Wacker Chemie AG Hanns-Seidel-Platz 4 81737 München, Germany www.wacker.com

# FEATURE SERVICE

Number 02, October 2014

Fitness from Within: Nature-Identical Hydroxytyrosol

Fresh fish, lots of fruits and vegetables, a serving of walnuts every day and, most importantly, olive oil. People who regularly put these foods on their menu are following what is known as the Mediterranean diet – and making very healthy eating choices. What makes this a healthy diet is that these foods contain plant secondary metabolites, a category of substances that includes hydroxytyrosol. This antioxidant is much sought after as an active agent, and WACKER can now produce a nature-identical version of it via a new, patented synthesis route. The process improves purity with no unwanted byproducts, while keeping the content of the active agent at a defined level – opening up entirely new possibilities for manufacturers of nutritional supplements and cosmetics.

> Thinly sliced carrots combined with layers of bell peppers and tomatoes, seasoned with olive oil and grated Parmesan – while certainly delicious, a colorful vegetable carpaccio like this also serves up a course of important vitamins, minerals and fiber. That eating an abundance of fruits and vegetables keeps mind and body in good shape is fairly well known, as is the promise of longevity that the Mediterranean diet offers. A large number of studies have put Southern European eating habits under the microscope and investigated physical health within these populations. What they have concluded is that the Mediterranean diet has a positive effect on our bodies – an effect due to the com-

Healthy and fit with the Mediterranean diet

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	pounds these foods contain. Nutrition experts have been paying
	particularly close attention to the plant secondary metabolites in
Plant secondary metab-	bell peppers, olives and the like, owing to the health benefits of
olites ward off disease	these chemicals. Examples of what these "natural wonders" are
	supposed to do include protecting against cardiovascular diseas-
	es, diabetes and Alzheimer's, strengthening immune system re-
	sponse, and inhibiting inflammatory processes.
	Plant secondary metabolites – polyphenols, to be precise – in-
	clude hydroxytyrosol, a compound that is among the most power-
Hydroxytyrosol is a	ful antioxidants, protecting human cells from harmful oxygen rad-
powerful antioxidant	icals. Olives and olive leaves are the most common naturally oc-
	curring source of this substance. "There are already a few sup-
	pliers out there who obtain this active agent via extraction," ex-
	plains Dr. Sebastian Schuck, senior manager for business devel-
but is expensive in its natural form and difficult to produce	opment at WACKER BIOSOLUTIONS, "and they sell it to manu-
	facturers of nutritional supplements and cosmetics who use it in
	their product formulations." Their olive extract, however, is a
	blend of a wide range of polyphenols rather than pure hydroxyty-
	rosol. Olive harvests are also subject to seasonal fluctuations,
	which affects more than just the composition of the product in-
	gredients – it also has an impact on the price of these valuable
	extracts.
	Nevertheless, demand for this health superstar is on the rise,
The solution: chemical synthesis	which gave WACKER experts an idea: "We follow trends on the
	nutritional-supplement market very closely, of course. That's
	what got us to thinking about developing a synthesis route for
	making hydroxytyrosol – a method that could supply the market

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An efficient process	with this coveted active in large enough amounts and at high
	levels of purity," Schuck recalls. No sooner said than done. With-
	in just a year and a half, WACKER researchers had managed to
	prepare a nature-identical hydroxytyrosol – a feat that included
	selecting the most suitable synthesis route, scaling up the pro-
	cess and ultimately submitting a successful patent application.
	That these chemists were able to develop a route in such a short
	period of time was primarily due to the enormous treasure trove
	of experience that has been gained over a great many decades
	in the laboratories at the main plant in Burghausen. And thanks
	to the new multistage synthesis method, the company now has a
	cost-effective process for manufacturing a highly pure form of the
	active. Because a great deal of expertise has gone into the pro-
	cess, Schuck can only reveal so much: "There are a lot of ways
independent of har-	of synthesizing a natural substance – but, as far as we're con-
vests and weather	cerned, our researchers found the best, most efficient method
yielding high purity and quality	currently available," says the WACKER expert. Another ad-
	vantage is that the route is based on commercially available raw
	materials and practical process conditions, ensuring that the
	compound can be produced on a large scale.
	"Plus, we don't end up with a mixture of natural substances any
	more – now we can directly produce metric tons of the molecule
	we want, without having to rely on harvests. Our hydroxytyrosol
	meets a precise set of specifications, and the quality is consistent
	and exceptionally high," Schuck is pleased to report. And that
	opens up the possibility of other applications: the active is highly
	concentrated, making it suitable, for instance, as a nutritional
	supplement in more compact forms, such as tablets and cap-

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	sules. In addition, WACKER also hopes that nature-identical hy-
	droxytyrosol – marketed under the name $HTEssence^{^{ ext{B}}}$ – will be
	more widely accepted among food manufacturers. "Cost pres-
	sures are especially high in that industry – and HTEssence $^{ extsf{@}}$
	means we can offer these manufacturers a cost-effective,
The global market for	high-quality alternative," Schuck points out. The food industry is
nutritional supplements	the most important market for WACKER BIOSOLUTIONS, con-
is growing	stituting 61 percent of the division's sales. Experts estimate glob-
	al sales of food supplements to be around 96 billion US dollar, a
	figure that rises to 112 billion US dollar when functional foods are
	taken into account. It is a promising market – and it is precisely
	the market that WACKER's new, nature-identical hydroxytyrosol
	is intended to serve and, in so doing, expand the company's
	functional ingredients business.
	Its unusually powerful antioxidant properties are what make hy-
Hydroxytyrosol protects	droxytyrosol such an interesting plant secondary metabolite for
cells from free	use in foods and cosmetics. The compound is exceptionally good
radicals	at capturing free radicals, which can damage our cells. At 45,000
	micromol Trolox equivalents <sup>1</sup> per gram, the oxygen radical ab-
	sorbance capacity of hydroxytyrosol is nearly ten times that of
	green tea and over twice as high as that of coenzyme Q10. As
	Schuck notes, "That means that hydroxytyrosol protects human
which heads off cardi-	cells and blood lipids from oxidative stress – a key factor influ-
ovascular diseases	encing the development of cardiovascular diseases."
	Also attributed to this active substance is the ability to strengthen

<sup>&</sup>lt;sup>1</sup> Trolox is a water-soluble vitamin E derivative with a pronounced antioxidant effect, prompting its use as a reference substance for determining the antioxidant capacity of a sample.

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	immune system response, lower blood pressure, inhibit inflam- matory processes, and exert a positive effect on the bones and joints – interesting properties that could be put to use in nutrition- al supplements or applications in sports nutrition, such as power bars or functional beverages.
Interest is high – also in the cosmetics industry Suppressing dark pig- ments	The cosmetics industry, however, discovered hydroxytyrosol some time ago as a useful additive in products such as anti-aging formulations. "Hydroxytyrosol affects melanin production in hu- man cells, suppressing dark pigmentation and making the skin lighter and more uniform. Antioxidants are also used for protect- ing the skin, where they counteract the aging process," Schuck explains.
makes the skin lighter and more uniform No impurities, no by- products	Light skin is particularly desirable for many people in Asia, which is why so many products there advertise a lightening effect. Pharmacies already carry a number of creams and lotions utiliz- ing the active substance in olives in order to lighten the skin. In- dustry experts expect the market for these skin-lightening prod- ucts to grow to some 20 billion US dollar by 2018. "Our HTEssence <sup>®</sup> is the perfect product for this market, since it doesn't contain any undesirable byproducts or impurities at all," Schuck observes. The substance will be sold as an odorless, water-soluble powder, although it will also be available in liquid form.
Positive customer feed- back	WACKER anticipates approval for HTEssence <sup>®</sup> by the end of the year. The company has already sent out samples of the product to a few customers for research and development purposes – and initial feedback on nature-identical hydroxytyrosol has put

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Schuck and his team in a very good mood.

### Background Information on Hydroxytyrosol:

A compound found in olives and olive leaves, hydroxytyrosol is a plant secondary metabolite – a phenol to be precise. Phenols and polyphenols are aromatic compounds present as bioactive substances in plants, where they produce color or flavor, or act as an antioxidant. Their considerable antioxidant power makes them interesting for the food and cosmetics industries: as free-radical scavengers, antioxidants prevent oxidative stress, which is associated with a number of diseases and signs of aging.

Antioxidants are present in many plants, but not all of these exhibit the same levels of activity – this is why hydroxytyrosol is considered so important: Olive oil polyphenols, which include hydroxytyrosol, are said to contribute to the protection of blood lipids from oxidative stress – a factor often linked to human aging and many diseases, including cancers or cardiovascular diseases.

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A WACKER employee is filling pure, nature-identical hydroxytyrosol dissolved in water in a spray dryer. Thanks to the controlled production process, HTEssence<sup>®</sup> is free of allergens and pesticides and does not contain any unwanted byproducts or contaminants (photo: Wacker Chemie AG).



Spray-dried nature-identical hydroxytyrosol: The odorless, water-soluble powder HTEssence<sup>®</sup> is available in great purity and with a defined amount of active ingredient (photo: Wacker Chemie AG).

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Ideal for cosmetic products: Hydroxytyrosol is able to influence the melanin pigments in human cells, so that the skin becomes lighter and dark patches disappear. As a free radical scavenger it can further prevent wrinkle formation and skin aging (photo: Wacker Chemie AG).

<u>Note:</u> These photos are available for download at: <u>http://www.wacker.com/featureservice</u>

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#### The company in brief:

WACKER is a globally-active chemical company with some 16,000 employees and annual sales of around €4.48 billion (2013). WACKER has a global network of 25 production sites, 21 technical competence centers and 52 sales offices.

#### WACKER SILICONES

Silicone fluids, emulsions, rubber and resins; silanes; pyrogenic silicas; thermoplastic silicone elastomers

#### WACKER POLYMERS

Polyvinyl acetates and vinyl acetate copolymers in the form of dispersible polymer powders, dispersions, solid resins and solutions used as binders for construction chemicals, paints and coatings, adhesives, plasters, textiles and nonwovens, as well as for polymeric materials based on renewable resources

#### WACKER BIOSOLUTIONS

Biotech products such as cyclodextrins, cysteine and biologics, as well as fine chemicals and PVAc solid resins

#### WACKER POLYSILICON

Polysilicon for the semiconductor and photovoltaic industries

#### Siltronic

Hyperpure silicon wafers and monocrystals for semiconductor components