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Evaluation of Ancestral Inbreeding Coefficients: *Ballou's* formula versus Gene Dropping

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Ancestral inbreeding coefficient (f_a)



$$f_a = [f_{a(s)} + (1 - f_{a(s)}) f_s + f_{a(d)} + (1 - f_{a(d)}) f_d] / 2$$
 Ballou's formula





Objectives

To determine the validity of Ballou's formula and propose a new method for an unbiased estimation of ancestral Inbreeding



Simulation model

- Population size: 50, 100, 200 with sex ratio 1:2
- Genome: 500 unlinked loci

genotype	relative viability (<i>w</i>)
AA	1
Aa	1- <i>h</i> s
aa	1- <i>s</i>

h is the dominance coefficient of *a*

s is the coefficient of selective against aa

Simulation model



- Genetic model: neutral; s = 0detrimental; s = 0.15h = 0lethal; s = 1
- Initial allele frequency: 0.005, 0.01
- The relative viability of individual is

 $W_i = W_1 W_2 W_3 \dots W_{500}$

Survival: w_i > random number (0- 1)

Simulation procedure







Ancestral inbreeding coefficients (f_a)

- *f_{a-true}* : true proportion of alleles of an individual
 that has undergone inbreeding in the past
- *f_{a-Ballou}* : *f_a* calculated by Ballou's formula
- f_{a-genedrop}: f_a estimated by gene dropping method





According to Ballou, f_a is the part of individual's genome that has undergone IBD in the past

$$f_{a} = \left[\left(f_{a(s)} \cup f_{s} \right) + \left(f_{a(d)} \cup f_{d} \right) \right] / 2$$

$$f_{a} = \left[f_{a(s)} + f_{s} - f_{a(s)} \cap f_{s} + f_{a(d)} + f_{d} \cap f_{a(d)} \cap f_{d} \right] / 2$$

$$\Pr\left(f_{a(s)} \cap f_{s} \right) = \Pr\left(f_{a(s)} | f_{s} \right) * \Pr\left(f_{s} \right)$$

$$= \Pr\left(f_{s} | f_{a(s)} \right) * \Pr\left(f_{a(s)} \right)$$
... dependent
$$= \Pr\left(f_{a(s)} \right) * \Pr\left(f_{s} \right) \dots \text{ independent}$$

 $f_a = \left[f_{a(s)} + f_s - f_{a(s)} f_s + f_{a(d)} + f_d - f_{a(d)} f_d \right] / 2 \quad \text{Ballou's formula}$











Conclusions

- Ballou's formula overestimated true ancestral inbreeding in all scenarios, because *fa* and *f* are not independent.
- Overestimation was not influenced by initial allele frequency and the genetic model.
- Overestimation occured earlier in smaller populations and reached higher values.



Conclusions

 The gene dropping method provided unbiased estimates for ancestral inbreeding in a neutral or a detrimental allele model but marginally biased estimates in a lethal allele model.



Thank you for your attention !!



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Background



Ancestral inbreeding coefficient (f_a)

$$f_a = [f_{a(s)} + (1 - f_{a(s)}) f_s + f_{a(d)} + (1 - f_{a(d)}) f_d] /2$$

Ballou's formula

$$f_a = [f_{a(s)} + f_s - f_{a(s)} f_s + f_{a(d)} + f_d - f_{a(d)} f_d] /2$$

- *f_a*: the cumulative proportion of an individual's genome that has been previously exposed to inbreeding in its ancestors
- f : inbreeding coefficient

