



Production plant video updates

In this section, you'll be able to view the videos updated every month and see the construction of the Bio-on plant in Castel San Pietro, Bologna as it develops. Click on an image to view the video.

VIDEO #009 ITA 720

video HD #008 30_04_2018

video HD #007 31_03_2018

video HD #006 28_02_2018

video HD #005 31_01_2018

video HD #003 31_12_2017

video HD #003 30_11_2017



video HD #001 14_09_2017

Special PHAs production plant (inauguration summer 2018)

A new production plant dedicated solely to special PHAs production for advanced niche products in rapid development and not in competition with the large-scale plants licensed by Bio-on around the world for PHAs output from 5,000 to 10,000 tons per year. This new plant based in Castel San Pietro Terme in the province of Bologna has 3,700 covered m², 6,000 m² land for development and a total area 30,000 m². The plant has a capacity dedicated to the research into and production of 1,000 tons per year rapidly expandable to 2,000 tons per year. The plant is equipped with the most modern technologies and the most advanced research and development laboratories. New agricultural waste carbon sources for producing biopolymers are continuously tested to increase the range of technologies offered by Bio-on.

Bio-on Plants SRL

Via Legnana 1900
Castel San Pietro Terme località Gaiana Bologna Italy

All enquiries: info@bio-on.it



Bio-on's technology has been adapted to by-products of the following:

- sugar beet
- sugar cane
- glycerol from biodiesel
- potatoes
- animal fat
- fruit
- vegetables

- wood
- domestic wet waste
- wine production waste

Bio-on Plants logo

Bio-on Plants is a new company set up by Bio-on to meet the increasing number of requests in PHAs production (polyhydroxyalkanoates) for special uses. Bio-on Plants is an exclusively productive unit for the study and development of PHAs through bacterial fermentation using the technologies developed by Bio-on and licensed around the world. The mission is to create new production of special biopolymers and at the same time offer a new range of products to be licensed all over the world. Bio-on Plants produces the varieties of MINERV PHAs invented and developed by Bio-on S.p.A.'s technicians.

The Bio-on Plants logo and all rights pertaining to it are the property of Bio-on spa.



bio-on plants logotype

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Minerv PHAs logo

MINERV is the logo owned by Bio-on spa for the generic production of Polyhydroxyalkanoate or PHAs - a linear polyester produced naturally through the bacterial fermentation of sugar. The family includes over 100 different monomers for producing materials with extremely different properties. Thermoplastics or elastomers can be created, with a melting point varying from 40 to over 180°C. **MINERV-PHA** is a high-performance PHA biopolymer. **MINERV-PHA** has excellent thermal properties. Through characterisation, it is possible to meet production requirements from -10°C to +180°C. The product is particularly suited for the production of objects by injection or extrusion. It also replaces highly polluting products such as PET, PP, PE, HDPE, LDPE.

The MINERV logo and all rights pertaining to it are the property of Bio-on spa.

The image shows the Minerv logo. The word "minerv" is written in a lowercase, sans-serif font. The letters "m", "i", and "n" are green, while the letters "e", "r", "v", and the registered trademark symbol (®) are grey.

Reference markets for Bio-On Plants direct production

Bio-on Plants offers a vast range of products for use in the cosmetics and biomedical sector. The characteristics of the oil-based products used today in these fields can be replaced by MINERV PHAs. The particular conditions designed in production enable the immediate use to replace microbeads (micro-particles of plastic) in the cosmetics sector.



minerv®
bio
cosmetics

natural
beauty
ingredients

General information

A number of associations are active in different parts of the world. The associations are particularly focused on constantly communicating the problem surrounding the use of polymers in beauty products (cosmetics). Since 2013, they have strongly denounced the problem of ocean pollution caused by some cosmetics and body care products, particularly skin exfoliants. This category of products has become the figurehead in a series of products (e.g. shampoo, creams, make-up and many more) cited in the recent law proposed in the USA and accepted by various states in the union.

The official publication of this new USA law, which prevents the use of oil-based polymers in body care products and particularly the category defined as "exfoliants" by the Obama administration (President of the U.S.A.), generates an enormous opportunity for the platform products generated by **Bio-on (PHAs, Polyhydroxyalkanoates)** as well as its particular applications, as is the case with the new patent for the beauty products sector called **minerv pha bio cosmetics type C1**.

The stance taken by the **USA** against the plastics that in nano and micrometric dimensions form polluting **Microbeads** is aimed exclusively at the "conventional" polymers widely used in cosmetics, as stated in the law: polyethylene (PE), polymethyl methacrylate (PMMA), nylon, **polyethylene terephthalate (PET) and polypropylene (PP)**. Oil-based polymers not compostable in the environment not only pollute by their very presence but also increase pollution due to the prolonged periods that they persist in the water and soil. Replacing these oil-based polymers with bio-based polymers or even biopolymers would not change the problem of pollution in the environment and, particularly, in the seas. We believe that the only choice capable of producing real change is to pursue not just **Plastic Free Seas**, but **Microbead Free Seas**. This is easy to achieve, using polymers that are truly capable of being completely broken down by the bacterial component within a short time and, therefore, not capable of forming **Microbeads**. For this problem, the time taken for biodegradation and the metabolites formed as a consequence of this degradation are very important. **Minerv bio cosmetics** guarantees a relatively short persistence of **PHA** in water and soil and that the derivatives of bacterial aggression are biocompatible, and even useful to the environment as nutrients for micro-organisms (a very positive factor for the environment). **Bio-on has a unique opportunity to propose to the cosmetics world to safeguard the environment without reducing the performance and effectiveness of their products by using bio-based PHAs**, which does not accumulate in the environment because it is broken down

by the bacterial component into molecules and metabolites, the presence of which is not only **NON-toxic** or harmful, but in certain aspects even necessary to feed specific biological processes, growth of plants, micro-organisms and insects.

Minerv PHA for cosmetics

Since 2007, Bio-on has been constructing a more sustainable future for the plastics sector, with proprietary know-how regarding the production of PHAs (polyhydroxyalkanoates), which are considered the best biopolymers by those consciously looking to the future. PHAs are plastics made 100% from renewable waste plant sources, with no competition with food supply chains, and are 100% biodegradable in a variety of environmental conditions, without dispersal of residue following biodegradation.

Bio-on designed and patented the world's first fully bio-based PHAs plastic (certified since 2014 by the United States Department of Agriculture - **USDA**) and 100% naturally biodegradable in water and soil (certified since 2008 by **Vinçotte**) without the use of chemical solvents. This exceptional product is obtained through the natural fermentation of bacteria fed by by-products from the agricultural industry (no human food). Bio-on biopolymers have exceptional properties that adapt to the injection and extrusion methods currently in use in the plastic industry and can cover a vast range of strategic applications: biomedical, packaging, design, clothing, automotive and many more.



Material ConneXion®

Minerv PHAs bio cosmetics (type C1)

These are new biocompatible complexes, including a polyhydroxyalkanoate (**PHA**) and a biomimetic inorganic compound, that can be used advantageously for the formulation of cosmetics and personal care products. The complex can act not only as a biodegradable polymeric component that remains for long periods on the skin and hair keratin even after prolonged rinsing, but also as an innovative carrier and dispenser of specific active substances, the bio-activity of which is prolonged after each application.

minerv bio cosmetics (type C1) brings together respect for the environment and human health.

Made from agriculture-based carbon sources, it is not made from oil-based products, but from the **atmospheric CO₂** captured by the crops from which it is made.

It is totally biodegradable, unlike any other biopolymer, and respects the environment.

PHAs molecules are also present in the human body; they are biocompatible and naturally absorbed once their action is carried out. Everyday products such as lipstick, lip gloss, mascara, eye-liner, nail polish, creams and shampoo contain plastic polymers, especially in the form of **microbeads**.

To date, all the plastic polymers used in the cosmetics world are obtained from fossil fuels, oil, and hydrocarbons in general.

This apparently limited yet immense worldwide use implies the extraction of oil, increased greenhouse effect, and the dispersion of non-biodegradable material into the environment.

This non-biodegradable oil-based plastic permeates the natural cycle down to the plankton in the rivers and oceans. Plankton is able to swallow the microbeads, thereby introducing them into the food chain.

All of these environmental impacts disappear with **MINERV PHA**; since it is biodegradable, **PHAs** microbeads do not accumulate in the environment. What is more, it is a bio-compatible and naturally bioabsorbable material coming into contact with the skin, lips, hair and mucous membranes.

Another important aspect: **Bio-on** has discovered that **minerv bio cosmetics type C1** is capable of binding active molecules and antioxidants, such as **coenzyme Q10**, vitamins, proteins, and active substances in general, transporting them naturally to parts of the body where cosmetics products are normally applied. After having located the active substance where it must act, **MINERV-PHA** is naturally washed off or absorbed without a trace.

 Video Micorbeads





[Download US Law](#)



Areas for development and production in the cosmetics world

MinervPHA Bio Cosmetics is the new range of intermediate products for cosmetics, designed to makes beads suitable for all uses in the cosmetics industry.

The product range is divided into 3 macro-families:

✓ SCRUBS

The **natural alternative to polyethylene beads (PE)**.

Available in sizes varying from **100 to 800 micron**, they are suitable for **soft, mild or hard** exfoliation and can be used in all applications: body and face scrubs, cleansers, toothpastes...

Easy to process and stabilise, they are a **versatile solution** for all types of formulation (W/O emulsions, O/W emulsions, Gels...)

✓ TEXTURING

Highly **spherical ultra-fine powders** (micropowders) available in different sizes from **5 to 20 micron** designed specifically for texturing, i.e. for modifying the **sensory and tactile experience** of

cosmetics products:

Added to the formulation, they help improve:

- initial feel
- ease of application
- post-application skin feel

Their special characteristics modify the structure of the cream, varying its adhesiveness and giving an "elegant feeling".

The micropowders in the **MinervPHA Bio Cosmetics** line are **hydrophobic**, so they do not absorb the skin's humidity, and highly **lipophilic**, which means they are very good at **absorbing oils and sebum**.

The latter characteristic gives even the greasiest skin a natural matt look. A further optical effect of **MinervPHA Bio Cosmetics** micropowders is the excellent **Soft Focus** they provide, i.e. the ability to flatten wrinkles and creases in the skin and add brightness to the face.

✓ ACTIVE INGREDIENTS

Thanks to **patented technology**, this line of products based on micropowders for texturing **incorporates a wide range of active substances** (UV filters, anti-ageing, antibacterial, anti-cellulite, toning, etc.).

The biodegradable carrier shell provides more effective, stable and processable protection than the unstable ingredients susceptible to chemical degradation found in conventional formulations. The polymeric structure also guarantees a more controlled, longer-lasting release.







Advanced research and production team minerv PHA bio cosmetics

The industrial process developed by **Bio-on** has features that are unique in the technological panorama of the production and commercialisation of biopolymers. Our process allows us to focus on peerless "naturalness", since we do not use organic solvents and do not genetically modify bacteria and plants. These aspects enable the successful use of the **PHAs** obtained from the Bio-on process in many food applications and applications directly linked to living beings, both intracorporeal and extracorporeal. This is why we have already successfully developed scaffolds for the production of bone and arterial tissue. All of these products further enrich the **Bio-on** products portfolio. The new line of **minerv bio cosmetics** products is now available for finalising their use in myriad cosmetics applications. Our research and development team has developed an intense product development schedule for the 2016-2020 period, in part thanks to the replacement of the polymer particles used today in toothpaste, shampoo, exfoliants (those cited in the USA law), face make-up, tanning and sun creams and many other cosmetics products.

Bio-on is willing to collaborate with other private research centres to develop new vertical applications of **minerv bio cosmetics type C1**.

Our products, based on a polymer obtained through fermentation and 100% naturally and totally biodegradable in a short time (especially in small particles) in soil and water, is currently the most advanced product in the world for easily replacing the plastic used in all beauty products without any legislative limitation.

Video Links:

https://www.youtube.com/watch?v=uAilGd_JgZc

<https://www.youtube.com/watch?v=mGzlz9Ld-sE>

<https://www.youtube.com/watch?v=pfq000AF1i8>

USA links

<http://www.fda.gov/Cosmetics/GuidanceRegulation/LawsRegulations/ucm2005209.htm>

<https://www.congress.gov/bill/114th-congress/house-bill/1321/text>

EU links

<http://eur-lex.europa.eu/legal-content/IT/TXT/?uri=URISERV%3Aco0013>

http://ec.europa.eu/growth/sectors/cosmetics/legislation/index_en.htm



Nanotechnologies and bioplastic to diagnose and combat tumours Bio-on researchers on the job.

For the very first time, bioplastic can be used to diagnose and treat tumours thanks to the first patent registered by Bio-on in the nanomedical field, particularly in nanodiagnostics (nano-imaging). Bio-on researchers use nanotechnologies to create **minerv BIOMEDS**: these are revolutionary and innovative nanocapsules in PHAs bioplastic (polyhydroxyalkanoates) capable of simultaneously containing two contrast media: magnetic nanoparticles and gold nanocylinders. These two elements flag up diseased areas of the body, e.g. a tumour mass, using traditional **Nuclear Magnetic Resonance** and the more innovative **Photoacoustic imaging***.

*"Using PHA bioplastic is very advantageous because it is safe for the patient and has no side effects. **These products are safe for the environment and human health**, especially when it comes to biomedical applications. Bio-on bioplastics fully meet these requirements and open up important, unexplored fields for nanotechnologies in medicine - a rapidly growing sector."*

This technology has a **diagnostic** as well as a **therapeutic** function, given that drugs can be inserted into the nanocapsules, for chemotherapy for example. This will enable **minerv BIOMEDS** nanocapsules to be used in targeted and selective cancer therapies in the future. Combining **Therapeutic** with **Diagnostics** has led the two terms to be fused, thus creating **Theranostics**.

This versatility makes nanocapsules multifunctional and this dual system will allow clinical theranostic applications in oncology and in neurodegenerative disorders, enabling medical professionals to work safely on patients.

The **PHAs bioplastics developed by Bio-on are made from renewable plant sources with no competition with food supply chains. They are completely eco-sustainable and 100% naturally biodegradable.** The research conducted and the patent registered by Bio-on show that they can be used successfully in the nanomedical field precisely because they are biocompatible and safe for human health.

The global market in contrast media alone is dominated by four multinational companies that generated an overall turnover of \$4.3 billion in 2015. According to the most recent estimates, this figure should rise to \$6 billion with 39.5% growth over the next 5 years.

APPLICATIONS

By synthesising inorganic and organic systems, Bio-On enters the field of biomedical and pharmaceutical applications through **Theranostics - a term coined by blending Therapeutic and Diagnostic.**

minervBiomedS innovative nanocapsules have a diameter of 80-100 nanometres and are made of polyhydroxyalkanoates.

Therapeutic applications in oncology are in the controlled and targeted release of cancer drugs. Thanks to its being biocompatible and the possibility of directing the nanocapsules, minerv-BiomedS product can be delivered selectively to the diseased area, releasing the drug where it is needed and eliminating side effects on healthy cells.

minervBiomedS nanocapsules can also incorporate smaller biocompatible metallic nanoparticles (20-30 nanometres) for diagnostic applications (imaging).

These metallic nanoparticles can be spherical or cylindrical and are iron oxide nanoparticles with magnetic properties and gold nanocylinders with plasmonic properties, respectively.

With the magnetic nanoparticles, we can perform Nuclear Magnetic Resonance using a product safe for human health and view diseased areas effectively.

With plasmonic gold nanocylinders, the innovative photoacoustic technique will enable high resolution images of diseased areas with greater sensitivity than conventional techniques.

Photoacoustic imaging is an innovative, revolutionary and non-invasive technique that produces extremely high resolution images of diseased areas of the body. It is highly precise and very safe because it does not use ionising radiation.

Advanced and exclusive research laboratories

Bio-on spa laboratories are housed at Bio-on Plants. These laboratories are a highly advanced research environment for the development of the fermentation technology and the development of the use of PHAs in the most advanced sectors.

COSMETICS - THERANOSTICS

Equipped with fume cupboards for chemical synthesis and separation with workbenches under suction hoods. Laboratories equipped with glassware for reactions, separations and preparation of

samples for chemical analysis.

The laboratories also have instruments for blending, discontinuous and continuous batch formulations, ovens, centrifuges, instruments for analysing chemical structure, molecular weight and more. The cellular biology part of the lab is particularly useful for the cosmetics field, since it includes extractor fans plus bench-top sterile hoods with UV lamp.

MICROBIOLOGICAL LABORATORIES

The microbiological laboratories are equipped with the best instruments for the following tasks:

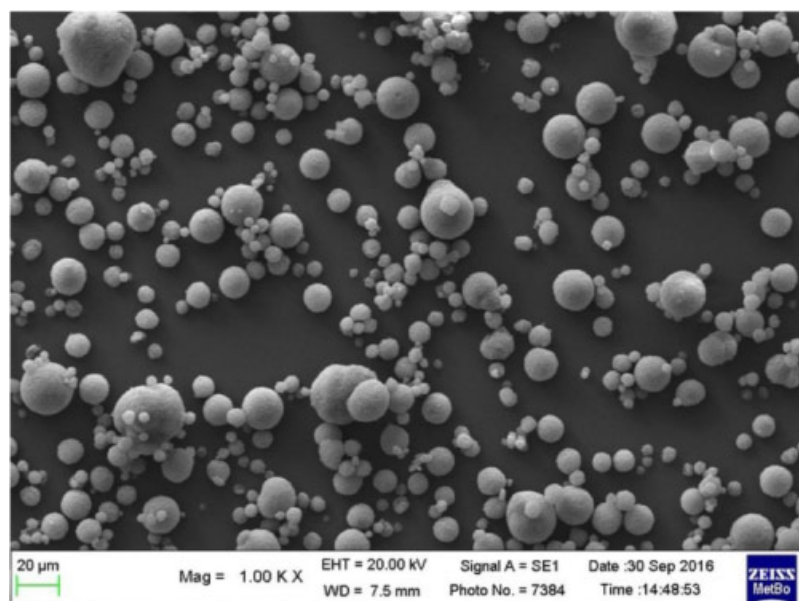
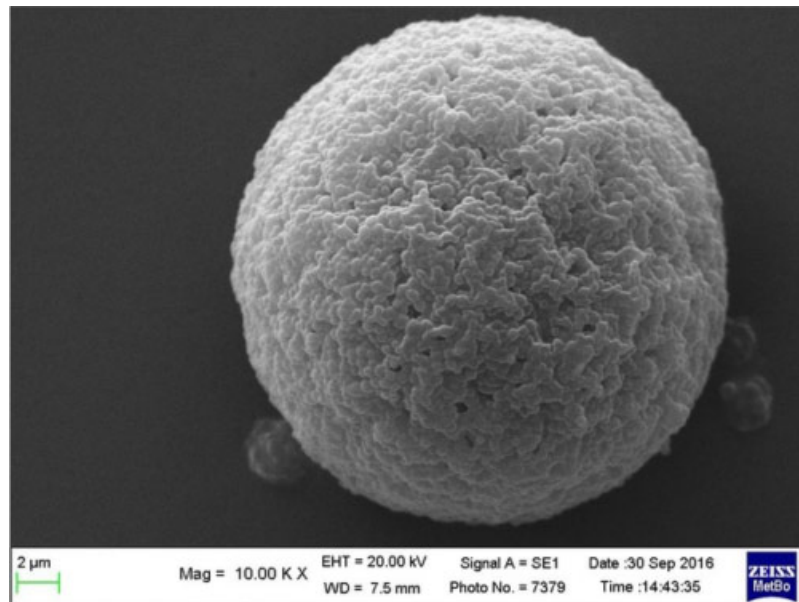
- cell clone improvement;
- providing production facility for aseptic inocula in the right physiological phase;
- studying various types of feedstock tested for PHA production.

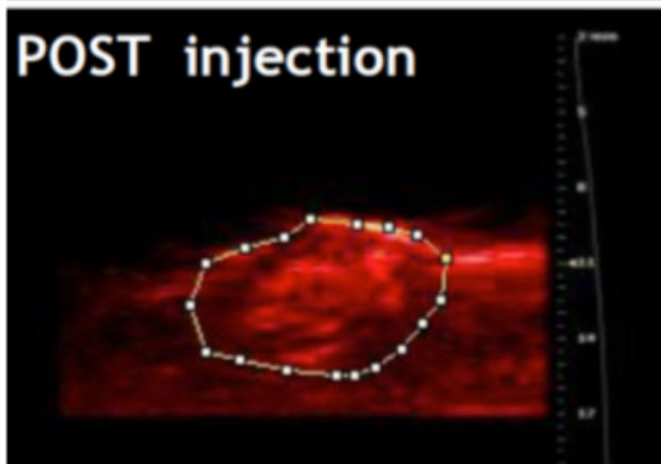
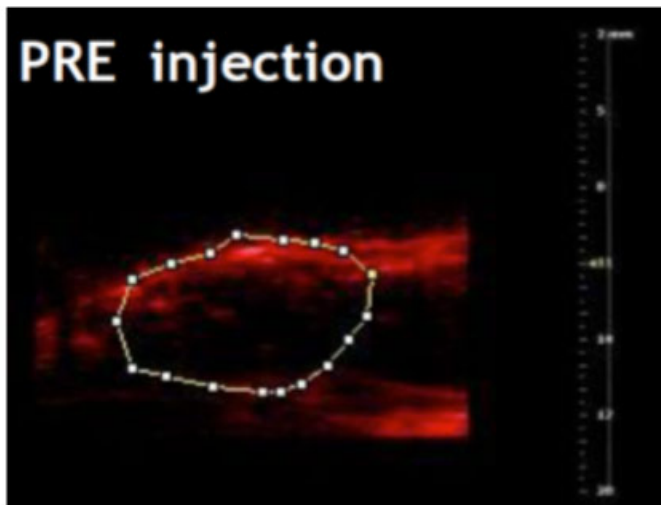
Particular attention is paid to aseptic conditions in cell preparation.

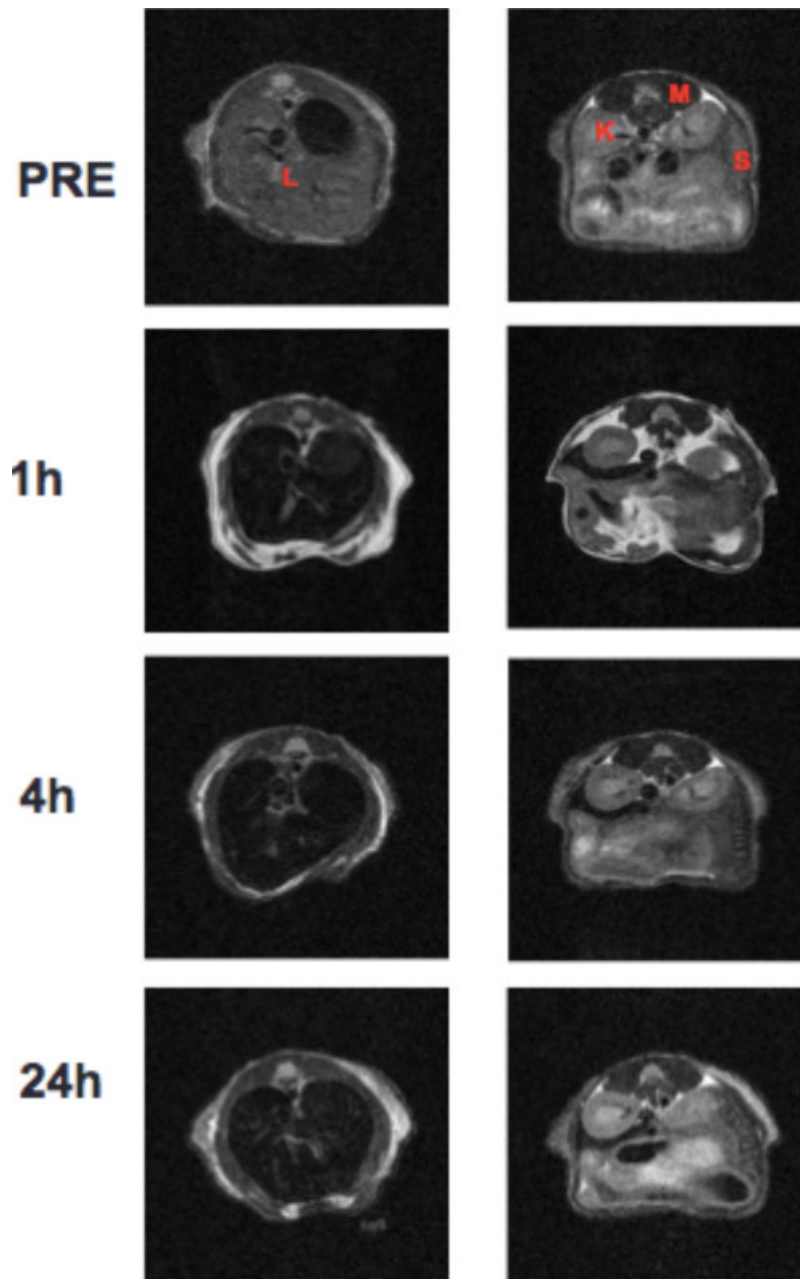
ANALYSIS LABORATORIES

The analysis laboratories perform a variety of functions:

- checking the nature of feedstock;
- supporting management of the fermentation process;
- determining production yields.







Fermentation and recovery plants

The best technologies associated with fermentation processes are implemented on fermenters of different sizes and scales:

- laboratory
- pilot
- industrial for special products in 2018.

Mechanical and electronic parts are subjected to a constant process of improvement in line with the relative technological advances.

Particular attention is paid to fermentation data detection systems and to process parameter management systems.

PHAs research and development

PHA production from carbon dioxide

Bio-on is working on a processor to produce PHA from CO₂.

The microorganism *Cupravidus necator* has the metabolic ability to accumulate PHA by metabolising CO₂ and converting it to PHA, with an autotrophic process. The heterotrophic process

from agricultural by-products also basically derives from CO₂, captured by cultures through photosynthesis and transformed into carbohydrates and lipids in plants.

During the industrial transformation phase, the sugars and fats then go into the transformation by-products, such as molasses and raw glycerides, an optimal carbon source for fermentation processes.

The process of transforming CO₂ is part of a new scientific and technological frontier that aims to capture CO₂ directly from the air, with a dual advantage:

zero cost carbon source

less CO₂ in the air, with a consequent reduction of the greenhouse effect.

The selection and improvement of the microorganism used for synthesis is being continuously developed.

An advanced engineering phase is currently setting out the characteristics of fermenters suitable for this cutting-edge technology.

The main obstacle to overcome in the industrial application of this new scientific and technological frontier is capturing from the air and the low solubility of CO₂ in water.

Research and development is advancing on three fronts:

microbiological;

processes;

engineering;

bearing in mind the need for a solid industrial-scale application to be implemented.

"TAILOR-MADE" special PHAs production

Bio-on makes characterisation grades on request.

Our laboratories choose a well-known, landmark plastic product to be characterised based on Bio-on's technology for producing PHAs from sugar beet or sugar cane by-products or other agricultural waste.

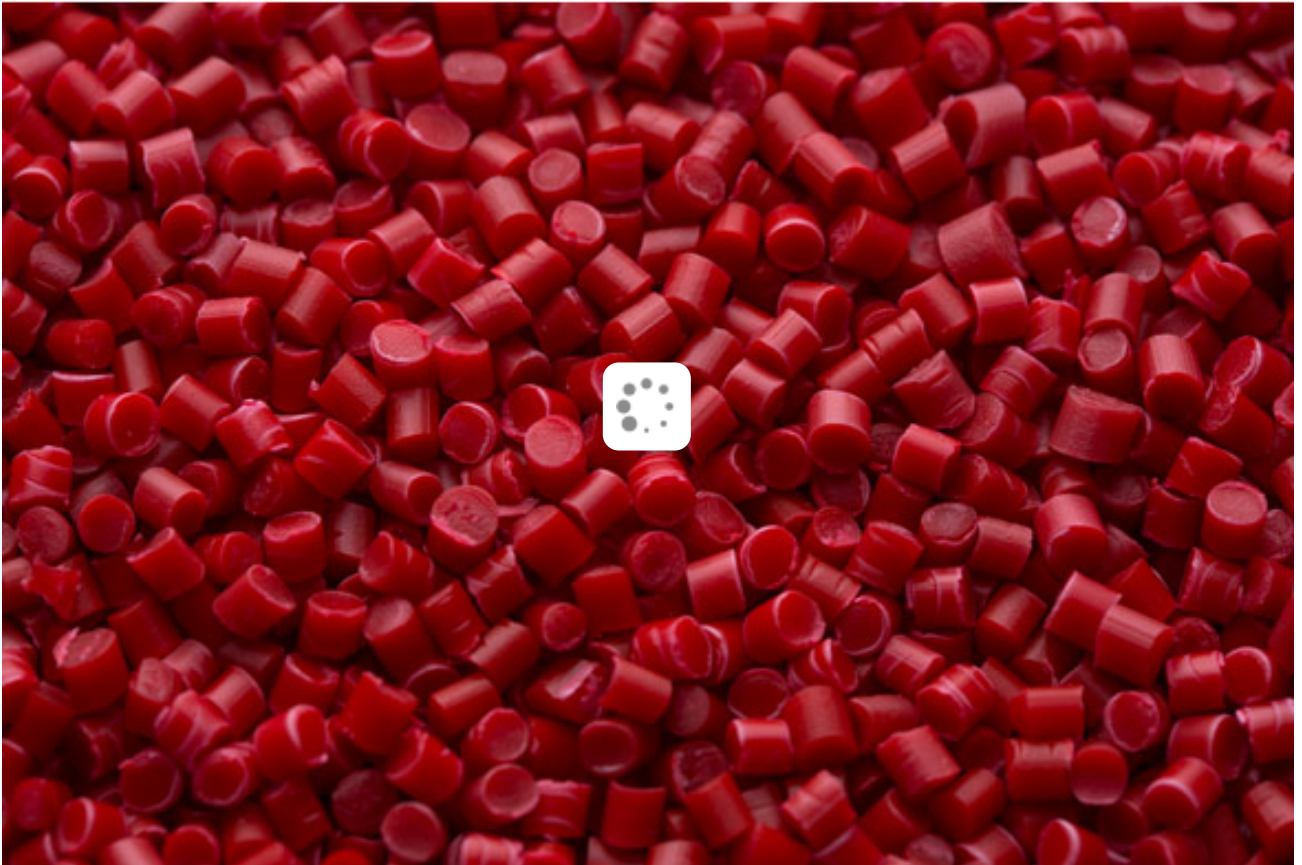
List of noted macro product families* with technical characteristics replicable by PHAs:

- Low density polyethylene (LDPE)
- High density polyethylene (HDPE)
- Polypropylene (PP)
- Polyvinyl chloride (PVC)
- Polystyrene (PS)
- Polyethylene (PE)
- Polyethylene terephthalate (PET)
- PMMA

(*)having identified the PHAs corresponding to the macro-family (e.g.: PP polypropylene), the individual grade is identified and characterised for a very precise use in terms of mechanical, physical and thermal properties.

For further technical information, contact us at: info@bio-on.it









Essential data on "basic" PHAs production

The technology for producing Minerv-PHATM features successive sub-processes that transform the carbon sources contained in agricultural co-products into polyhydroxyalkanoates (PHA), whose properties enable them to replace many oil-based plastics at commercial level, with the added benefit of being biodegradable.

Polyhydroxyalkanoates (PHA) are biodegradable in compost, like other bioplastics, but also in free water, a unique property thanks to the metabolic activity of microorganisms naturally present in the environment.

The Bio-on technology was designed to respect and enhance the eco and bio character of polyhydroxyalkanoates (PHA).

There are two platform products:

PHB, poly-hydroxy-butirate

PHBVV, poly-hydroxy-butirate-valerate-valerate.

The two platform products have considerably different characteristics, enabling different types of non-biodegradable, conventional plastics to be replaced.

Indeed, PHBVV is a family of molecules the flexible nature of which enables it to be modulated according to the needs of the application.

The sub-processes are as follows:

a) fermentation or up-stream-phase;

b) recovery and purification or down-stream-phase;

a) Fermentation.

The fermentation process is based on the ability of a bacteria microorganism, of the *Ralstonia eutropha* species, to metabolise carbon sources and convert them to polyhydroxyalkanoates (PHA).

The carbon sources used have variable origins, e.g.:

beet industry by-products

cane industry by-products

glycerol, including crude glycerol such as the biodiesel by-product oils and fats of various origin.

An initial growth phase encourages the proliferation of the bacterial biomass, i.e. the number of cells present; to this end:

- we provide all the necessary nutritional elements in order for the microorganism to duplicate as quickly as possible;
- we adjust process parameters (pH, temperature and pressure) to create the optimum environment for rapid growth;
- we have several vegetative propagation phases in a series of fermenters, in order to separate productive time from the dead time of biomass growth.

The microorganism used is entirely free from pathogenicity, so there is no danger to operators or those living in the surrounding area.

The essential purposes of the fermentation process are:

- to obtain a high conversion yield from substrate to polyhydroxyalkanoate (PHA);
- to obtain a high yield of polyhydroxyalkanoate (PHA) per unit of time.

As indicated previously, some production phases have no impact from a time point of view on hourly output, because they occur in parallel with the actual fermentation phase; these ancillary phases are:

- growth of the microorganism in the vegetative phases, in conical flask and fermenter;
- sterilising treatment of the co-product containing the carbon source;
- preparation of feed solutions integrating the bacterial metabolism.

At the end of the fermentation process, the resulting broth rich in polyhydroxyalkanoate (PHA) is rapidly removed in order to wash and sterilise the fermenter in preparation for the subsequent inoculum.

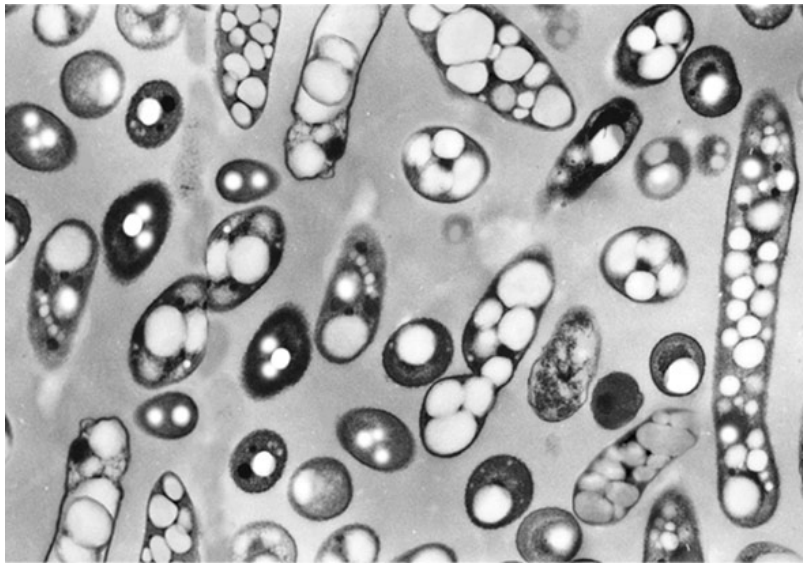
b) Recovery and purification.

The purpose of the recovery phase (the term used in the fermentation field) is to extract the molecule in question from the fermented broth mixture, until obtaining a highly pure raw product, which is refined to commercial level in the purification phase.

At the end of the process, the microorganism has a biomass of up to 80% polyhydroxyalkanoate (PHA), contained within the cell wall; therefore the purification phase of Bio-on's technology is part of the family of processes in which extraction involves the destruction of the cell. 20% of the biomass is made up of protein, plasma membranes, cytoplasmic content in general, treatable with standard purification processes.

The merit of Bio-on's technology, designed firstly to maintain its eco credentials and secondly to guarantee the biobased origin of the carbon contained and used in the process, lies in the fact that it does not use organic solvents such as chloroform or acetone, which have a negative impact on the environmental and are a high-cost part of the production process.





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