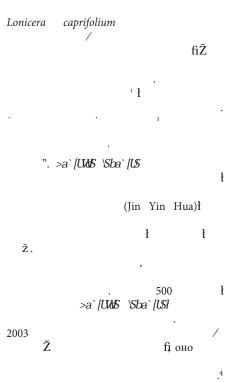
# Экстракт жимолости как альтернатива традиционным консервантам

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сегодняшний день консервирования косметических средств стал одним из главных в парфюмернокосметической отрасли, пробуждая сильный интерес к альтернативным системам консервирования, обладающим спектором активности, широким дерматологически и токсикологически безопасным и имеющим "благозвучные" названия по например, ботанические названия. Нечасто все вышеперечисленное сочетается вместе в одной системе. Редким исключением являются экстракты цвет ов жимолости из растений Lonicera japonica и Lonicera caprifolium в составе продукта Plantservative WSr от Campo Research.

В данной статье детально рассматрена данная комбинация экстрактов, включая ее происхождение, химию, применение, антибактериальные свойства, безопасность устойчивость, а также содержание (или же их отсутствие) парабенов и формальдегида в составе.



овременные исследования подтверждают невероятный потенциал Lonicera japonica, демонстрируя его антибактериальные, противовоспалительные и противовирусные свойства.



Only a couple of years ago a team of Chinese researchers identified the Lonicera japonica actives capable of targeting various influenza viruses directly, suggesting it to be the first natural product to directly target influenza A viruses. This makes the Lonicera japonica extract the first natural virological penicillin, with potential activity against other types of viruses as well. Lonicera japonica is a great example of the untapped potential in plant phytochemistry when scientific research is

## Chemistry

Given Lonicera japonica's interesting medicinal properties, there is a great deal of literature on this plant's phytochemistry. In fact Lonicera japonica has been found to contain quite a few notable substances thirty two organic acids (none of which is para-hydroxy benzoic acid), thirty flavones (flavonoids based on 2-phenylchromen-4 one (2-phenyl-1-benzopyran-4 one), thirty eight iridoids, (i.e. monoterpenes based on cyclopentanopyran), twenty eight saponins and nineteen other compounds 4

The ethanolic extracts were found to be rich in phenolic compounds, such as gallic acid, with interesting antioxidant as well as anti tyronase activity.  $^6$ 

The volatile components of both loniceras flowers obtained via hydrodistillation were analysed and found to contain fifty seven and thirty eight compounds respectively. Farnesol and germacrene D were the molecules present at the highest concentration in both Loniceras <sup>7</sup>

The above composition also applies to extracts obtained via traditional physical extraction methods. Interesting properties can be achieved by applying green chemistry techniques to the loniceras, where internal molecular rearrangement takes place generating newmolecular structures such as the Lonicerin complex. Lonicerin is a flavonoid normally present in the plant. This complexity is key in this loniceras extract combination, making it different from other conventional extracts obtained from the same plant. The lonicerin complex is suitable for cosmetic applications, being highly conditioning for the skin, with the additional benefit of being a broad spectrum antimicrobial agent.

The manufacturer only discloses part of the lonicerin complex molecular structure, to protect its proprietary intellectual property. The same applies to the green chemistry manufacturing method. Because of the lack of the full structure disclosure there were many speculations as to whether this extract contains formaldehyde. parabens or p-hydroxybenzoic acid, however these questions were not backed by any scientific evidence, and indeed the speculation seems misplaced and not based on any relatable experience or indeed expertise. This aspect will be covered later on in the article.

# Applications, performance and regulatory status

The extract of Lonicera japonica and caprifolium based on the Ionicerin complex is water soluble, pH independent, nearly colourless and with a faint odour, making it suitable for all sorts of cosmetic applications, from skin care to personal care, and it can be used as an alternative preservative system. It is compatible with the most cosmetic ingredients (including cationic and anionic surfactants), with the exception of citral, with which it reacts giving a slight pink colour, without affecting its antimicrobial properties.

The extract does not affect surface tension and therefore it does not impact an emulsion's stability or a surfactant's viscosity. Being pH independent it can provide antimicrobial protection in a wide array of pHs and in pH shift scenarios, a rare property in alternative preservative systems. It can be used by itself as a broad spectrum alternative preservative system at a concentration of 0.5%-1% depending on



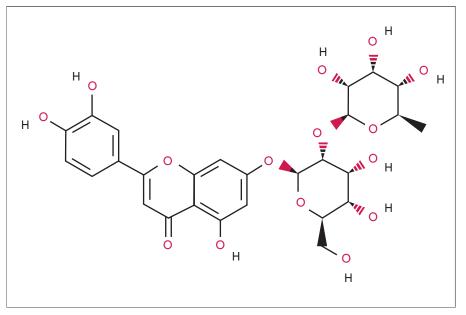


Figure 1: Multi-layered complexes of aqueous vegetal extractum of Lonicera caprifolium and L. japonica (only partial structure(s) of the multi-layered complexes of Lonicera caprifolium and Lonicera japonica shown here).

the formulation, without affecting product stability.

The extract also possesses low volatility, making it a reliable presence in formulations stored in open containers such as jars. It can also be used at a lower concentration, as a preservative booster or in combination with other alternative preservative systems. Papageorgiou et al<sup>8</sup> tested the lonicera extracts at 0.2% with 1% glyceryl caprylate, passing the EP challenge test in a couple of rinse-off formulations, but it failed in emulsion systems against A. niger, indicating this combination is very much formulation dependant. To boost the killing rate toward A. niger, the author suggests 0.1% anisic

The loniceras extract is also registered in index number. In addition each lonicera has its own CTFA monograph, and it is exempt from California proposition 65 making it globally compliant.

With regards to Japan where formaldehyde is banned, imported cosmetics undergo formaldehyde screening via a colorimetric method involving Schiff reagents. Because of the incompatibility of the loniceras with this test, it is advisable to have the loniceras extract tested using the manufacturer's recommended methods i.e. the AOAC 931.08 or the AOAC 964.21 used to test formaldehyde in maple syrup. If the colorimetric method screening test is used, it will give a false positive test result.

The false positive can also be verified by a dermatological test performed on a product containing 3.5% of the loniceras extract. The test was based on a single application with an occlusive patch test for 24 hours on a panel of 20 volunteers. The final report conclusion was 'non-irritating on intact human skin'. If formaldehyde was present in the loniceras extracts then this result would not have been possible being formaldehyde a known skin irritant and sensitiser. This is also confirmed by the in vitro tests performed on the raw material.

## Safety data

The Ioniceras extract manufacturer commissioned an in vitro toxicity test using Matrex, a reconstituted skin model to evaluate irritancy potential. The test was performed using propylene glycol and morpholine as comparison controls, propylene glycol as a non-irritating one and morpholine as a moderately irritating one. The results showed the loniceras extract has no irritation potential, even less than propylene glycol.

The loniceras extract was also tested for

mutagenicity according to the bacterial mutation Ames test (OECD 471) and the conclusion was that it did not induce mutations. This test is used to screen for mutagenic and potential carcinogenic hazards.

Other *in vitro* tests were performed using reconstructed human epidermis, such as skin corrosivity following the OECD testing guidelines, where the conclusion was that it was a non-corrosive material. Acute eye irritation potential was also evaluated in vitro reconstituted human corneal epithelium and again the result was that it was non-irritant.

# Parabens and formaldehyde content

A lot of opinions have been shared on the internet in the last ten years regarding the loniceras extract, some claiming that it contains p-hydroxybenzoic acid, a paraben precursor, and formaldehyde.

The first claim was based on an another raw material that did contain phydroxybenzoic acid from the same manufacturer - not currently used in cosmetics, but still used as an industrial grade water sanitiser (especially for Legionnaires' disease, *Legionella spp.*) for air-conditioning cooling towers. The material is also used in sanitising chicken abattoirs and poultry breeding farms from *Campylobacter jejuni*.

It is important to notice that in the articles found on the *Lonicera japonica* phytochemistry no *p*-hydroxybenzoic acid was mentioned, therefore not even the original plant seems to contain the paraben precursor.

The second claim regarding formaldehyde being present in the loniceras extract is due to the incompatibility of the loniceras extract with Schiff reagents. The Schiff test was originally developed in the 1800s as a reaction to detect organic aldehydes. Aliphatic aldehydes and aldose sugars react with a solution of rosaline decolourised with sulphurous acid (dye formulation and Schiff reagent). If aldehydes are present a magenta colour appears due to the formation of a chromophore compound produced by the chemical combination with the aldehyde. Therefore, if a sugar has an aldehyde group, like for example non adulterated maple syrup, and gets tested using a method where Schiff reagents or pseudo-Schiff reagents are used, a false positive for formaldehyde will appear. The same will happen if an aldehyde group is present in a fragrance compound in a finished product. The recommended method by the manufacturer, the AOAC 964.21, avoids false positives by distilling the formaldehyde from the sample and taking a spectrophotometric reading at 415



nm of the distillate, giving a quantitative result. The test was developed in the 1960s to detect formaldehyde in maple syrup to avoid the false positives given by other aliphatic aldehydes or aldose sugars. Not many laboratories perform the AOAC tests mentioned, the one found is TUV Sud PSB Singapore and Chemservice Srl in Milan, Italy. The manufacturer recommends the use of AOAC accredited laboratories that can be found on the AOAC International member directory.

# Sustainability

The loniceras extract comes from wildly cropped plants in East Asia, therefore it is very much a sustainable, product, coming from non-intensive cultivation and with no forest destruction involved.

The extract does not have organic certification, however it complies with the standards of several green certification bodies, this includes Cosmos (Organic and Natural), Ecocert, BDIH, ICEA and Soil Association, via the relevant raw material forms.

# Заключение

The Plantservative Wsr is a successful example of green chemistry and the great potential present in every plant, waiting to be tapped into with modern green chemistry techniques. Produced from wildly cropped Japanese honeysuckle and Italian honeysuckle, without the use of any synthetic substances, it contains novel phytochemicals based on lonicerin complex with broad spectrum antimicrobial properties for cosmetic applications.

The Plantservative WSr has an excellent dermatological profile (*in vitro* and *in vivo*) and broad antibacterial activity in several cosmetic applications, including organic

certified products. Given its novel molecular structure it is not compatible with Schiff reagents and modified Schiff reagents, therefore the manufacturer recommends particular methods need to be followed for the formaldehyde screening test i.e. AOAC 931.08 or AOAC 964.21. The full molecular structure and manufacturing method are protected by the manufacturer's intellectual property rights, however the above tests prove the authenticity of this green chemistry extract.

# Библиографические ссылки

- 1 Ghirardini MP, Carli M, Del Vecchio N, et al. The importance of a taste. A comparative study on wild food plant consumptions in twenty-one local communities in Italy. *Journal of Ethnobiology and Ethnomedicine* 2007; 3:22, DOI 10.1186/1746-4269-3-22
- 2 Grieve M. A modern herbal, 1971; London, Dolver Publications
- 3 Culpeper's complete herbal, 1995; pg 273, Wordsworth Reference
- 4 Shang X, Pan H, Li M, Miao X, Ding H. Lonicera japonica Thunb: ethnopharmacology, phytochemistry and pharmacology of an important traditional Chinese medicine, *Journal of Ethnopharmacology* 2011; 138:1-21
- 5 Zhou Z, Li X, Liu J, et al. Honeysuckle-encoded atypical microRNA2911 directly targets influenza A viruses. Cell Research 2015; 25:39-49. 6
- Dung NT, Bajpai VK, Rahman A. Phenolic contents, antioxidant and tyronase inhibitory activities of Lonicera Japonica Thumb, *Journal of Food Biochemistry* 2011; 35 (1):148-160
- 7 Ilies DC, Radulescu V, Dtu L. Volatile constituents from the flowers of two species of Honeysuckle (Lonicera Japonica and Lonicera Caprifolium), Farmacia 2014; 62 (1):194-201
- 8 Papageorgiou S, Varvaresou A, Tsirivas E,
  Demetzos C. New alternatives to cosmetics
  preservation, *J. Cosmet.* Sci 2010; 61:107-123