Trans Fat and Cardiovascular Disease
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Abstract
Cardiovascular disease remains the number one killer in the United States. According to the American Heart Association statistics, CVD accounted for 35.2% of total deaths in 2005 in the United States. This is also true for developing countries, such as China and India: CVD has also become the first or second leading causes of death; it is responsible for around 30% of total deaths in developing countries. Of many risk factors for CVD, dietary intake of fats received much attention and has been extensively studied due to the fact that blood cholesterol levels are a strong predictor for CVD and that dietary fats are a strong determinant for serum lipids. Although the effects of some fatty acids on coronary health are still a matter of debate, the relationship between trans fatty acids and risk of CVD has been established. [N AJ Med Sci. 2008;1(1):34-35.]

Biochemistry of trans fatty acids
Fatty acids are categorized into different sub-groups according to the chain length of carbon-backbone and presence of unsaturated double bonds. Saturated fatty acids have a carbon-backbone of 4-24 carbons and no unsaturated double bonds. In contrast, unsaturated fatty acids have a carbon-backbone of 16-22 carbons and at least one unsaturated double bond. Trans fatty acids are a special kind of unsaturated fatty acid in that the hydrogen atoms are on opposite (trans configuration) sides of the double bonds, whereas for the rest of unsaturated fatty acids these hydrogen atoms are on the same (cis configuration) sides of the double bonds. It is because of this trans configuration, trans fat is similar to saturated fat in terms of physical properties and is semi-solid at room temperature. More importantly, it is because of this subtle structural difference, trans fatty acids are entirely different from other unsaturated fatty acids with respect to coronary effects.

History
Trans fatty acids are naturally produced in the rumen through bacterial fermentation; about 5% of ruminant fat is trans fat. However, most trans fats in the US diet are produced by the food industry via partial hydrogenation vegetable oils. Nobel Prize winner Paul Sabatier developed the methods of hydrogenation in 1890s which enables the production of trans fats. Hydrogenation of oils is a process that adds hydrogen atoms to the unsaturated double bonds and thus unsaturated fatty acids are converted to saturated fatty acids. In partial hydrogenation, only a proportion of the double bonds are fully saturated and some double bonds are converted from cis to trans configuration. Partially hydrogenated oils first entered US food supply by 1920. But it is until 1950s-1960s the use of trans fats (primarily margarine) became substantial. Ironically, at that time, margarine was regarded as a healthy substitute of butter because of low saturated fat content. Total consumption of trans fat was approximately 2-3% of total energy in the US.

Physiological Effects of Trans Fatty Acids
The evidence regarding adverse effects of trans fatty acids on blood lipids and other CVD risk factors is substantial. Of many potential mechanisms through which trans fatty acids may increase the risk of CVD, the effect of trans fatty acids on blood lipid profiles has been most extensively studied in clinical trials. Mensink et al. summarized results from 8 clinical trials that specifically examined the effect of 18:1 trans isomer intake in a meta-analysis. Results indicated that of all classes of fatty acids, trans fatty acids had a strong effect on raising serum LDL cholesterol (bad cholesterol) concentrations and were the only class of fatty acids that did not raise HDL cholesterol (good cholesterol) concentrations when replacing carbohydrates. As a result, trans fatty acids had the strongest effect on raising total cholesterol to HDL ratio, which is a stronger predictor of CHD risk than total cholesterol. Trans fatty acids may increase the risk of CVD through other pathways, including increasing lipoprotein(a) and blood triacylglycerol concentrations, and interfering with essential fatty acid metabolism and eicosanoid balance by inhibiting delta-6-desaturase. Furthermore, trans fat intake has been shown to be associated with systemic inflammation, endothelial cell dysfunction, and insulin resistance. By their incorporation into the phospholipids in cell membranes, trans fatty acids may also alter the membrane function and decrease the membrane fluidity in a way similar to saturated fat.

Epidemiologic Evidence
Because of the clear adverse effects of trans fatty acids on blood lipids and inflammatory markers, it is unethical to conduct clinical trials that use the incidence of CVD as the outcome. Evidence from several well-designed prospective studies consistently showed a strong association between trans fatty acid intake and the risk of CVD. In a meta-analysis, 2% of total energy from trans fat was associated with a 23% increase of CVD risk. It is worth noticing that 2% of total energy is a small unit. These results were
corroborated by several case-control studies that used trans fatty acid contents in human tissues as an indicator of trans fat intake. For example, in the Nurses’ Health Study, women whose trans fatty acid levels in red blood cells were in the highest quartile had more than 3 times elevated risk of developing coronary heart disease than women in the lowest quartile.\textsuperscript{15} Data from developing countries were rare. In a Costa Rican population, trans fatty acids in adipose tissue were also associated with an increased risk of CVD.\textsuperscript{16}

**Food Sources and Customers’ Choices**

On January 1, 2006, US Food and Drug Administration (FDA) ruled that nutrition labels for all foods and supplements must indicate the content of trans fatty acids. At the end of 2006, New York City passed a trans fat ban at restaurants. Currently, more and more cities have passed or are considering to pass such a ban in the US and worldwide. This is probably the first time that, without clinical trial evidence, a nutrient is being eliminated from the food supply in the US. Traditionally, primary food sources of trans fatty acids are fast foods and bakery products, including French fries, breaded fish burgers, breaded chicken nuggets, pies, sweet rolls, doughnuts, cookies, cakes, brownies, and muffins. This is not at all a complete list. Consumers are recommended to read the food labels. It is necessary to point out those foods containing less than 500 mg trans fatty acids per serving are allowed to be labeled as zero trans fat content. Therefore, several servings of such foods can still provide substantial trans fatty acids.

**Conclusions and Future Implications**

Based on the evidence from basic research, clinical trials, and epidemiologic studies, it is safe to conclude that trans fatty acid intake bears no benefit to human health and can bring substantially harmful effects to overall coronary health. In developed countries, actions have already been taken to eliminate trans fats from the food supply. In contrast, data regarding trans fat content in food is lacking in third world countries. In China, unpublished data showed that among a small middle-aged population, trans fatty acid content in plasma was 20% of that of US population. However, given the dramatic lifestyle and nutrition transition that is currently undergoing in China, whether younger Chinese populations also have low trans fat content in their bodies is questionable. It is important to shed more light on this issue. Governments of developing countries should seriously not hesitate to ban trans fat use for the sake of their people’s health.

\textbf{References}