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(54) Title: AIDING OF COGNITIVE FUNCTION

(57) Abstract: The invention provides the use of a bioavailable iron compound and a bioavailable zinc compound in a weight ratio of at least 2:1, and at least one B vitamin, in the manufacture of an edible composition, for use in aiding the cognitive development or cognitive performance of humans having an age of up to 18 years.

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## AIDING OF COGNITIVE FUNCTION

### FIELD OF THE INVENTION

The present invention relates to the use of certain micro-  
5 nutrients in the preparation of an edible composition for  
aiding the cognitive development or performance of humans  
having an age of up to 18 years. It also relates to a method of  
aiding the cognitive development or performance of such humans  
by using the aforementioned edible composition.  
10

### BACKGROUND OF THE INVENTION

In recent years there has been an increasing interest in the  
link between nutrition and cognitive performance in humans,  
15 especially, in the impact of nutrition on the development of  
cognitive abilities during childhood.

Iron is thought to be an important nutrient for the cognitive  
performance and development of children because the areas of  
20 the brain that are sensitive to iron deficiency (such as the  
cortex, frontal lobes and hippocampus) are the areas  
responsible for higher level cognitive functions such as  
executive functions and memory (Beard, 1995, American Journal  
of Nutrition, 62, 709-710).  
25

Zinc may have independent effects on cognitive performance, as  
well as an indirect effect through the enhancement of the  
efficacy of iron (Black & Miguel, 2001, The emerging roles of  
zinc in human nutrition, development and infectious disease,  
30 part 1, Nutrition Today, 36, 281-290). Zinc is important for  
cognitive performance and development of children because it is  
important for myelination and release of the neurotransmitters

GABA and glutamate that are key modulators of neuronal excitability (Bhatnagar & Taneja, 2001, Zinc & cognitive development. British Journal of Nutrition, 85, S139-S145). Intervention studies show that 20mg/day zinc plus other  
5 micronutrients has a positive effect on reasoning and visual recognition in Chinese and Mexican American school-aged children (Penland, 2000, Behavioural data and methodological issues in studies of zinc nutrition in humans, Journal of Nutrition, 130, 361S-364S).

10

Vitamin B-6 may affect the glutaminergic system and N-methyl-D-aspartate (NMDA) receptor that is important for learning and memory (Calvaresi & Bryan, 2001, B vitamins, cognition and ageing: A review. Journal of Gerontology: Psychological  
15 Sciences, 56, P327-P339). Correlational studies in children in India (Refsum et al., 2001 - Hyperhomocysteinemia and elevated methylmalonic acid indicate a high prevalence of cobalamin deficiency in Asian Indians. American Journal of Clinical Nutrition, 74, 233-241) and Indonesia (Setiawan, et al., 2000  
20 Vitamin B-6 inadequacy is prevalent in rural, and urban Indonesian children. Journal of Nutrition, 130, 553-558) have shown associations between B-12 and B-6 deficiency and poorer cognitive performance.

25 Certain omega-3 Polyunsaturated Fatty Acids (PUFAs) have been linked with cognitive performance. Studies have shown that infants who are fed formula milk supplemented with docosahexaenoic acid (DHA),  $\alpha$ -linolenic acid (ALA) and arachidonic acid (AA), and those who are breast-fed demonstrate  
30 better visual acuity and cognitive performance, including problem solving (Uauy et al, 2001, Essential fatty acids in

somatic growth and brain development. Nutrition and fitness: Diet, genes, physical activity and health. NY: Karger. pp.134-160).

- 5 WO03/003981 discloses compositions which are claimed to be beneficial in improving mental performance. These compositions comprise vitamin B12 on ion exchange resin, docosahexaenoic acid, certain herbal extracts and may comprise iron and/or zinc.
- 10 W000/62812 discloses nutritional compositions which are claimed to be beneficial for the improvement of cognitive performance. The composition comprise caffeine, choline, gamma aminobutyric acid, L-phenylalanine and taurine.
- 15 US2003/0044472 A1 discloses compositions for treating Attention Deficit/Hyperactivity Disorder. The compositions comprise omega 3-fatty acids, certain B vitamins and zinc.

US 2002/0045660 A1 discloses infant milk formulae comprising PDA.

- 20 Supplementation of infant formula milk with DHA is said to enhance neurological development of a pre-term infant.

Neumann et al (2002, contribution of animal source foods in improving diet quality and functions in children in the  
25 developing world, Nutrition Research, 22, 193-220) review the effect at iron, zinc, vitamins B12 and A on child growth, cognitive development and health.

- Abstract No. XP-002257665, Behavioral Responses of Young Anemic  
30 Indian Children to Iron Folic Acid Supplements discloses the results of supplementing young anemic Indian children with iron and folic acid supplements. The supplements are reported to raise

Hb levels and to provide improved intelligence test results, particularly in the performance section.

There is an increasing demand in the art for edible compositions  
5 that will aid the cognitive development or performance of humans, especially those having an age of up to 18 years. Preferably, such edible compositions can be added to ordinary foodstuffs, or, they can be consumed separately e.g. as a foodstuff in its own right.

10

The present invention seeks to provide edible compositions that aid the cognitive development and/or performance of humans, especially those having an age of up to 18 years.

15 In particular, it is an object of the invention to provide compositions that aid the cognitive development and/or performance of humans, especially those having an age of up to 18 years and that can be added to ordinary foodstuffs, or, can be consumed separately.

20

#### SUMMARY OF THE INVENTION

Surprisingly, it has now been found that edible compositions according to the invention, can aid the cognitive development or  
25 performance of humans, especially those having an age of up to 18 years. These edible compositions can be added to ordinary foodstuffs, or, can be consumed separately.

According to a first aspect, the present invention provides  
30 the use of;

1. a bioavailable iron compound and a bioavailable zinc compound in a weight ratio of at least 2:1, and
2. at least one B vitamin,

in the manufacture of an edible composition, for use in aiding the cognitive development or cognitive performance of humans having an age of up to 18 years.

5 According to a second aspect, the present invention provides a method for aiding the cognitive development or cognitive performance of humans having an age of up to 18 years, the method comprising the step of administering to said human by means of an edible composition;

- 10 1) a bioavailable iron compound, and a bioavailable zinc compound in a weight ratio of at least 2:1, and  
2) at least one B vitamin.

It is preferred according to both the second and first aspect  
15 of the invention that the edible composition after consumption aids the cognitive development or performance of humans having an age of from 6 to 9 years.

The terms "cognition" and "cognitive function" as used herein  
20 are interchangeable and refer to a construct of a range of functions:

- Intelligence (fluid, crystallised),
- Creativity (generate and open to new ideas, perceive new relationships),
- 25 - Memory (encoding, storage, retrieval),
- Executive functions (dealing with novelty, planning & implementation, monitoring performance, vigilance, inhibition of task irrelevant info),
- Cognitive resources (speed of information processing, working  
30 memory capacity, attentional capacity).

The term "cognitive development" as used herein refers to

the development of cognition (cognitive function) over time. The quality of cognition (cognitive function) in a human can be assessed by tests measuring cognitive performance. As cognition (cognitive function) of a growing child develops over time with  
5 brain development, measuring the cognitive performance over time for a given child, provides a measure of cognitive development. For example, for children aged 6-9 years, cognitive development is generally recognised by :

- 10 — A spurt in information processing capacity and/or executive functions.
- Development of the frontal lobes of the brain (involved in executive functions). These growth spurts may also occur at other periods during childhood (e.g. 0-2 and mid-teens).

15 The present invention provides a convenient and effective way of aiding the cognitive development or performance of humans having an age of up to 18 years.

Except in the operating and comparative examples, or where  
20 otherwise explicitly indicated, all numbers in this description indicating amounts of material or conditions of reaction, physical properties of materials and/or use are to be understood as modified by the word "about". All amounts are as percentages by weight unless otherwise stated. For the edible  
25 compositions, all percentages are by weight based on the total weight of the composition unless otherwise stated.

The term "comprising" is meant not to be limiting to any subsequently stated elements but rather to encompass non-  
30 specified elements of major or minor functional importance. In other words the listed steps, elements or options need not be exhaustive. Whenever the words "including" or "having" are

used, these terms are meant to be equivalent to "comprising" as defined above.

A serving of the edible composition as referred to herein  
5 refers to the amount of the edible composition that is provided to be consumed as a single portion, typically in a single sitting. For beverages, the typical serving size is in the range of from 100 to 500 ml and for a bar product or cookie etc will be in the range of from 20 to 100 g. However, it will be  
10 understood that the serving size will vary according to the type of edible composition.

#### DETAILED DESCRIPTION

15 According to the present invention, edible compositions that comprise the specific mixture of nutrients according to the present invention can be used to aid the cognitive development or performance of humans having an age of up to 18 years. Further details of the edible compositions and their components  
20 are given below.

#### Iron compound

The present invention uses a bioavailable iron compound, that is, an iron compound that is at least partially bioavailable to  
25 the body. Any bioavailable iron compound may be used according to the present invention although ferrous iron is generally better utilized by the body than ferric iron.

Bioavailable iron compounds which can be used include edible  
30 inorganic iron compounds such as edible ferrous compounds, for example ferrous sulfate. Whilst ferrous iron is typically more bioavailable, certain ferric salts can also provide highly bioavailable compounds of iron and these may also be used

according to the present invention. Suitable examples include ferric sulfate and ferric chloride and mixtures thereof.

Edible bioavailable iron organic compounds may also be used  
5 according to the present invention. Preferred examples include ferrous carboxylate salts such as ferrous fumarate, ferrous succinate, ferrous tartarate, ferrous lactate, ferrous gluconate, ferrous citrate and mixtures thereof. Whilst, as stated above ferrous iron is typically more bioavailable,  
10 highly bioavailable edible ferric compounds that may be used in the present invention include ferric carboxylate salts such as ferric ammonium citrate, ferric citrate and mixtures thereof.

An especially preferred bioavailable iron organic compound is  
15 ferric EDTA (sodium ferric ethylenediaminetetraacetic acid or sodium iron ethylenediaminetetraacetic acid, NaFeEDTA). Ferric EDTA has the advantage that it does not appreciably interchange with other cations often present in a edible compositions with added vitamin/mineral mixes (e.g., sodium, calcium, potassium,  
20 zinc, iodine, vitamin C, vitamin E, and the like) and thus no significant free iron is generated. This aids in the colour and flavour stability of the food product. It is also relatively stable in edible compositions and has good bioavailability.

25 Other edible iron organic compounds which can be used according to the present invention include bioavailable iron-sugar-carboxylate complexes. These materials are referred to herein as "complexes," but they may, in fact, exist in solution as complicated, highly hydrated, protected colloids. However, the  
30 term "complex" is used for the purpose of simplicity. Such materials are described in US 6 509 045 which is incorporated by reference herein.

In these iron-sugar-carboxylate complexes, the carboxylate provides the counterion for the ferrous (preferred) or ferric iron. Sugars that can be used to produce these complexes  
5 include any of the edible saccharidic sugars such as glucose, sucrose and fructose, mannose, galactose, lactose, maltose, and the like, with sucrose and fructose being preferred. Mixtures thereof may also be used. The carboxylic acid used may be any edible carboxylic acid such as citric acid, malic acid,  
10 tartaric acid, lactic acid, succinic acid, propionic acid, and mixtures thereof. Particular examples include ferric saccharide, ferric saccharate and mixtures thereof.

US 6 509 045 also discloses that it is known that iron is more  
15 bioavailable if administered in the form of chelates wherein the chelating ligands are amino acids or protein hydrolysates (see U.S. 3,969,540 and U.S. 4,020,158). These chelated iron compounds are known in the art by various names such as iron proteinate, iron amino acid chelates and peptide or  
20 polypeptide chelates. Such materials may also be used according to the present invention.

Particularly suitable ferrous amino acid chelates are those where the reacting ligands are glycine, lysine, and leucine,  
25 including the ferrous amino acid chelate where the reacting ligand is glycine.

Ferrous amino acid chelates particularly suitable as highly bioavailable amino acid chelated irons for use in the present  
30 invention are those having a ligand to metal ratio of at least 2:1. For example, suitable ferrous amino acid chelates having a ligand to metal mole ratio of two are those of formula:  $\text{Fe}(\text{L})_2$  where L is an alpha amino acid, dipeptide, tripeptide or

quadrapeptide reacting ligand. Thus, L can be any reacting ligand that is a naturally occurring alpha amino acid selected from alanine, arginine, asparagine, aspartic acid, cysteine, cystine, glutamine, glutamic acid, glycine, histidine, 5 hydroxyproline, isoleucine, leucine, lysine, methionine, omithine, phenylalanine, proline, serine, threonine, tryptophan, tyrosine and valine or dipeptides, tripeptides or quadrapeptides formed by any combination of these alpha amino acids. See U.S. Pat. No. 3,969,540 and U.S. Pat. No. 4,020,158 10 which are incorporated by reference.

Elemental iron may also be used according to the present invention and is considered herein to be a bioavailable compound even though, strictly speaking, it is not a compound. 15

Mixtures of one or more of any of the above bioavailable iron compounds may also be used. It is preferred that the bioavailable iron compound is selected from ferrous inorganic compounds, ferrous carboxylate salts, iron-sugar-carboxylate 20 complexes, chelated iron compounds and mixtures thereof. The bioavailable iron compounds, where appropriate, may be in a hydrated form. An especially preferred bioavailable iron compound is sodium ferric ethylenediaminetetraacetic acid or sodium iron ethylenediaminetetraacetic acid, known as NaFeEDTA. 25

The bioavailable iron compounds, may if desired, be in an encapsulated form.

The amount of bioavailable iron compound used in the edible 30 composition may be any amount which aids the cognitive development or performance of humans having an age of up to 18 years provided that the requirement for the weight ratio of iron to zinc according to the present invention is met. It is

preferred that an amount of up to 50mg of iron from the bioavailable iron compound is used in the edible compositions, more preferably 0.5 to 30 mg, most preferably 1 to 20 mg, such as 2 to 15mg per serving of the edible composition. These  
5 amounts are based on the weight of the iron in the bioavailable iron compound and not on the total weight of the bioavailable iron compound. Thus the weight of the bioavailable iron compound will be correspondingly greater depending upon the compound used.

10

#### Zinc Compounds

The present invention uses a bioavailable zinc compound, that is, a zinc compound that is at least partially bioavailable to the body.

15

Bioavailable zinc compounds which can be used include edible inorganic zinc compounds such as zinc oxide, zinc sulfate, zinc chloride and mixtures thereof.

20 Edible bioavailable zinc organic compounds may also be used according to the present invention. Preferred examples include zinc carboxylate salts such as zinc acetate, zinc gluconate, zinc ascorbate, zinc citrate, zinc aspartate, zinc picolinate and mixtures thereof.

25

Amino acid chelated zinc compounds of the typed referred to above for iron compounds may also be used.

Mixtures of one or more of any of the above bioavailable zinc  
30 compounds may also be used. Preferably the bioavailable zinc compound is selected from inorganic zinc salts, zinc carboxylate salts, chelated zinc compounds and mixtures

thereof. The compounds, where appropriate, may be in a hydrated form.

The bioavailable zinc compounds, may if desired, be in an  
5 encapsulated form.

According to one embodiment of the invention, it is particularly preferred that iron compound comprises NaFeEDTA and the zinc compound comprises zinc sulphate.7H<sub>2</sub>O.

10

The amount of bioavailable zinc compound used in the edible composition may be any amount which aids the cognitive development or performance of humans having an age of up to 18 years provided that the requirement for the weight ratio of  
15 iron to zinc according to the present invention is met. It is preferred that an amount of up to 50 mg of zinc from the bioavailable zinc compound is used in the edible compositions, more preferably 0.5 to 20 mg, most preferably 0.7 to 15 mg, such as 1 to 10 mg per serving of the food product. These  
20 amounts are based on the weight of the zinc in the bioavailable zinc compound and not on the total weight of the bioavailable zinc compound. Thus the weight of the bioavailable zinc compound will be correspondingly greater depending upon the compound used.

25

It is especially preferred that the weight ratio of the bioavailable iron compound to the bioavailable zinc compound is in the range of from 2:1 to 5:1, most preferably 2:1 to 4:1.

30 B vitamins

The present invention uses at least one B vitamin, preferably, selected from vitamins B6, B11 and B12. It is especially

preferred that the edible compositions comprises any two of vitamins B6, B11 and B12, even more preferably each of B6, B11 and B12.

5 The at least one B vitamin is preferably used in amounts of up to 5 mg in a serving of the edible composition, preferably 0.05 to 3 mg, more preferably 0.1 to 2mg.

It is preferred that Vitamins B6 and B11 are individually used  
10 in amounts of from 0.05 to 5 mg, more preferably from 0.075 to 3 mg, most preferably from 0.1 to 2 mg per serving of the edible composition.

It is preferred that Vitamin B12 is used in amounts of from  
15 0.0001 to 1 mg, preferably from 0.001 to 0.5 mg, most preferably from 0.0015 to 0.2 mg per serving of the edible composition.

For all the B vitamins above, the amount of the ingredient  
20 referred to is to the amount of the active ingredient.

Polyunsaturated fatty acid

The present invention preferably further uses a polyunsaturated fatty acid in the preparation of the edible composition.

25 Preferably, any oil comprising omega 3 fatty acids, especially docosahexaenoic acid (DHA, C20:5) and/or eicosapentaenoic acid (EPA, C22:5) may be used. Preferred omega 3 fatty acids are the following C18:3, C18:4, C20:4, C20:5, C22:5 and C22:6.

30 Preferably the polyunsaturated fatty acid may be incorporated by the inclusion in the edible composition of fish oil and/or an oil derived from algae.

The polyunsaturated fatty acid preferably comprises DHA and EPA in a weight ratio of at least 2:1, more preferably at least 3:1, such as at least 4:1.

- 5 The polyunsaturated fatty acid is preferably used in amounts of up to 2g per serving of the edible composition, more preferably 50 mg to 1 g per serving, more preferably 100 to 500 mg per serving.
- 10 It is especially preferred according to the present invention that the polyunsaturated fatty acid is used with either a bioavailable iron compound or a bioavailable zinc compound in the edible composition. Most preferred is that a polyunsaturated fatty acid, especially a mixture of DHA and EPA
- 15 is used with both a bioavailable iron compound and a bioavailable zinc compound in the edible composition.

#### Vitamins A and C

- It known that concurrent supplementation with vitamin C can
- 20 improve iron absorption by the body. Vitamin A may enhance iron mobilisation and vitamin A and beta carotene increase the absorption of non-heme iron (Mejia & Chew, 1988, Hematological effects of supplementing anaemic children with vitamin A alone and in combination with iron. American Journal of Clinical
- 25 Nutrition, 48, 1271-1276).

Vitamin A and/or vitamin are C preferably included in the edible composition.

- 30 Vitamin A is preferably used in amounts of from 0.5 to 50 mg, preferably 1 to 20 mg, such as 2 to 10 mg based on the retinol acetate equivalent amount. Vitamin C is preferably used in

amounts of from 1 to 100 mg, preferably 5 to 75 mg, such as 10 to 65 mg.

It is especially preferred according to the present invention 5 that the edible composition comprises in addition to the B vitamins recited herein a polyunsaturated fatty acid, a bioavailable iron compound, a bioavailable zinc compound and vitamins A and/or C, especially both vitamins A and C.

10 Types of edible composition

The edible composition of the invention may be in the form of a nutritional supplement (such as a tablet, powder, capsule or liquid product) or in the form of a food composition (product) including a beverage. Food compositions are especially 15 preferred according to the present invention.

It is preferred that the edible composition is a food composition which further comprises at least one of protein, carbohydrate or fat. It is especially preferred that the food 20 composition comprises at least protein and carbohydrate, most especially in the amounts given hereinbelow. Alternatively, a source of one of protein, carbohydrate and/or fat can be given as the same time as the edible composition, for example, if the edible composition does not contain substantial amounts of 25 these ingredients. In this case, the edible composition of the invention and an additional food product would provide a kit that could be used as according to the present invention. The comments herein to the protein, carbohydrate and fat content etc apply equally to the additional food product in such a kit.

30

The edible composition may be for example; a solid product, a powdered product, a liquid, a flowable, spoonable, pourable or spreadable product or a bar etc. The edible composition may be

a powder which is mixed with a liquid, such as water or milk, to produce a liquid or slurry product.

The edible compositions may be suitably selected from dairy based products (such as milk based products and drinks), soy based products including drinks, breads and cereal based products (including pasta or noodle based products and cereal bars), cakes, biscuits, ice creams, desserts, soups, porridge-type products, beverages including a ready-to-drink liquid, a liquid produced from a soluble powdered product, bar products, confectionery, snack foods, ready-to-eat meal products, pre-packed meal products, and dried meal products etc and soluble powdered products.

It is especially preferred that the edible compositions are in the form of a beverage (including dairy and soy based beverages), a bar product, a porridge, a biscuit or a cake.

Where an edible powder is provided according to the invention, this will be usually added to another food product, such as a beverage or a biscuit, to make consumption more convenient.

#### Protein

The edible composition further preferably comprises protein, this is preferred to provide energy to the consumer of the edible food product as well as the specific micronutrients described hereinabove. Any suitable protein may be used depending upon the type of edible composition.

Suitable sources of protein include dairy sources such as milk, yoghurt, kefir, cheese or cream. Other animal sources may also be used depending upon the type of edible composition. Alternatively the edible composition may comprise vegetable

derived protein such as soy-protein, rice protein, pea protein or wheat protein.

The amount of protein in the edible composition is preferably  
5 an amount of from 0.1 to 30 or 40% by weight of the edible  
composition, preferably 0.5 to 25%wt, most preferably 1 to 20  
%wt.

#### Carbohydrate

10 The edible compositions may also comprise one or more  
carbohydrates, preferably in an amount of from 1 to 90% by  
weight based on the weight of the composition, more preferably  
of from 5 to 80%wt, most preferably 10 to 75%wt. This is  
preferred to provide energy to the consumer of the edible food  
15 product as well as the specific micronutrients described  
hereinabove.

Any suitable carbohydrate may be used, for example sucrose,  
lactose, glucose, fructose, corn syrup, maltodextrins, starch,  
20 modified starch or mixtures thereof.

The edible composition may also comprise dietary fibres, for  
example in an amount of from 0.1 to 10% by weight based on the  
weight of the composition, preferably 0.5 to 5wt.

25

It is especially preferred that the edible composition  
comprises both protein and carbohydrate, preferably in the  
amounts given above.

#### 30 Fat

The edible composition may comprise edible fats in addition to  
any polyunsaturated fatty acid referred to above. The  
compositions preferably comprise an amount of up to 40% by

weight based on the weight of the composition of other fats, more preferably from 0.5 to 30 or 35%wt, most preferably from 0.75 to 10 or 20% fat.

5 Any suitable fat may be used with vegetable fats being especially preferred. Suitable examples include vegetable fats, plant oils, nut oils, seed oils, or mixtures thereof. Saturated or unsaturated (mono-unsaturated and poly-unsaturated) fats may be used, although saturated fats are less preferred.

10

The amounts of protein, fat, carbohydrate and other ingredients in the edible composition will vary according to the product format of the composition and also, where required, according to national or regional legislation.

15

#### Optional ingredients

The edible composition may optionally comprise fruit or vegetable pieces or particles, concentrates, juice or puree, preferably in amounts of up to 60% by weight of based on the  
20 weight of the edible composition. Preferably the compositions comprise 0.1 to 40%wt, more preferably 1 to 20%wt of these ingredients. The amount of these ingredients will depend upon the type of edible composition.

25 The edible composition may also comprise 0.1 to 15% by weight of edible salts based on the weight of the composition, preferably 3 to 8%wt. Any edible salts may be used, for example, sodium chloride, potassium chloride, alkali metal or alkaline earth metal salts of citric acid, lactic acid, benzoic  
30 acid, ascorbic acid, or, mixtures thereof. Calcium salts may also be used such as calcium chloride and calcium caseinate.

The edible composition may further comprise ingredients selected from other added vitamins and/or minerals, herbs, spices, flavourings, stabilisers, emulsifiers, aromas, antioxidants, colourants, preservatives or mixtures thereof.

5 These ingredients are used in conventional amounts.

These other added vitamins and/or minerals are preferably selected from at least one of vitamins D, E, H, K and calcium, magnesium, potassium, iodine, manganese, molybdenum,

10 phosphorus, selenium and/or chromium.

#### Preparation

The edible compositions of the invention may be prepared by any suitable method of preparation, including mixing, blending,

15 homogenising, high-pressure homogenising, emulsifying, dispersing, and/or encapsulating depending upon the type of edible composition. It is well within the skill of the person skilled in the art to select the most appropriate preparation method for different types of edible compositions.

20

#### Method of aiding the cognitive development or performance

The invention also provides the aforementioned method for aiding the cognitive development or performance of humans having an age of up to 18 years according to the second aspect

25 of the invention.

The total effective administered to a subject according to the invention may vary according to the needs of the person to whom it is administered, e.g. age, general nutrition status etc.

30 Typical amounts administered by the administration of the edible composition for each of the micronutrients according to the invention are listed above under the heading referring to that micronutrient. These are the typical amounts administered

per day per micronutrient. An effective daily amount may be administered by a single dose or by multiple doses.

The consumption of the edible composition of the invention preferably occurs as a part of the daily diet. It is preferred that the edible composition is consumed daily.

The invention is further described by way of the following examples that are to be understood as not limiting. Further examples within the scope of the invention will be apparent to the person skilled in the art.

#### EXAMPLES

##### 15 Example 1 - beverage

5 g of a powder having the composition shown in table 1 was added to 100 ml of a commercially available aqueous soy-based drink (ADES soy based fruit drink) comprising 0.6% wt of soy protein. The resulting beverage was mixed until the powder had dissolved.

Table 1 - composition of the powder

Ingredient	Amount of compound	Amount of active ingredient
FeNaEDTA	84 mg	10 mg iron
ZnSO <sub>4</sub> .7H <sub>2</sub> O	23.1 mg	5 mg zinc
Fish oil powder *1	763 mg	88 mg DHA and 22 mg EPA
Vitamin B6 (pyridoxine.HCl)	1.34 mg	1 mg
Vitamin B11 (folic acid)	0.17 mg	0.15 mg
Vitamin B12 (cyanocobalamin, 0.1 % WS)	1.65 mg	0.0015 mg

Vitamin A (retinol acetate dry, 325.000 IU/g)	4.51 mg	0.4 mg (as retinol equivalent)
Vitamin C (Ascorbic acid)	54 mg	54 mg
Maltodextrin	2640 mg	2640 mg
Lecithin	300 mg	300 mg
Citric acid	40 mg	40 mg
Powdered sugar	1067 mg	1067 mg
Flavours	20 mg	20 mg

\*1 - the fish oil powder contained 20% wt in total of DHA/EPA in a weight ratio of 4.5:1.

The edible composition can be consumed by humans aged up to 18 5 years of age to aids cognitive development or performance.

**Claims**

1. The use of:
  1. a bioavailable iron compound and a bioavailable zinc compound in a weight ratio of at least 2:1, and
  2. at least one B vitamin,in the manufacture of an edible composition, for use in aiding the cognitive development or cognitive performance of humans having an age of up to 18 years.
2. The use according to claim 1, wherein the edible composition comprises 0.5 to 30 mg per serving of the edible composition of iron from the bioavailable iron compound.
3. The use according to either one of claims 1 or 2, wherein the bioavailable iron compound is selected from ferrous inorganic compounds, ferrous carboxylate salts, iron-sugar-carboxylate complexes, chelated iron compounds and mixtures thereof.
4. The use according to any one of the preceding claims, wherein the edible composition comprises 0.5 to 20 mg per serving of the edible composition of zinc from the bioavailable zinc compound.
5. The use according to any one of the preceding claims, wherein the bioavailable zinc compound is selected from inorganic zinc salts, zinc carboxylate salts, chelated zinc compounds and mixtures thereof.

6. The use according to any one of the preceding claims, wherein the wherein the weight ratio of the bioavailable iron compound to the bioavailable zinc compound is in the range of from 2:1 to 5:1.
7. The use according to any one of the preceding claims, wherein the edible composition further comprises at least one polyunsaturated fatty acid.
8. The use according to claim 7, wherein the edible composition comprises up to 2 g per serving of the polyunsaturated fatty acid.
9. The use according to either one of claims 7 or 8, wherein the polyunsaturated fatty acid comprises docosahexaenoic acid and eicosapentaenoic acid in a weight ratio of at least 2:1.
10. The use according to any one of the preceding claims, wherein the at least one B vitamin is selected from vitamins B6, B11 and B12.
11. The use according to any one of the preceding claims, wherein the edible composition further comprises vitamin A and/or vitamin C.
12. The use according to any one of the preceding claims, wherein the edible composition is a food composition.
13. The use according to any one of the preceding claims, wherein the food composition further comprises protein and/or carbohydrate.

- 14.A method for aiding the cognitive development or cognitive performance of humans having an age of up to 18 years, the method comprising the step of administering to said human by means of an edible composition;
- 1) a bioavailable iron compound, and a bioavailable zinc compound in a weight ratio of at least 2:1, and
  - 2) at least one B vitamin
- 15.A method according to claim 13, wherein the edible composition in a food composition.

# INTERNATIONAL SEARCH REPORT

International Application No  
PCT/EP2004/006817

<b>A. CLASSIFICATION OF SUBJECT MATTER</b> IPC 7 A23L1/302 A23L1/304 A23L1/30 A61K33/26 A61K33/30				
According to International Patent Classification (IPC) or to both national classification and IPC				
<b>B. FIELDS SEARCHED</b> Minimum documentation searched (classification system followed by classification symbols) IPC 7 A23L A61K				
Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched				
Electronic data base consulted during the international search (name of data base and, where practical, search terms used) EPO-Internal, WPI Data, PAJ, FSTA, MEDLINE, BIOSIS				
<b>C. DOCUMENTS CONSIDERED TO BE RELEVANT</b>				
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.		
X	WO 03/003981 A (SMITH KYL L ;VITAL BASICS INC (US)) 16 January 2003 (2003-01-16) page 4, line 14 - page 8, line 21 page 10, lines 17-28 page 12, lines 9-19 page 15, line 30 - page 16, line 23 table 1 page 23, line 16 - page 25, line 8 ----- -/--	1-15		
<input checked="" type="checkbox"/> Further documents are listed in the continuation of box C. <span style="margin-left: 200px;"><input checked="" type="checkbox"/> Patent family members are listed in annex.</span>				
° Special categories of cited documents :				
<table style="width: 100%; border: none;"> <tr> <td style="width: 50%; border: none; vertical-align: top;"> <ul style="list-style-type: none"> <li>*A* document defining the general state of the art which is not considered to be of particular relevance</li> <li>*E* earlier document but published on or after the international filing date</li> <li>*L* document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)</li> <li>*O* document referring to an oral disclosure, use, exhibition or other means</li> <li>*P* document published prior to the international filing date but later than the priority date claimed</li> </ul> </td> <td style="width: 50%; border: none; vertical-align: top;"> <ul style="list-style-type: none"> <li>*T* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention</li> <li>*X* document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone</li> <li>*Y* document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.</li> <li>*Z* document member of the same patent family</li> </ul> </td> </tr> </table>			<ul style="list-style-type: none"> <li>*A* document defining the general state of the art which is not considered to be of particular relevance</li> <li>*E* earlier document but published on or after the international filing date</li> <li>*L* document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)</li> <li>*O* document referring to an oral disclosure, use, exhibition or other means</li> <li>*P* document published prior to the international filing date but later than the priority date claimed</li> </ul>	<ul style="list-style-type: none"> <li>*T* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention</li> <li>*X* document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone</li> <li>*Y* document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.</li> <li>*Z* document member of the same patent family</li> </ul>
<ul style="list-style-type: none"> <li>*A* document defining the general state of the art which is not considered to be of particular relevance</li> <li>*E* earlier document but published on or after the international filing date</li> <li>*L* document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)</li> <li>*O* document referring to an oral disclosure, use, exhibition or other means</li> <li>*P* document published prior to the international filing date but later than the priority date claimed</li> </ul>	<ul style="list-style-type: none"> <li>*T* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention</li> <li>*X* document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone</li> <li>*Y* document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.</li> <li>*Z* document member of the same patent family</li> </ul>			
Date of the actual completion of the international search	Date of mailing of the international search report			
13 August 2004	02/09/2004			
Name and mailing address of the ISA	Authorized officer			
European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016	Couzy, F			

## INTERNATIONAL SEARCH REPORT

International Application No  
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## C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	DATABASE WPI Week 1997 Derwent Publications Ltd., London, GB; AN 1997-213445 XP002292286 GAO C, HUANG T, ZHANG W: "Nutritious oral liq. contg. natural ingredients. vitamins and trace elements, promotes bone growth and brain development" & CN 1 097 607 A (MEDICAL INST 999 ENTERPRISE GROUP) 25 January 1995 (1995-01-25) abstract	1-15
X	----- DATABASE WPI Week 1997 Derwent Publications Ltd., London, GB; AN 1997-458010 XP002292287 CHEN W, JIANG X, LIU J: "Nutritious soy sauce and its production" & CN 1 117 816 A (CHEN W) 6 March 1996 (1996-03-06) abstract	1-6, 10-15
Y	abstract	7-9
X	----- DATABASE WPI Week 2003 Derwent Publications Ltd., London, GB; AN 2003-514435 XP002292288 ZHAO G: "Production method of coated and shaped tablet health-care food with the functions of increasing body height and reducing weight for youngsters" & CN 1 411 747 A (ZHAO G) 23 April 2003 (2003-04-23) abstract	1-6, 11-15
Y	abstract	7-9
X	----- US 2002/045660 A1 (O'CONNOR DEBORAH L ET AL) 18 April 2002 (2002-04-18) paragraphs '0064!', '0066!', '0068!', '0071!' - '0073!', '0075!', '0116!' - '0129! page 7; table B	1,7,9, 11,12
X	----- NEUMANN C, HARRIS DM, ROGERS LM: "Contribution of animal source foods in improving diet quality and function in children of the developing world" NUTRITION RESEARCH, vol. 22, 2002, pages 193-220, XP002257664 paragraphs '2.1.1!' - '2.1.3! paragraphs '3.3.1!' - '3.3.2.C! pages 206-207, paragraph 3.3.2A pages 207-208, paragraphs 3.3.2B,,3.3.2C paragraph '3.3.1!	1-6, 10-15
Y	paragraph '3.3.1!	7-9
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## INTERNATIONAL SEARCH REPORT

International Application No  
PCT/EP2004/006817

## C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y	WO 00/62812 A (ADVOCARE INTERNAT LLC) 26 October 2000 (2000-10-26) page 12, lines 1-5 example 1 -----	1-15
Y	US 2003/044472 A1 (LANG PHILIP C) 6 March 2003 (2003-03-06) page 1, paragraphs 2,13-19,28 page 2, paragraphs 32,33,36,37 page 3, paragraph 46 claim 1 -----	1-15
Y	DATABASE BIOSIS 'Online! BIOSCIENCES INFORMATION SERVICE, PHILADELPHIA, PA, US; SESHADRI S, HIRODE K, NAIK P, MALHOTRA S: "Behaviorial responses of young anemic Indian children to iron folic-acid supplements" XP002257665 retrieved from BIOSIS Database accession no. PREV198375074283 abstract -----	1-10,13, 14
Y	KOLETZKO B ET AL: "GROWTH, DEVELOPMENT AND DIFFERENTIATION: A FUNCTIONAL FOOD SCIENCE APPROACH" BRITISH JOURNAL OF NUTRITION, CAMBRIDGE, GB, vol. 80, no. SUPPL 1, 1998, pages S05-S45, XP009027350 page S29 - page S30 -----	1-15

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International application No.  
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### Box II Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)

This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1.  Claims Nos.:  
because they relate to subject matter not required to be searched by this Authority, namely:  

Although claims 14-15 may encompass a method of treatment of the human body, the search has been carried out and based on the alleged effects of the composition.
2.  Claims Nos.:  
because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3.  Claims Nos.:  
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

### Box III Observations where unity of invention is lacking (Continuation of item 3 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:

1.  As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2.  As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3.  As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4.  No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:

Remark on Protest

- The additional search fees were accompanied by the applicant's protest.
- No protest accompanied the payment of additional search fees.

# INTERNATIONAL SEARCH REPORT

Information on patent family members

International Application No  
PCT/EP2004/006817

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			WO 0062812 A1	26-10-2000
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US 2003044472	A1	06-03-2003	NONE	
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