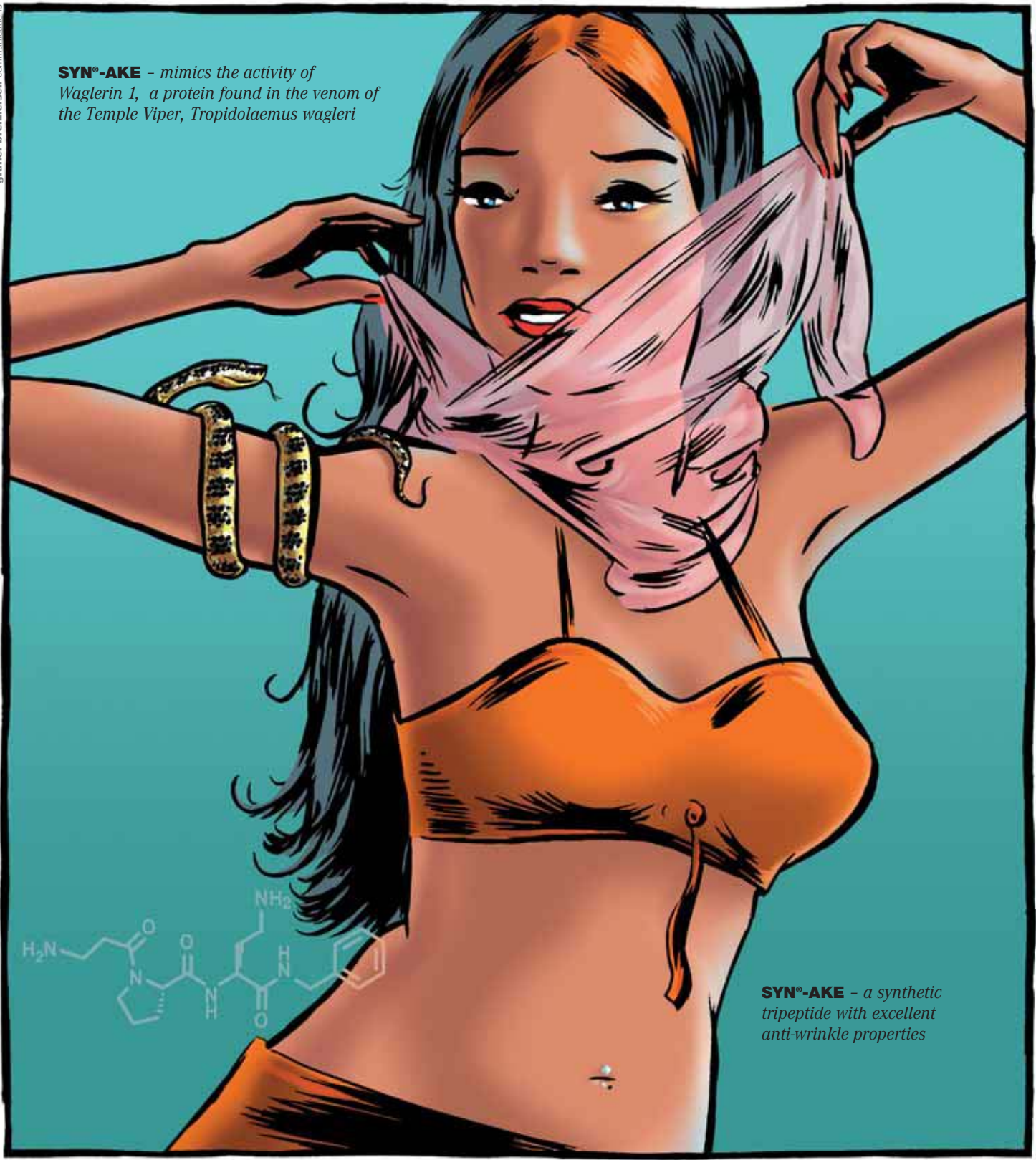


SYN[®]-AKE – Age Killing Effect.

SYN[®]-AKE – mimics the activity of
Waglerin 1, a protein found in the venom of
the Temple Viper, *Tropidolaemus wagleri*



SYN[®]-AKE – a synthetic
tripeptide with excellent
anti-wrinkle properties

**SWISS
TECHNOLOGY
AWARD**
2006

NA DISTRIBUTOR
Centerchem, Inc
20 Glover Avenue, Norwalk, CT 06850
Tel: 203-822-9800 * fax 203-822-9820
www.centerchem.com

pentapharm
benefiting society through science



SYN®-AKE

SYN®-AKE is a new anti-wrinkle active compound based on a synthetic tripeptide that mimics the effect of *Waglerin 1*, a peptide that is found in the venom of the Temple Viper, *Tropidolaemus wagleri*.

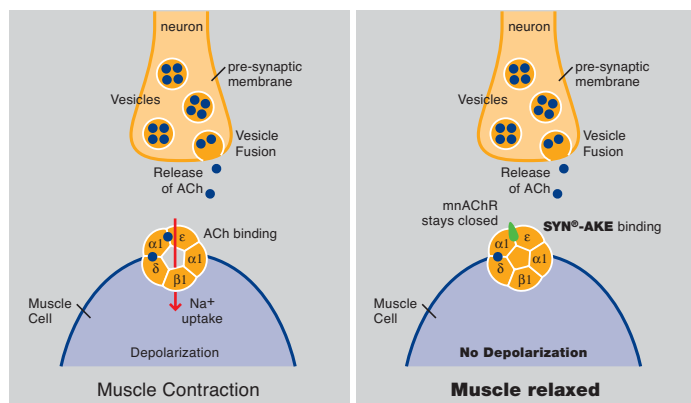
PENTAPHARM has developed, over the past 30 years, a unique approach for the breeding and housing of venomous snakes and particularly the Brazilian Lance Adder Bothrops moojeni whose venom is used for therapeutic (anticoagulants, haemostatics) and diagnostic products. Currently, some 10 000 specimens of this snake species are bred and housed at **PENTAPHARM DO BRASIL**, making **PENTAPHARM** the largest snake breeder and keeper in the world.

The long experience of **PENTAPHARM** in snake venom research has made possible the investigation of venom peptides for cosmetic applications. A special focus has been the investigation of the Temple Viper's venom.



SYN®-AKE acts in a manner similar to that of *Waglerin 1* which acts at the post-synaptic membrane. The peptide is an antagonist of the muscular nicotinic acetylcholine membrane's receptor (mnAChR). As the muscular nicotinic ACh receptors are blocked, the ion canal remains closed. There is no uptake of Na⁺ and the muscles stay relaxed.

Source: McArdle, JJ, T L Lentz, V Witzemann, H Schwarz, SA Weinstein & JJ Schmidt. 1999. *Waglerin-1 selectively blocks the epsilon form of the muscle nicotinic acetylcholine receptor. J. Pharmacol. Exp. Therap.* 289:543-550.



SYN®-AKE has been thoroughly tested and is considered to be appropriate for cosmetic applications.

Properties:

- Antagonist of the muscle nicotinic acetylcholine receptor (mnAChR)
- Blocks Na⁺ uptake at the post-synaptic membrane
- Inhibits muscle contractions

Function:

- **SYN®-AKE** is an excellent anti-wrinkle active compound with a snake venom-like mode of activity.
- **SYN®-AKE** smoothes mimic wrinkles in a short period.

Cosmetic applications:

- Age killing effect particularly effective against expression lines
- Intensive anti-wrinkles care

Suggested concentration:

1-4%

Formulation:

SYN®-AKE is a clear glycerine-based aqueous solution that can easily be incorporated into the aqueous phase of a formulation.

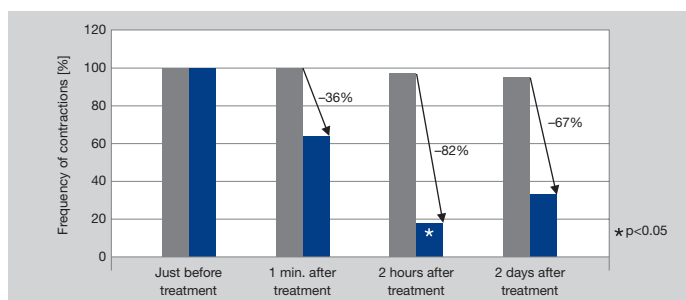
INCI Name:

Glycerine, Water, Dipeptide Diaminobutyroyl Benzylamide Diacetate

Efficacy Tests

in vitro tests:

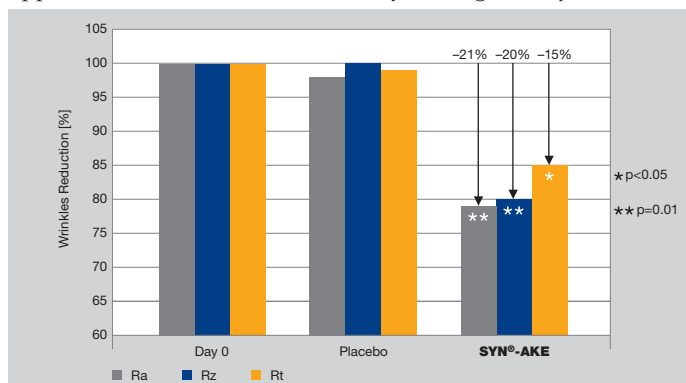
The efficacy of the **SYN®-AKE** tripeptide (at a concentration of 0.5mM) has been demonstrated *in vitro* by measuring the frequency of contraction of the innervated muscle cells as a function of the incubation time.



SYN®-AKE peptide reduces muscle cell contraction and its action is reversible.

in vivo tests:

The measurement of the smoothing and anti-wrinkle effect of **SYN®-AKE** (4%) was compared to a placebo. A cream was applied to the forehead twice daily during 28 days.



SYN®-AKE - Age Killing Effect.

The smoothing effect (Ra) was measured on 80% of the volunteers and the anti-wrinkle effect (Rz, Rt) measured on 73% of the volunteers.

Results showed up to -52% of wrinkle size after 28 days application!