

	<p>LOUISIANA STATE UNIVERSITY College of Agriculture School of Plant, Environmental, and Soil Sciences AGRO 7075 Prediction-based Breeding</p>	
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Covariance between relatives

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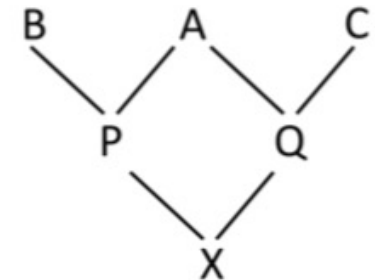
Baton Rouge, Feb 6th, 2023

Definitions

- **Coefficient of kinship (f)**
 - Probability that two gametes taken at random from two individuals are identical by descent (IBD, \equiv)
 - Expresses the degree of relatedness between individuals - **coefficient of parentage**
- **Coefficient of relationship (r)**
 - It is the additive genetic relationship between individuals
 - This is the twice the coefficient of kinship
 - $r = 2f$
 - It is also equal the inbreeding coefficient of their progeny
- **Additive covariance between relatives**
 - The covariance between the breeding values
 - $COV_{a(x,y)} = r_{xy} V_a$
 - It can actually be due to additive genetic effects, as well as dominance and epistatic effects
 - In general the contribution of dominance and epistatic effects to the genetic covariance is low

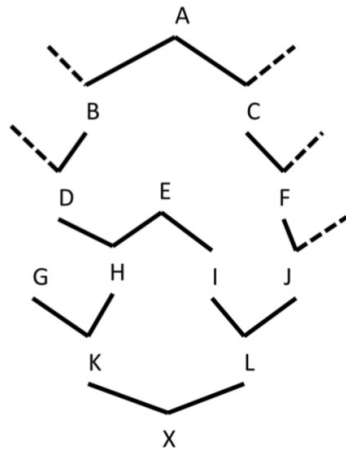
Calculating the inbreeding coefficient of X

- Let's consider one bi-allelic locus with two alleles A_1 and A_2
- We assume that A, the common ancestor of P and Q, is not inbred, thus its genotype is A_1A_2
- The probability that X receives A_1 from A via P, is the probability that A passes A_1 to P multiplied by the probability that P passes A_1 to X
- This probability is $1/2 \cdot 1/2 = 1/4$
- Now we need to know the probability that X receives A_1 from both P and Q
- $1/4 \cdot 1/4 = 1/16$
- **We now know the probability that A_1 is IBD in X**
- X could also be IBD by receiving two copies of A_2
- Probability IBD in X via either A_1 or A_2 is $1/16 + 1/16 = 2/16 = 1/8$
- $P(\text{IBD}) = 1/2^3 = 1/2^n$, where n is the number of common ancestral individuals
- **However, if the parent A is inbred, the IBD increases and should be considered**
- $(1/2)^n F_A$, where F_A is the inbreeding coefficient of the common ancestor
- **Thus, IBD is the sum the two probabilities:**
- $F_X = (1/2)^n + (1/2)^n F_A$
- $F_X = (1/2)^n (1 + F_A)$



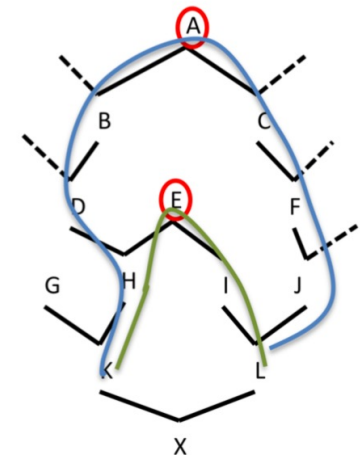
Calculating the inbreeding coefficient of X

- In more complex pedigrees, parents may be related to each other through more than one common ancestor, or from the same common ancestor, but through different paths
- The general formula is
- $F_X = (1/2)^n (1 + F_A)$
- where n is the number of individuals in any path of relationship counting the parents of X and all individuals in the path which connects the parents to the common ancestor
- The summation is over all paths



There are 2 common ancestors, A and E
There are 2 possible paths

Paths	n	F of common ancestor	Contribution of F_X
KHDB A CFJL	9	0	$(1/2)^9 = 0.002$
KH E IL	5	0	$(1/2)^5 = 0.031$
			Total= 0.033

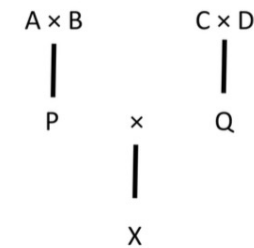


The coefficient of kinship

- Probability that two gametes taken at random (one from each individual carry alleles that are IBD
- The kinship (f) between two individuals is equal to the inbreeding coefficient of their progeny
- $F_X = f_{p_1 p_2}$
- where X is the progeny and p_1 and p_2 are the parents

- **Basic rules to estimate f**
- First: the f between P and Q is the mean of the four co-ancestries

$$f_{PQ} = \frac{1}{4}f_{AC} + \frac{1}{4}f_{AD} + \frac{1}{4}f_{BC} + \frac{1}{4}f_{BD}$$



- Second: the coefficient of kinship of an individual with itself f_{AA} is the inbreeding coefficient of progeny that would be produced by self-mating
- $f_{AA} = \frac{1}{2}(1 + F_A)$
- Third: the coefficient of kinship between parent and offspring f_{PA} is the mean coefficient of kinship between A and both the parents of P, (A and B)
- $f_{PO} = \frac{1}{2}(f_{AO} + f_{BO})$

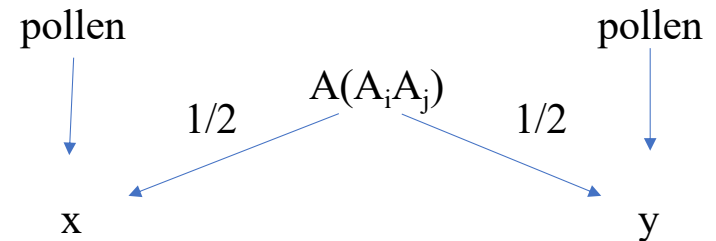
Covariance between relatives

- $f_{xy} = \frac{1}{4} [P(x_i \equiv y_i) + P(x_j \equiv y_j) + P(x_i \equiv y_j) + P(x_j \equiv y_i)]$
- $u_{xy} = [P(x_i \equiv y_i; x_j \equiv y_j) + P(x_i \equiv y_j; x_j \equiv y_i)] - \text{simultaneous events (the same genotype - dominance)}$

- **NON-INBRED RELATIVES**

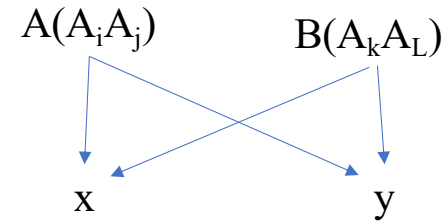
- **Half-sibs**

- $F = 0$, thus, $F_A = 0$
- $f_{xy} = \frac{1}{4} [P(x \equiv y \equiv A_i) + P(x \equiv y \equiv A_j) + P(x \equiv A_i \equiv y \equiv A_j) + P(x \equiv A_j \equiv y \equiv A_i)]$
- If x and y are non-inbred, thus the last two parts are zero, because their parents are not inbred either
- $f_{xy} = \frac{1}{4} [1/4 + 1/4 + 0 + 0]$
- $f_{xy} = 1/8$
- Since $r = 2f$
- $r = 1/4$
- $u_{xy} = 0$
- Probability of transmit the genotype – **dominance effect**



Covariance between relatives

- **Full-sibs**
- $F = 0$, thus, $F_A = F_B = 0$
- $f_{xy} = \frac{1}{4} [P(x \equiv y \equiv A_i) + P(x \equiv y \equiv A_j) + P(x \equiv y \equiv A_k) + P(x \equiv y \equiv A_L) +$
- $P(x \equiv A_i \equiv y \equiv A_j) + P(x \equiv A_j \equiv y \equiv A_i) + P(x \equiv A_k \equiv y \equiv A_L) + P(x \equiv A_L \equiv y \equiv A_k)]$
- Since A and B are non-inbred, $P(A_i \equiv A_j) = 0$
- $f_{xy} = \frac{1}{4} [1/4 + 1/4 + 1/4 + 1/4 + 0 + 0 + 0 + 0]$
- $f_{xy} = 1/4$
- Since $r = 2f$
- $r = 1/2$
- $(\frac{1}{2} \cdot \frac{1}{2}) \cdot (\frac{1}{2} \cdot \frac{1}{2})$
- $u_{xy} = [P(x \equiv y \equiv A_i; x \equiv y \equiv A_k) + P(x \equiv y \equiv A_i; x \equiv y \equiv A_L) +$
- $P(x \equiv y \equiv A_j; x \equiv y \equiv A_k) + P(x \equiv y \equiv A_j; x \equiv y \equiv A_L)]$
- $u_{xy} = [(0.25 \times 0.25) + (0.25 \times 0.25) + (0.25 \times 0.25) + (0.25 \times 0.25)]$
- $u_{xy} = 1/16 + 1/16 + 1/16 + 1/16$
- $u_{xy} = 1/4$



Covariance between relatives

- **INBRED RELATIVES**

- **Half-sibs**

- $F_p \neq 0; P(A_i \equiv A_j) = P(A_k \equiv A_L) \neq 0$

- $f_{xy} = \frac{1}{4} [P(x \equiv y \equiv A_i) + P(x \equiv y \equiv A_j) + P(x \equiv A_i \equiv y \equiv A_j) + P(x \equiv A_j \equiv y \equiv A_i)]$

- these cases can be $A_i \equiv A_j$

- $f_{xy} = \frac{1}{4} [1/4 + 1/4 + F(1/4) + F(1/4)]$

- $f_{xy} = \frac{1}{4} [1/2 + F(1/2)]$

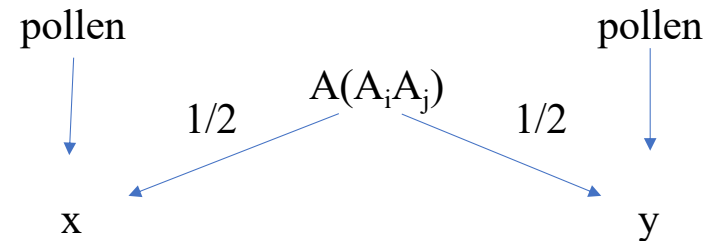
- $f_{xy} = 1/8[1 + F]$

- Since $r = 2f$

- $r = \frac{1}{4}[1+F]$

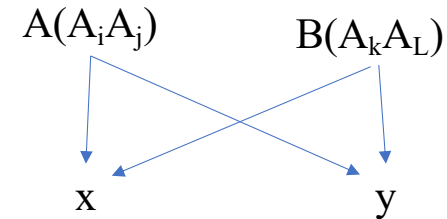
- $u_{xy} = 0$

- Probability of transmit the genotype – **dominance effect**



Covariance between relatives

- **Full-sibs**
- $F \neq 0$, thus, $F_A = F_B = 0$
- $f_{xy} = \frac{1}{4} [P(x \equiv y \equiv A_i) + P(x \equiv y \equiv A_j) + P(x \equiv y \equiv A_k) + P(x \equiv y \equiv A_L) +$
- $P(x \equiv A_i \equiv y \equiv A_j) + P(x \equiv A_j \equiv y \equiv A_i) + P(x \equiv A_k \equiv y \equiv A_L) + P(x \equiv A_L \equiv y \equiv A_k)]$
- Since A and B are non-inbred, $P(A_i \equiv A_j) = 0$
- $f_{xy} = \frac{1}{4} [1/4 + 1/4 + 1/4 + 1/4 + F(1/4) + F(1/4) + F(1/4) + F(1/4)]$
- $f_{xy} = \frac{1}{4}[1 + F]$
- Since $r = 2f$
- $r = \frac{1}{2}[1 + F]$



- $(\frac{1}{2} \cdot \frac{1}{2}) + (\frac{1}{2} \cdot \frac{1}{2})F$
- $u_{xy} = [P(x \equiv y \equiv A_i; x \equiv y \equiv A_k) + P(x \equiv y \equiv A_i; x \equiv y \equiv A_L) +$
- $P(x \equiv y \equiv A_j; x \equiv y \equiv A_k) + P(x \equiv y \equiv A_j; x \equiv y \equiv A_L)]$
- $u_{xy} = [(0.25 \times 0.25) + (0.25 \times 0.25) + (0.25 \times 0.25) + (0.25 \times 0.25)]$
- $u_{xy} = 1/16 (1 + F)^2 + 1/16 (1 + F)^2 + 1/16 (1 + F)^2 + 1/16 (1 + F)^2$
- $u_{xy} = \frac{1}{4}(1 + F)^2$

Covariance between relatives

- $P(x \equiv y \equiv A_i) = P(x \equiv y \equiv A_k) + P(x \equiv A_i \equiv y \equiv A_j)$
- $= \frac{1}{2} \cdot \frac{1}{2} + F(\frac{1}{2} \cdot \frac{1}{2})$
- $= \frac{1}{4} + \frac{1}{4}F$
- $= \frac{1}{4}(1 + F)$

- $P(x \equiv y \equiv A_k) = P(x \equiv y \equiv A_k) + P(x \equiv A_k \equiv y \equiv A_L)$
- $= \frac{1}{2} \cdot \frac{1}{2} + F(\frac{1}{2} \cdot \frac{1}{2})$
- $= \frac{1}{4} + \frac{1}{4}F$
- $= \frac{1}{4}(1 + F)$

- $P(x \equiv y \equiv A_i; x \equiv y \equiv A_k) = P(x \equiv y \equiv A_i) \cdot P(x \equiv y \equiv A_k)$
- $= \frac{1}{4}(1+F) \cdot \frac{1}{4}(1+F)$
- $= \frac{1}{16}(1+F)^2$

Why $r = 2f$?

- $x_{ij} = u + \alpha_{ix} + \alpha_{jx} + \mathcal{S}_{ijx}$
- $y_{ij} = u + \alpha_{iy} + \alpha_{jy} + \mathcal{S}_{ijy}$
- $E(x_{ij}) = E(y_{ij}) = u$
- $E(\alpha_i) = \sum_i p_i \alpha_i = 0$
- $E(\alpha_j) = \sum_j p_j \alpha_j = 0$
- $E(\mathcal{S}_{ij}) = \sum_{ij} p_i p_j \mathcal{S}_{ij} = 0$
- $E(\alpha_i, \alpha_j) = E(\alpha_i) E(\alpha_j) = 0$
- $E(\alpha_i, \mathcal{S}_{ij}) = E(\alpha_i) E(\mathcal{S}_{ij}) = 0$
- $E(\alpha_j, \mathcal{S}_{ij}) = E(\alpha_j) E(\mathcal{S}_{ij}) = 0$
- $\text{COV}(x_{ij}, y_{ij}) = E[x_{ij} - E(x_{ij})] \cdot E[y_{ij} - E(y_{ij})]$
- $= [u + \alpha_{ix} + \alpha_{jx} + \mathcal{S}_{ijx} - u] \cdot [u + \alpha_{iy} + \alpha_{jy} + \mathcal{S}_{ijy} - u]$
- $= E(\alpha_{ix}, \alpha_{iy}) + E(\alpha_{ix}, \alpha_{jy}) + E(\alpha_{ix}, \mathcal{S}_{ijy}) + E(\alpha_{jx}, \alpha_{iy}) + E(\alpha_{jx}, \alpha_{jy}) + \dots + E(\mathcal{S}_{ijx}, \mathcal{S}_{ijy})$
- $E(\alpha_{ix}, \mathcal{S}_{ijy}) = 0$
- Covariance between allele effect and genotype (all cases)

Why $r = 2f$?

- $E(\alpha_{ix}, \alpha_{iy}) = \sum_i p_i \alpha_i P(x_i \equiv y_i) \alpha_i = \sum_i p_i \alpha_i^2 P(x_i \equiv y_i) = 1/2 Va. P(x_i \equiv y_i)$
- $E(\alpha_{ix}, \alpha_{jy}) = \sum_i p_i \alpha_i P(x_i \equiv y_j) \alpha_i = \sum_i p_i \alpha_i^2 P(x_i \equiv y_j) = 1/2 Va. P(x_i \equiv y_j)$
- $E(\alpha_{jx}, \alpha_{iy}) = \sum_j p_j \alpha_j P(x_j \equiv y_i) \alpha_j = \sum_j p_j \alpha_j^2 P(x_j \equiv y_i) = 1/2 Va. P(x_j \equiv y_i)$
- $E(\alpha_{jx}, \alpha_{jy}) = \sum_j p_j \alpha_j P(x_j \equiv y_i) \alpha_j = \sum_j p_j \alpha_j^2 P(x_j \equiv y_i) = 1/2 Va. P(x_j \equiv y_i)$
- $E(\mathcal{S}_{ijx}, \mathcal{S}_{ijy}) = \sum_{ij} p_i p_j \mathcal{S}_{ij} [P(x_j \equiv y_i, x_j \equiv y_j) + [P(x_i \equiv y_j, x_j \equiv y_i)] \mathcal{S}_{ij}$
- $= \sum_{ij} p_i p_j \mathcal{S}_{ij}^2 [P(x_j \equiv y_i, x_j \equiv y_j) + [P(x_i \equiv y_j, x_j \equiv y_i)]$
- $= Vd. [P(x_j \equiv y_i, x_j \equiv y_j) + [P(x_i \equiv y_j, x_j \equiv y_i)]$
- $COV(x_{ij}, y_{ij}) = E[x_{ij} - E(x_{ij})] \cdot E[y_{ij} - E(y_{ij})]$
- $= [u + \alpha_{ix} + \alpha_{jx} + \mathcal{S}_{ijx} - u] \cdot [u + \alpha_{iy} + \alpha_{jy} + \mathcal{S}_{ijy} - u]$
- $= E(\alpha_{ix}, \alpha_{iy}) + E(\alpha_{ix}, \alpha_{jy}) + E(\alpha_{ix}, \mathcal{S}_{ijy}) + E(\alpha_{jx}, \alpha_{iy}) + E(\alpha_{jx}, \alpha_{jy}) + \dots + E(\mathcal{S}_{ijx}, \mathcal{S}_{ijy})$
- $COV_a(x_{ij}, y_{ij}) = 1/2 Va [P(x_j \equiv y_i) + P(x_j \equiv y_j) + P(x_i \equiv y_j) + P(x_i \equiv y_i)] = 1/2 Va [4f_{xy}] = 2fVa$
- $COV_d(x_{ij}, y_{ij}) = 1/2 Vd. [P(x_j \equiv y_i, x_j \equiv y_j) + [P(x_i \equiv y_j, x_j \equiv y_i)] = Vd[u_{xy}] = u_{xy} Vd$
- $COV_g(x_{ij}, y_{ij}) = 2.f_{xy}.Va + u_{xy} Vd$