



November 27, 2014

To whom it may concern,

**STATEMENT OF FOOD USE FOR
BIOKYOWA's L-CITRULLINE**

We, KYOWA HAKKO BIO ITALIA SRL, on behalf of BIOKYOWA INC., hereby confirm that BIOKYOWA's L-Citrulline can be used as food ingredient in accordance to EU regulation for the following reason.

FOOD SAFETY

BIOKYOWA's L-Citrulline can be used as food ingredient because:

- is manufacturer in accordance to Good Manufacturing Procedure (GMP) with ISO9001 and HACCP certification/accreditation;
- its production process conforms to the regulations in EU for the manufacture of alimentary ingredients;
- it meets the requirements of Regulation (EC) No. 178/2002 "laying down the general principles and requirements of food law, establishing the European Food Safety Authority and laying down procedures in matters of food safety".

ORIGIN

BIOKYOWA's L-Citrulline is made in USA, from fermentation followed by purification in USA.

The ingredient of fermentation is Glucose, manufactured in USA and derived from corn (*Zea mays*).

Throughout the manufacturing process of BIOKYOWA's L-Citrulline, neither raw material of animal nor human origin is used or added.



ALLERGENS

BIOKYOWA's L-Citrulline is free from allergen in accordance to According REGULATION (EU) No 1169/2011 on the provision of food information to consumers.

The REGULATION (EU) No 1169/2011 aims at the obligation of clearly labelling the allergens below, used in food products and still present in the product, even in a modified form, including additives and processing aids.

In addition we can declare that there is no risk of Cross-Contamination with the material listed below.

	Direct incorporation as a base raw material or in a derivative form (<u>don't forget the carriers, the additives and the processing aids</u>)	
	Yes (precise the nature)	NO
Cereals containing Gluten (1) and products thereof		X
Crustaceans and products thereof		X
Eggs and products thereof		X
Fish and products thereof		X
Peanuts and products thereof		X
Soybeans and products thereof		X
Milk and products thereof (including Lactose)		X
Nuts and products thereof (2)		X
Celery and products thereof		X
Mustard and products thereof		X
Sesame seeds and products thereof		X
Sulphur dioxide and sulphites at conc. of more than 10 mg/kg or 10 mg/litre expressed as SO ₂		X
Lupin and products thereof		X
Molluscs and products thereof		X

(1) The word Gluten is here the general term. It applies to all the cereals which contains gluten, of which: Wheat, Rye, Barley, Oats, Spelt, Kamut or their hybridised strains.

(2) Almond, Hazelnuts, Walnut, Cashew, Pecan nut, Brazil nut, Pistachio nut, Macadamia nut or Queensland nut



CONTAMINANTS

There are no possibilities for BIOKYOWA's L-Citrulline to be contaminated with materials shown below during manufacturing processes because:

- BIOKYOWA's L-Citrulline can be considered free from materials listed above.
- BIOKYOWA's L-Citrulline is pure material with assay value of "99.0–101.0%", and no materials are added after the purification.

For the above reason we can declare that:

- MELAMINE:
BIOKYOWA's L-Citrulline is considered as free from MELAMINE and its analogues (AMMELIDE, AMMELINE and CYANURIC ACID). In addition is in compliance with Commission Regulation (EC) No. 1881/2006 and its amendments (including 594/2012).
- PESTICIDES
BIOKYOWA's L-Citrulline is considered as free from PESTICIDE and complies with Regulation (EC) No. 396/2005 and its amendments (including 91/414/EC).
- LATEX AND GLUTEN
BIOKYOWA's L-Citrulline is considered free from Gluten and Latex and there is no risk of cross-contamination with these materials.
- OTHERS CONTAMINANTS
BIOKYOWA's L-Citrulline is considered free from:
 - Aflatoxin
 - Dioxin
 - Casein
 - Ethylene Oxide
 - Lactose
 - Mesilate Esters

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- Phthalate
- Preservatives

BSE/TSE

Throughout the manufacturing process of BIOKYOWA's L-Citrulline neither source materials nor production processes as defined in Section 2 of the "Note for Guidance on Minimizing the Risk of Transmitting Animal Spongiform Encephalopathy Agents via Human and Veterinary Medical Products" (EMEA/410/01 rev. 3) and in chapter 5.2.8 in the current EP are used.

GMO

BIOKYOWA's L-Citrulline is not applicable to the:

- a) "REGULATION (EC) No 1829/2003 OF THE EUROPEAN PARLIAMENT AND OF THE COUNCIL of 22 September 2003" as well as
- b) "REGULATION (EC) No 1830/2003 OF THE EUROPEAN PARLIAMENT AND OF THE COUNCIL of 22 September 2003"

on genetically modified food and feed as well as related traceability and labelling.

In Regulation (EC) No 1829/2003, as a Scope, there is following description.

CHAPTER II GENETICALLY MODIFIED FOOD

Section 1 Authorization and supervision

Article 3 Scope

1. This Section shall apply to:

- (a) GMOs for food use;
- (b) food containing or consisting of GMOs;
- (c) food produced from or containing ingredients produced from GMOs.

As BIOKYOWA's L-Citrulline is manufactured using processes out of this scope, we are in a position to confirm that food produced from or containing BIOKYOWA's L-Citrulline is excluded from the scope of above mentioned guidelines.



HEAVY METAL

During the strictly controlled manufacturing process of BIOKYOWA's L-Citrulline, it will comply with the specifications shown below if tested, in accordance to EC Regulation No. 629/2008.

Lead: Not More Than 3.0 ppm

Cadmium: Not More Than 1.0 ppm

Mercury: Not More Than 0.1 ppm

IRRADIATION

BIOKYOWA's L-Citrulline is non-irradiated and comply with DIRECTIVE 1999/2/EC OF THE EUROPEAN PARLIAMENT AND OF THE COUNCIL on approximation of the laws of the Member States concerning foods and food ingredients treated with ionising radiation.

NANOMATERIALS

BIOKYOWA's L-Citrulline is out of scope of engineered nanomaterial defined in REGULATION (EU) No 1169/2011 OF THE EUROPEAN PARLIAMENT AND OF THE COUNCIL of 25 October 2011.

Also, engineered nanomaterials are not used in the manufacturing process or the package of BIOKYOWA's L-Citrulline.

WADA

BIOKYOWA's L-Citrulline is not a material listed in the following documents:

- 2014 prohibited list issued by WADA (World Anti-Doping Agency)
- NFL-NFLPA Banned Substances List
- Athletic Banned Substances of NSF International Certification Guideline

During the manufacturing process of BIOKYOWA's L-Citrulline and that there is no possibility to be contaminated with any substances in the list into those products.

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SOLVENTS

BIOKYOWA's L-Citrulline is in accordance to *Directive 2009/32/EC on the approximation of the laws of the Member States on extraction solvents used in the production of foodstuffs and food ingredients.*

Any solvent listed in the directive above mentioned is not used as "Extraction Solvent".

PACKAGING STATEMENT

The packaging of BIOKYOWA's L-Citrulline is packed into double polyethylene bags which are packed into fiber drums.

The primary packaging materials are suitable for packaging of pharmaceutical ingredients, feed or food stuffs.

They are in compliance with the following regulations:

<u>CPMP/QWP/4359/03:</u>	Guideline on plastic immediate packaging materials for active substances and medicinal products.
<u>EC 1935/2004:</u>	materials and articles intended to come into contact with food
<u>EU 10/2011:</u>	plastic materials and articles intended to come into contact with food
<u>EC 2023/2006:</u>	good manufacturing practice for materials and articles intended to come into contact with food.

In addition they are stable and safe.

If you have any additional questions or further requirements, please feel free to contact us.

Sincerely yours;


Riccardo Carcano

Qualified Person

KYOWA HAKKO BIO ITALIA SRL