Can human micronutrient status be improved by supplementing domestic animals?

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Micronutrient deficiencies are a major problem throughout the world and hundreds of millions of the world's population are affected by micronutrient deficiency disorders. In Europe the prevalence of clinical micronutrient deficiency disorders is less than that in the Third World. However, marginal deficiency of some of the micronutrients might be involved in the aetiology of many of the so-called lifestyle diseases, e.g. cancer, cardiovascular diseases, diabetes, osteoporosis. Supplementing domestic animals with micronutrients in excess of their needs could be one strategy to increase the intake and, thereby, status of micronutrients in the human population. This approach should, however, be considered carefully, in relation to both animal and human welfare. Many micronutrients that may accumulate in animal foods are toxic in high doses. It would also be unethical to expose animals to doses that might have deleterious effects on their health, and concentrations in animal products that might have adverse effects when consumed by man should be avoided. Furthermore, food quality should not be impaired by the supplement. On the other hand, to be relevant in relation to human nutrition, the given micronutrient should accumulate in animal tissue in concentrations that make an important contribution to total intake. Finally, the micronutrient should be incorporated in a way and in a form that is bioavailable to man, i.e. is well absorbed and utilized.

Micronutrients: Animal food: Man: Micronutrient status

Prevalence of micronutrient deficiencies

Millions of the world's population are micronutrient deficient or are at risk of becoming deficient. In Europe alone 140 million of the population are assumed to be at risk of I-deficiency disorders and approximately 100 million Europeans currently have goitres (World Health Organization, 1996). Fe deficiency and Fe-deficiency anaemia are widespread among women of childbearing age worldwide, including Europe (World Health Organization, 1996). In Denmark it has been estimated that approximately 30 % of the women of childbearing age have low Fe stores (Milman & Kirchhoff, 1996). Vitamin D deficiency may be a problem among the elderly who are confined to their homes and in Muslim women living in Europe (World Health Organization, 1996). Low or marginal intakes of some micronutrients might also be involved in lifestyle diseases, such as cancer, cardiovascular diseases, osteoporosis, diabetes, etc. (Richard & Roussel, 1999). The optimal intake required to reduce the risk for these conditions is not known, but some European countries have relatively low intakes of folate, Se and Cu, and it has been suggested that higher intakes might be beneficial for human health (Van Dokkum, 1995).

Strategies to increase intake of micronutrients

Changing the dietary habits of a population is difficult and may take years. An alternative or complementary approach would be to increase the concentration of nutrients in foods that are already consumed. This approach could be implemented by the use of micronutrient-enriched fertilisers for crops, by food enrichment or by supplementing domestic animals with micronutrients. Meat, milk and eggs are good sources of protein, readily-available Fe, Ca, Mg, Se, Zn, vitamins A and D and a range of B-vitamins (Verbeke *et al.* 1999). In Denmark meat provides approximately 30 % of the Zn, Cu, Se and niacin intake (Andersen *et al.* 1996), approximately 25 % of the Fe

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400 S. Bügel

and vitamin B_6 intake and > 50 % of the vitamin B_{12} intake. Milk supplies approximately 15 % of the intake of retinol, vitamin D, Mg, Zn, Se and some of the B vitamins, and 25 % of the I intake. Eggs also make a small contribution to the intake of the same micronutrients as milk (Andersen et al. 1996). These levels are probably very similar in other countries where the national diet is based on animal protein. An increase in the total content of micronutrients in animal products could therefore be an effective way of improving human micronutrient status.

Animal welfare and food quality

Animal health and welfare, as well as meat quality, are important issues when considering supplementation in animals. The micronutrients most likely to be accumulated in animal tissue (e.g. Fe, Zn, Se and vitamins A and D) are also the nutrients that are most toxic in high doses. Supplementing animals with doses that are toxic for them is unethical and should be discouraged. In many countries with low soil Se concentrations animals are sometimes supplemented with Se injections. However, many animals have died because of accidental administration of toxic doses resulting from errors in dilution (Shortridge *et al.* 1971).

Supplementation should not affect food quality, e.g. colour, oxidative stability, taste, texture etc. of meat, which determines whether a given population will buy the food and eat it. Many studies have been conducted to improve meat quality, and oxidative stability, particularly, is of great concern to the food industry. Vitamin E supplementation has been shown to improve the oxidative stability of meat from a number of different animals, including chicken, veal and pork (O'Neill et al. 1998; Corino et al. 1999; Dufrasne et al. 2000; Granit et al. 2001; Guo et al. 2001). The addition of Fe or Cu, on the other hand, could have deleterious effects on meat quality, as they accelerate auto-oxidation (Snyder & Skrydlant, 1966), the photooxidation of oxymyoglobin (Assef et al. 1971), the oxidation of reducing agents (Kanner et al. 1988) and the propagation phase in lipid peroxidation (Harel & Kanner, 1985). Consequently, free Fe and Cu either directly or indirectly stimulate discoloration of meat, which is considered by customers to be a sign of unwholesomeness and product deterioration (Faustman & Cassens, 1990).

Efficacy of animal micronutrient supplementation

In order to be of any benefit, the micronutrient added to animal feed should accumulate in tissues that are normally consumed by man on a regular basis. Potential candidates are the fat-soluble vitamins A, D, E and K, some of the B-vitamins and some of the trace elements, such as Se, Fe, Cu and Zn. The fat-soluble vitamins are mainly deposited in fat and liver, while trace elements are deposited both in muscle and liver, as well as in other tissues. Mattila *et al.* (1992) found remarkable variation in the cholecalciferol content of eggs produced by hens receiving different types of feed. Subsequently, it was shown by feeding hens different amounts of cholecalciferol (26·6, 62·4 and 216·0 µg/kg feed) that there is a strong positive correlation

between the cholecalciferol content of the feed and the cholecalciferol $(r\,0.995)$ and 25-hydroxycholecalciferol $(r\,0.941)$ contents of the egg yolk (Mattila *et al.* 1999).

Vitamin E easily accumulates in meat (for review, see Liu *et al.* 1995) primarily in muscle, liver and adipose tissue (Machlin, 1994) and, in general, the greater the amount of vitamin E fed and/or the longer the supplementation, the higher tissue concentration of α -tocopherol (Arnold *et al.* 1993). Furthermore, it has been suggested that the L- α -tocopherol concentration of beef can be increased from $1.4 \,\mu\text{g/g}$ to the critical concentration of $3.5 \,\mu\text{g/g}$ by feeding a saturation dose of at least $1300 \,\text{mg/d}$ for 44 d (Arnold *et al.* 1993).

It is generally accepted that Se supplementation results in increased Se tissue concentrations, and many studies have shown correlations between dietary Se content and Se concentrations in animal products (for review, see Beale et al. 1990). Ruminants, however, absorb less of an oral dose than non-ruminants, probably because rumen flora are capable of converting various forms of Se into elemental Se (Pehrson et al. 1999). In single-stomached animals the absorption of Se depends on many variables, including the current Se status, the form of Se and the complexity of the diet (Levander, 1983).

Meat from Fe-supplemented animals may provide an alternative to existing methods of Fe fortification of food, as Fe from meat is highly available (Hurrell, 1997). Some studies have shown that increasing dietary Fe levels increases the concentration of Fe in muscle tissue in pigs in a dose-dependent manner (Miller *et al.* 1994).

The addition of Cu to animal feed has given conflicting results in relation to the accumulation of Cu in animal tissue. In studies in which pigs were fed acorns (with a high content of Cu; Rey & Lopez-Bote, 2001) or a Cu supplement (Lauridsen *et al.* 1999) no differences were found in muscle Cu concentrations. In contrast, increasing the amounts of Cu (from 10 mg CuSO₄/kg to 30 mg CuSO₄/kg) in the feed offered to ewes increased the liver Cu concentration of the slaughtered ewes (Eckert *et al.* 1999). Adding 250 mg Cu/kg to feed offered to chickens increased the Cu concentration of breast muscle by 14·5% (Bakalli *et al.* 1995). The concentrations were, however, relatively low (<0·5 mg/kg). This finding suggests that the accumulation of Cu in animals might depend on many factors, including animal species.

Bioavailability in man

Studies in which animal feed has been supplemented with micronutrients with the main purpose of increasing human micronutrient status are scarce. However, a number of studies have shown that many of the micronutrients present in meat are highly available, including Se (Shi & Spallholz, 1994*a*,*b*; Wen *et al.* 1997), Zn (Zheng *et al.* 1993; Sandström, 1997) and Fe (Hurrell, 1997).

Bauch *et al.* (1990) have demonstrated the beneficial effect of I supplementation. Addition of I to animal feed was initiated in 1985; almost 1 year after iodized salt was introduced onto the market in Germany. During the first year, in which only iodized salt was available, urinary I excretion (a marker of I intake) increased only marginally. Introduction of I to animal feed increased urinary excretion

markedly. During a 6-year period, the number of the population with I deficiency decreased from >90% to approximately 30% (Bauch *et al.* 1990).

Addition of vitamin E to pig feed has been shown to substantially increase the vitamin E content of both fat and organ meat in pigs (Morrissey *et al.* 1996). In one of our studies, however, the addition of vitamin E to pig feed only marginally increased the vitamin E content of experimental human diets containing pig products (Sandström *et al.* 2000).

In conclusion, supplementing animals with I might have a beneficial effect on the I status of the human population and is believed to be an effective means of eliminating most I-deficiency disorders in Europe (World Health Organization, 1996). Vitamin D supplementation of hens in order to produce enriched eggs might be a promising way of decreasing the low vitamin D status in some population groups during wintertime. The addition of Se to animal feed may be a means of improving human Se status, but its effectiveness will depend on the chemical form of Se added. The addition of Fe to animal feed might be beneficial to human Fe status, but only if the possible deleterious effect on meat quality can be overcome. Vitamin E supplementation of animal feed has a positive effect on meat quality but probably a very limited effect on human vitamin E status, as the content in the meat, even at very high levels of supplementation, is relatively low. Cu given in excess to animals does not appear to accumulate in animal muscle and therefore no benefits would be expected to human Cu status. The addition of vitamin A in excess of the animal's needs is not recommended, as the risk of reaching tissue concentrations known to be toxic to man is high. The effect on human status of supplementing domestic animals with other micronutrients remains to be explored.

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402 S. Bügel

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