

EPITHEORESE KLINIKES FARMAKOLOGIAS
 KAI FARMAKOKINETIKES, INTERNATIONAL
 EDITION 16: 77 (2002)
 PHARMAKON-Press

Reconsidering the Toxicology of GM Food

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If we go a few years back, we can see what WHO was saying in terms of the first regulatory action on the problem of GM food: "Considerations that should be made on the biosafety of GM food are practically the same with natural food" (substantially equivalence dogma). This is a misleading and erroneous stance to take, because technically speaking, production of animals or plants by taking advantage of biotechnology has no resemblance to the production by traditional breeding methods.

Organization of genetic material in plants or animals is extremely complicated. Every time that nature tries to create new varieties it preserves the natural hierarchy of the genome. In GM organisms that are used in GM food, the genetic modification employs artificial units of DNA that are introduced in animal or plant cells, which always disrupt the natural organization of the host genome. The consequences of this phenomenon known as position effect have been well described (Palmiter and Brinker: *Ann. Rev. Genet.* 20: 645-660, 1986). This effect has been taken as minor problem by the scientists which were first experimented with GM food, but someone can assume that considering the knowledge at that time, this misleading stance was taken in purpose.

Once exogenous DNA is injected into reproductive cells of the organism, the introduced gene is randomly incorporated into the host genome, resulting in disruption of the order of the host genome. Therefore the genetic modification in an organism, actually results in a loss, of the balanced functioning that is preserved through traditional breeding.

As a result, a new plant or animal is produced, that is not able to synthesize proteins or substances that are not made or made in a different quantity by the parent organism. These proteins can be allergenic or toxic as shown in the literature. The same is also true with the other substances of the GM foods. The latter is well pronounced in the example of the nutritional tragedy in USA in 1989, when many people died from the eosinophilia-myalgia syndrome. One

company that was making tryptophan from genetically engineered bacteria, changed the manufacturing conditions during the process. The result was a by-product of the fermentation that was finally consumed by many people, and created the problem. The absence of testing of GM foods at that time has been changed over the years, but in a way that does not exclude the risk to the consumer.

Another example has been published by Reddy and Thomas (*Nature Biotechnology* 14: 639-642, 1996). During their effort to create GM tobacco to produce gamma-linoleic acid, the GM tobacco produced also a highly toxic fatty acid that was not expected. There are other numerous examples that describe the production of substances in GM food that were not expected and all these events can be explained by the position effect phenomenon.

We see a problem arising from all above. The problem of incomplete testing of GM food. Since almost all GM organisms as explained earlier may pose risks to contain new products that are not known if are toxic to the consumers, all food that is produced by GM organisms must be tested pre-clinically and clinically with human volunteers in trials similar to what used to assess drugs and pharmaceuticals before marketing (a product with no toxicity in animals does not exclude the possibility being toxic to humans, remember thalidomide). Today the tests that are made by biotech companies are incomplete and require few weeks and are made in cell lines and mice for a period that does not make them biosafe. The tests should include processed food that has been derived from GM organisms (oil from soya) or food that contains ingredients from GM organisms (like enzymes from GM yeast used in the fermentation of cheese). This is a time consuming process but is mandatory to happen in order to minimize health hazards to the consumers.

As the genetic modification in the production of GM food may pose unexpected health risks for us and our children, new directives should be made to exclude acute or chronic toxicity, if we want the tragedy of tryptophan in 1989 not to happen again.