

Chance, often an Ignored Factor in Genetic Studies

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Genetic studies have made a substantial progress in identifying the genetic basis of many human diseases. However, search for genes or loci that might underlie a disorder among family members have sometimes been disappointing, especially if the disorder is common.

One reason for that could be not considering chance association among family members affected by the disorder. For example among 158 868 females with breast cancer, 5088 first-degree relatives could be identified¹. Noteworthy, in the matched (by year of birth, sex and year of death) control cohort with the same number of healthy females, no less than 3326 first-degree relatives were found (65%) and thus representing the normal hereditary component of such a big cohort.

The impact of chance association in searching for genes in first-degree relatives for some other common forms of cancer can be estimated from that study [1].

Most affected cancer by chance:

Stomach	74%
Breast	65%
Urinary organs	57%

Colon 53%

Lung 51%

Least affected cancer by chance:

Eye 6%

Testis 11%

Hodgkin's disease 15%

Thyroid gland 16%

Thus, when performing genetic studies on family members with a certain form of cancer, one has to be aware of the chance association. Even though a mother and her daughter both have breast cancer there is only 35% of such pairs that are true genetic and this must be considered when interpreting the result.

References

1. Lindelöf B, Eklund G. (2001) Analysis of hereditary component of cancer by use of a familial index by site. *Lancet* 358: 1696-98.

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