

# R- Package (*inctools*) and Shiny Apps for Incidence Estimation.

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## Background

### Incidence

- Determining disease incidence, the rate of occurrence of new cases in a population, presents a greater challenge than prevalence. For enduring conditions such as HIV, prevalence is not a good proxy for recent incidence because it depends in detail on historical incidence, demography, and survival.
- Reliable estimates of HIV incidence are critical for epidemiological monitoring, understanding transmission patterns, and in the design and evaluation of intervention programs.

### Incidence Estimation

We look at two primary alternative techniques to 'directly observed' incidence estimates from cohort studies:

1. Inferring incidence from cross-sectional surveys using biomarkers to test for 'recent infection' [1]. Kassanjee et al proposed a simple estimator for incidence

$$\hat{I}_T = \frac{P_R - \beta_T P_+}{P_S(\Omega_T - \beta_T T)}$$

where  $P_R$ ,  $P_+$  and  $P_S$  denote proportion of recent infections, proportion of infected individuals and proportion of susceptible individuals respectively.  $\Omega_T$  denotes the mean duration of recent infection within cutoff time T and  $\beta_T$  is the false-recent rate.

2. Inferring incidence from population renewal equations, given suitable age-stratified prevalence and mortality [2]. According to Mahiane et al, the incidence rate  $\lambda(a, t)$  is given by

$$\lambda(a, t) = \frac{1}{1 - P(a, t)} \left( \frac{\partial}{\partial t} P(a, t) + \frac{\partial}{\partial a} P(a, t) \right) + \Delta(a, t) P(a, t)$$

where  $P(a, t)$  and  $\Delta(a, t)$  represent the prevalence and excess mortality rate of infected individuals respectively.

## R- Package: *Inctools*

*Inctools* is a new R-package on CRAN that provides tools for estimating incidence based on Kassanjee et al, and maintained by SACEMA. Functionality can be categorized into three areas:

- **Support of survey design** using the two primary exposed functions *incprecision* and *incpower* which are essential for sample size and power calculations.
- **Analysis of survey data** consisting of subject level records indicating demographic factors and clinical indicators such as HIV and recency status. The primary inctool functions include *inccount*, *prevcount* and *incprop*.
- **Estimation of key performance characteristics of tests for recent infection** using the functions *frrcal* and *mdrical* to estimate false recent-rate and mean duration of recent infection respectively.

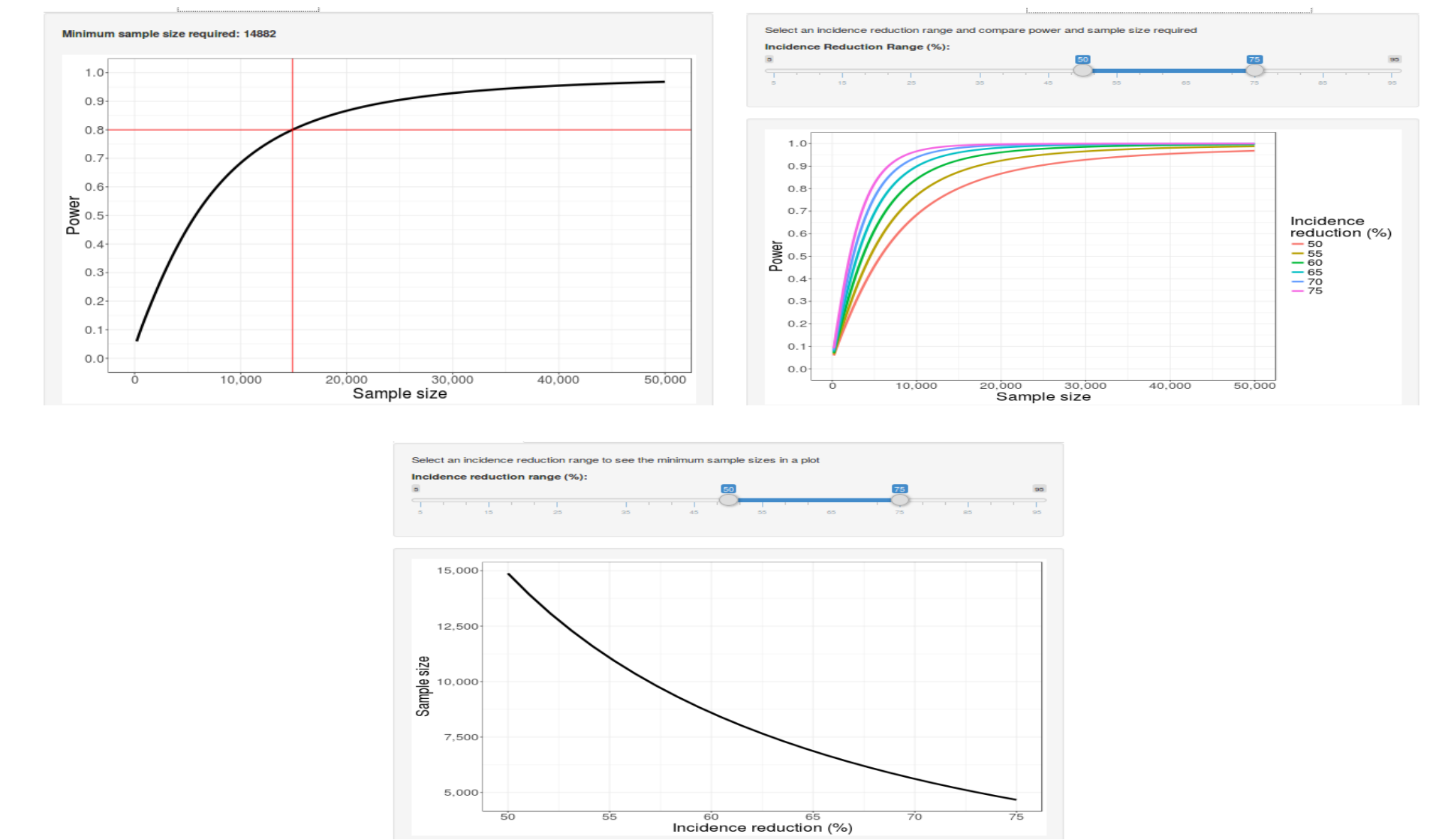
## Apps for incidence estimation tools:

E.g. Shiny App for **Sample size and power calculations** built based on inctools. Determines the required sample size for power (baseline survey and cohort).

The screenshot displays the Inctools Shiny App interface with several sections:

- Context:** Baseline incidence (cases/100PY) set to 1, Baseline prevalence (%) set to 10.
- Recency test:** Mean Duration of Recent Infection (days) set to 180, Relative Standard Error on MDRI (%) set to 10, False-Recent Rate (%) set to 0.5, Relative Standard Error on FRR (%) set to 25, Time cutoff T (days) set to 180.
- Baseline survey:** Recency test coverage rate (%) set to 100, Design effect on HIV prevalence set to 1, Design effect on prevalence of recency set to 1.
- Cohort:** Proportion of negatives recruited (%) set to 100, Follow-up time (years) set to 1, Design effect on cohort incidence set to 1.
- Significance:** Required power set to 0.8, Level of significance (α) set to 0.05.

The bottom of the interface features the SACEMA logo and the text 'DST-NRF Centre of Excellence in Epidemiological Modelling and Analysis'.



**Figure 1:** Output– Gives the minimum required sample size to achieve a specified power where respondents are recruited into the cohort to observe an expected decline in incidence. Also evaluates power vs sample size for different values of incidence reduction.

## Research Directions

- To extend and recode existing shiny applications using the *inctools* package based on Kassanjee et al.
- Implement key components of Mahiane et al [2] via:
  - A major R-package release for HIV Incidence estimation based on likelihood of individual level data and population renewal equations.
  - Vignettes to demonstrates the use of functions.
  - Shiny Apps for some key components and functionality.

## References

- [1] Reshma Kassanjee, Thomas A McWalter, Till Bärnighausen, and Alex Welte. A new general biomarker-based incidence estimator. *Epidemiology (Cambridge, Mass.)*, 23(5):721, 2012.
- [2] Guy Severin Mahiane, Rachid Ouifki, Hilmarie Brand, Wim Delva, and Alex Welte. A general hiv incidence inference scheme based on likelihood of individual level data and a population renewal equation. *PloS one*, 7(9):e44377, 2012.