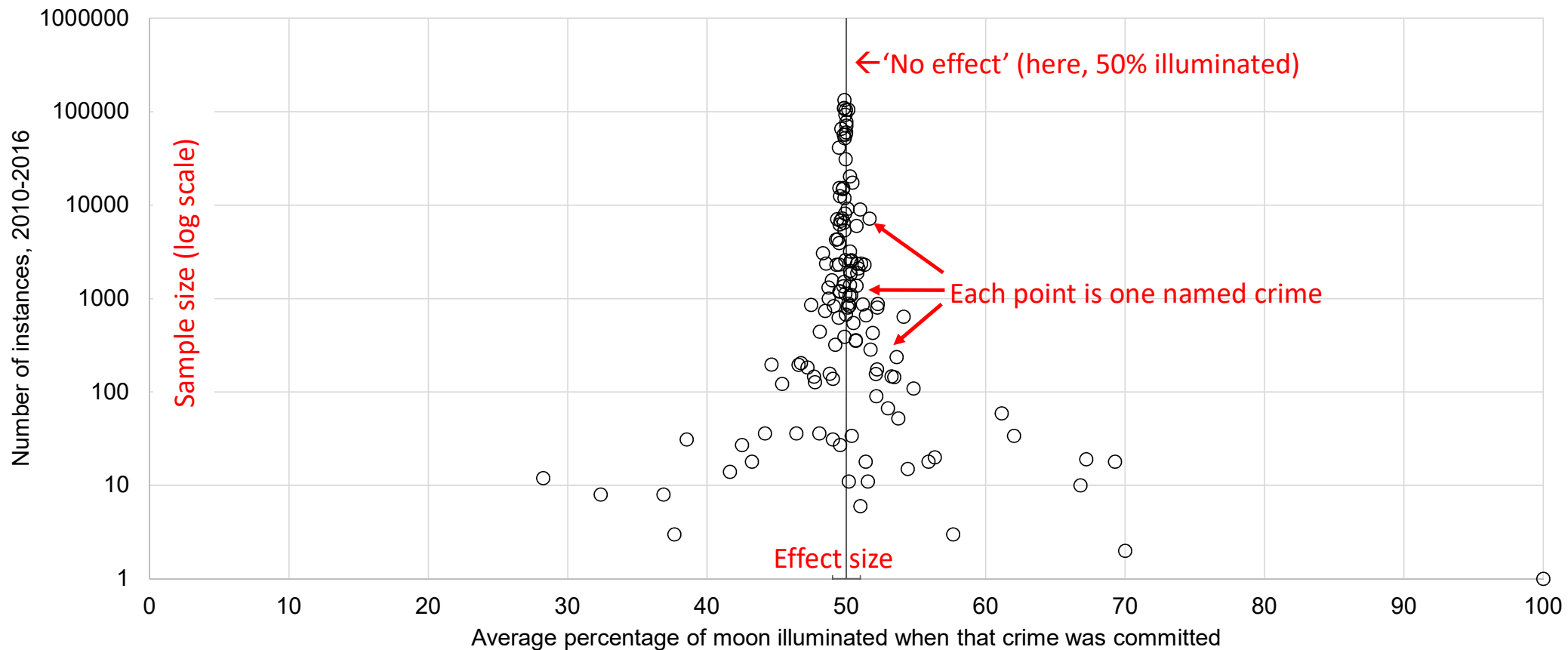
1. Does overall crime rate vary with lunar phase?
2. Is one sex at greater risk of victimization?

→ Is there effect heterogeneity / subgroup effects?

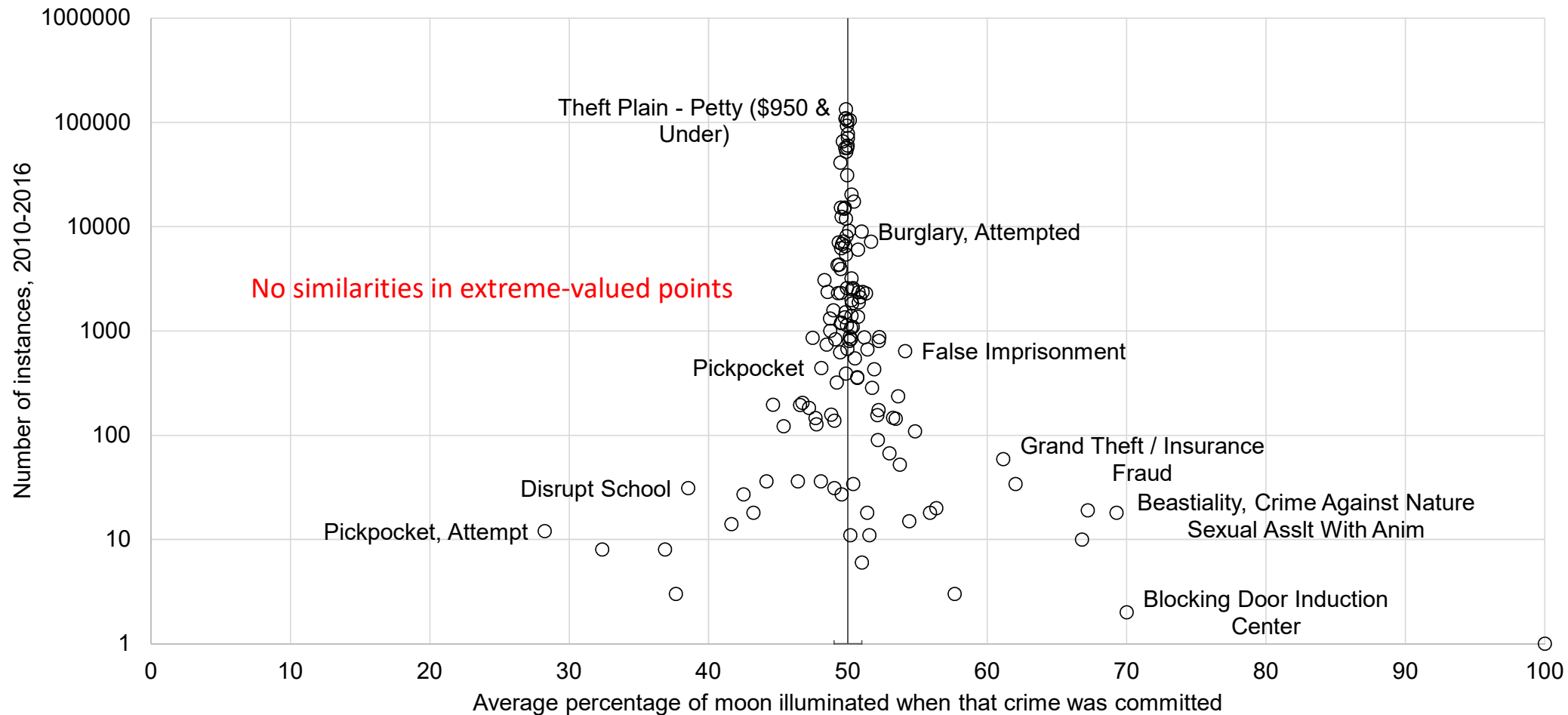
1. Are certain crimes (or crime types) more or less common at a given lunar phase?
2. Are certain crimes (or crime types) more likely to victimize one sex?

Example 1: No main effect of lunar phase on crime rate: no clear heterogeneity

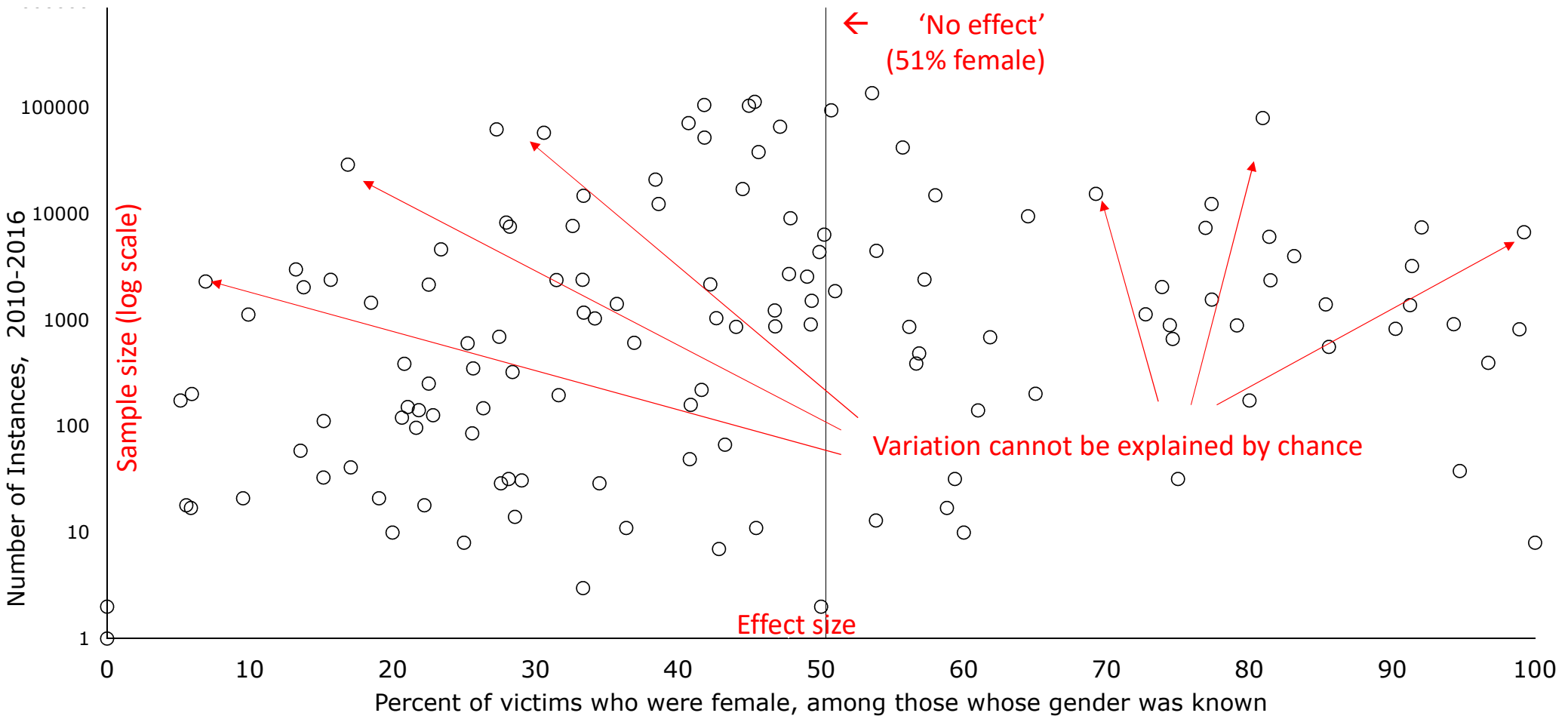
Lunar phase and crime incidence



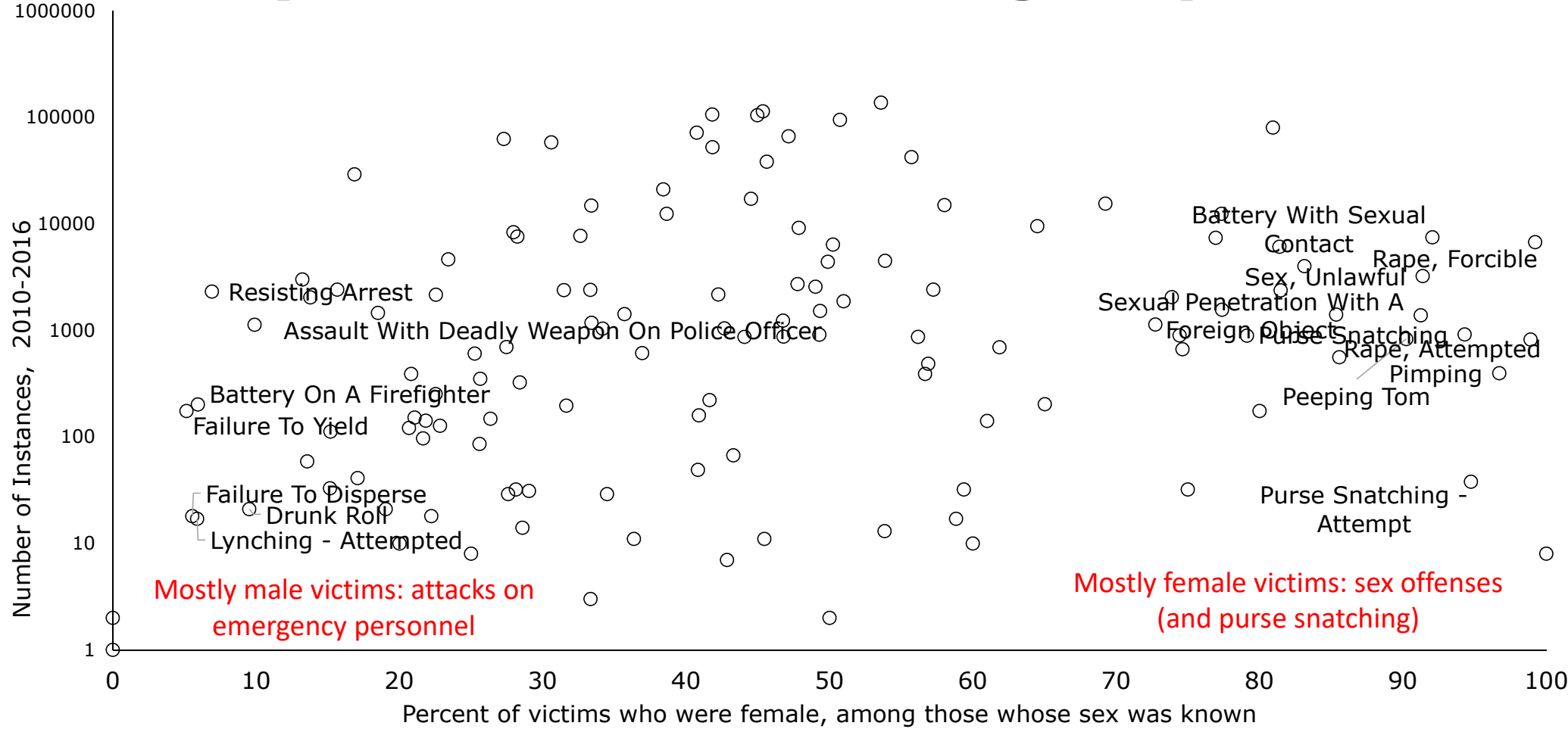
Example 1: Examination of extreme-valued points: no subgroup effects



Example 2: No main effect of sex on crime risk: however significant heterogeneity



Example 2: Examination of extreme-valued points: obvious subgroup effects



Conclusion: Association of rate of crime(s) with lunar phase or victim gender

→ Is there a main effect?

- Is crime more common at full or new moon? **Not really**
- Is one sex at greater risk of victimization? **Not really**

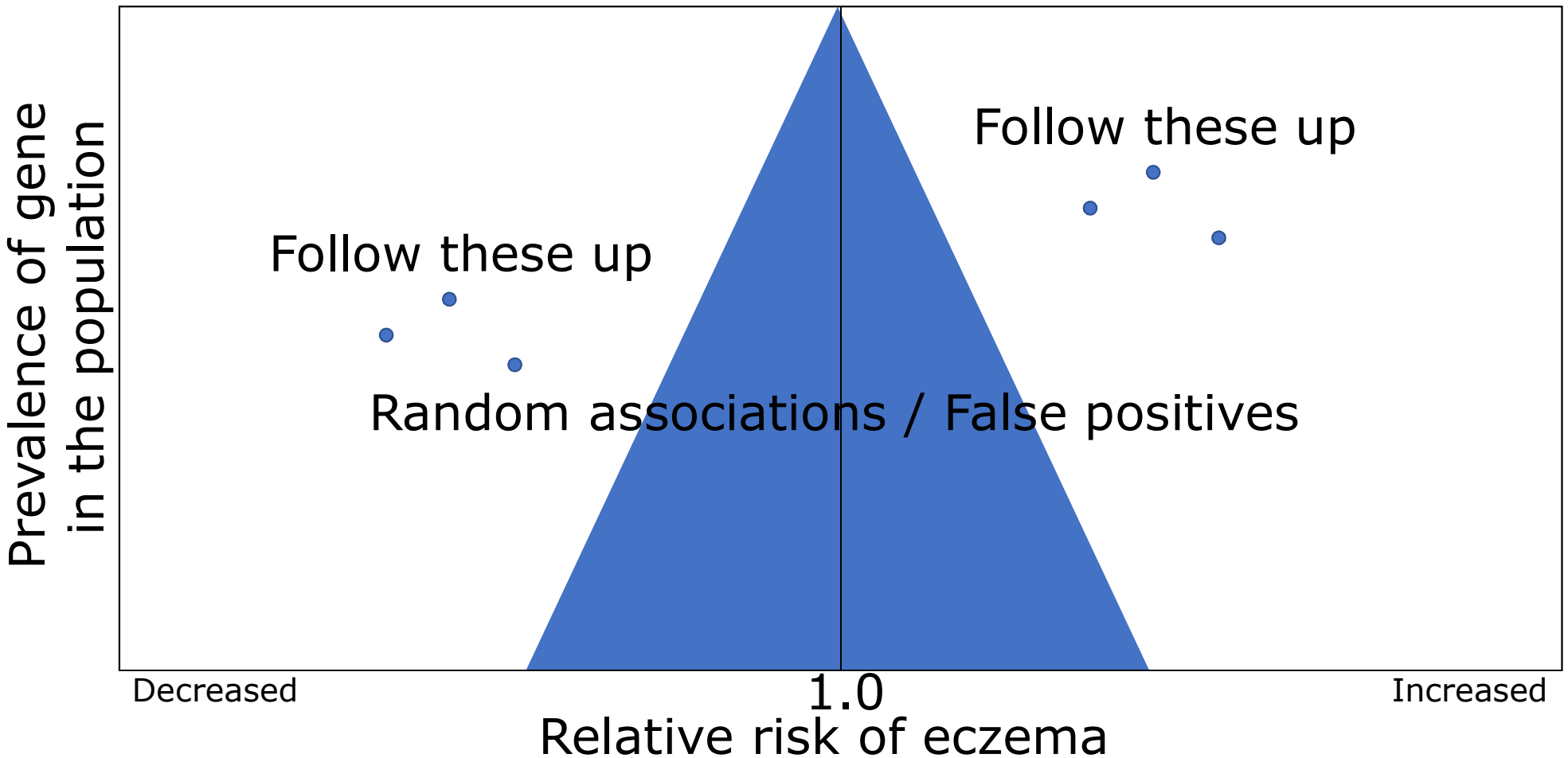
→ Is there effect heterogeneity?

- Are certain types of crime more common at a given lunar phase? **No**
- Are certain types of crime more likely to victimize one gender? **Yes**

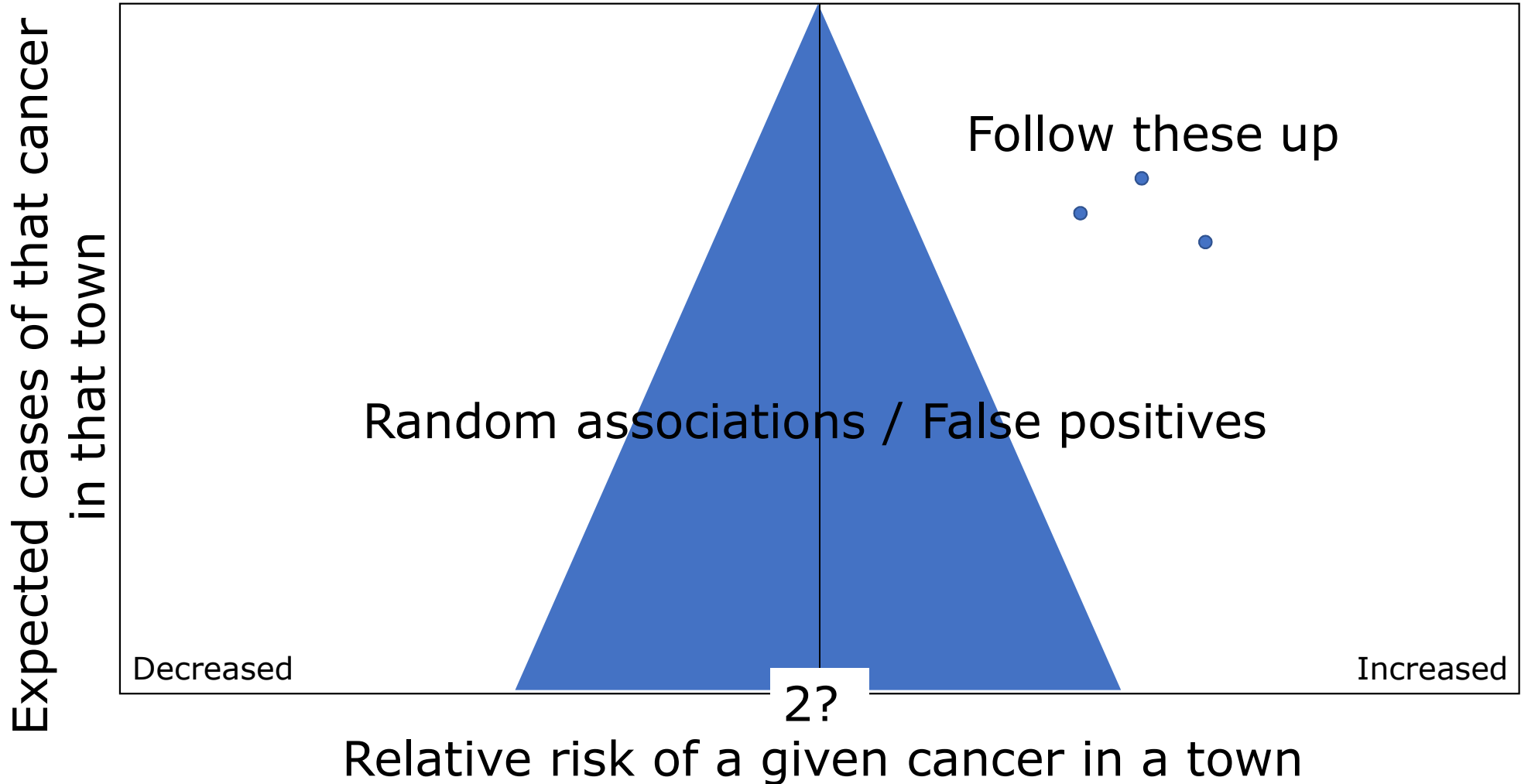
Conclusions:

- Effects of lunar phase on crime are **consistent with artifact** and can be ignored, **regardless of individual p-value**
- Effects of sex on crime victimization are **consistent with a true effect** and should not be ignored, **regardless of p-value**
 - Base rate of victimization does not vary much by sex: no main effect
 - Large within-outcome heterogeneity: 'crime' is not a single outcome
 - Studies of sex and crime should consider the clusterings we found (e.g. treat sex offenses as one cluster, and crimes against emergency personnel as another).

Application: Genome-wide association studies (GWAS)



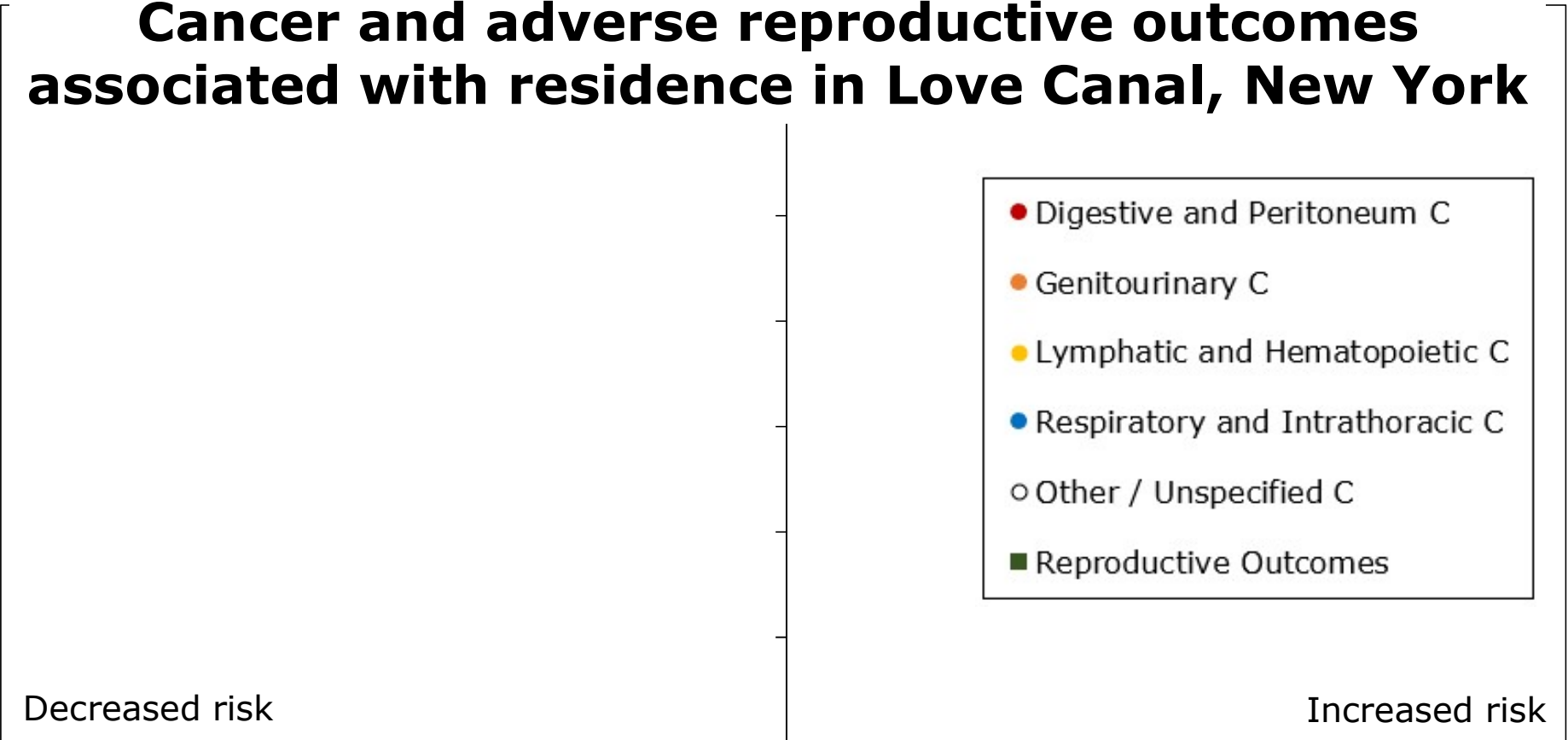
Further application: Environmental health



Final example: Environmental health

Cancer and adverse reproductive outcomes associated with residence in Love Canal, New York

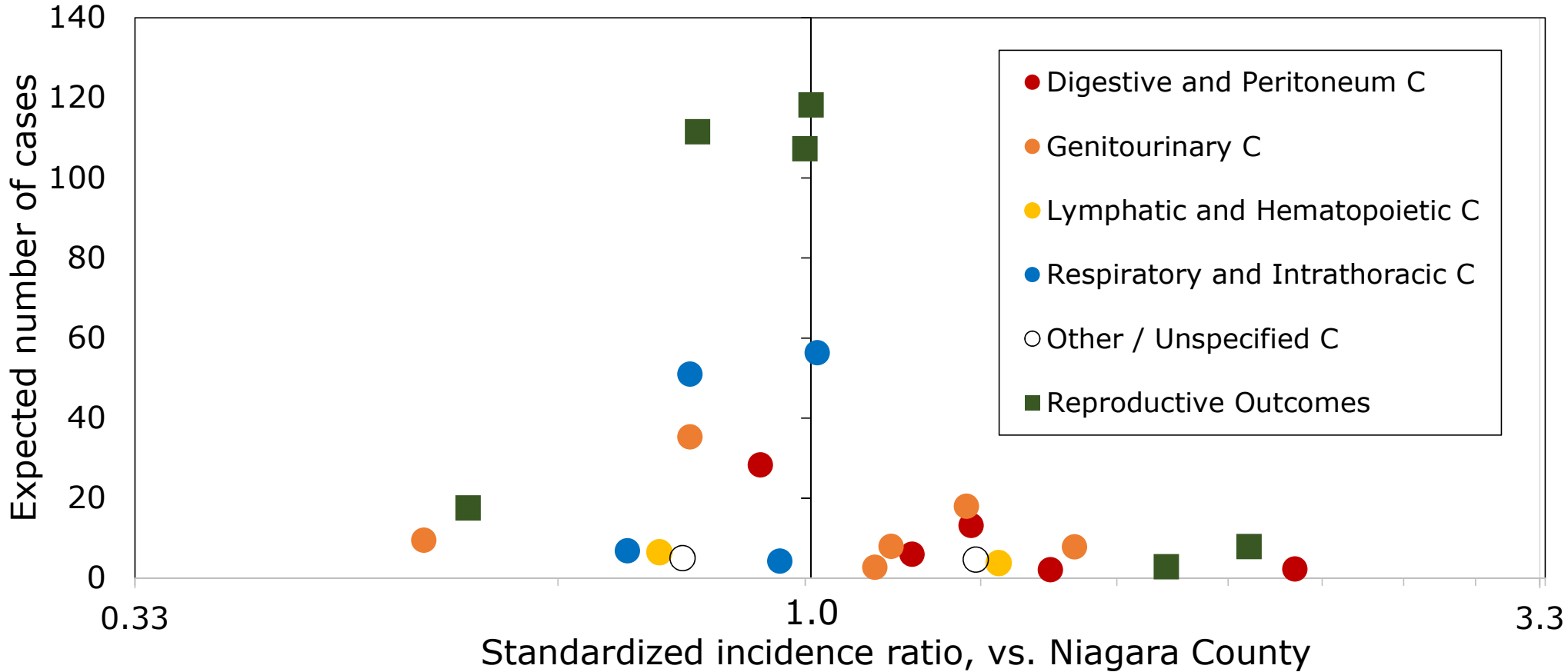
Expected number of cases based on population



Standardized incidence ratio, vs. a reference population

Final example: Environmental health

Cancer ('C') and adverse reproductive outcomes associated with residence in Love Canal, New York



Data from Tables 10 and 17 in Love Canal Follow-up Health Study, New York State Health Department; available at health.ny.gov/environmental/investigations/love_canal/docs/report_public_comment_final.pdf.

Thanks for listening!
Questions? Comments?