

PENTAPHARM SERVICE CENTER COSMETICS

Formulation Guidelines for PEPHA[®]-TIGHT (Code no 800356)

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SUMMARY

PEPHA[®]-TIGHT was tested with different groups of raw materials like preservatives, hydrocolloids, sunscreens, detergents, commonly used water-soluble substances and other PENTAPHARM actives.

As PEPHA[®]-TIGHT is an opalescent to turbid liquid, it's usual to observe a slight turbidity or clouding in aqueous solutions containing this active. However these phenomena don't influence the activity of PEPHA[®]-TIGHT in a cosmetic formula.

INTRODUCTION

PEPHA[®]-TIGHT (INCI: Algae Extract, Pullulan) is a yellow to amber, opalescent to turbid, water-soluble, natural active ingredient in highly viscous liquid form. PEPHA[®]-TIGHT is a unique, biotechnologically produced, microalgae-based skin tightener: it tightens the skin immediately after application and at the same time provides a long-term firming effect by strengthening the skin's connective tissue.

PENTAPHARM has screened a wide range of common ingredients to test their compatibility with PEPHA[®]-TIGHT.

The following guidelines give an overview of those raw materials tested and found to be either compatible or incompatible with PEPHA[®]-TIGHT.

GENERAL RECOMMENDATIONS

- Dosage:** For skin care preparations, concentrations of 1 to 5% PEPHA[®]-TIGHT are recommended.
- Processing:** PEPHA[®]-TIGHT must be processed cold (<50°C) and is incorporated right at the end of a cosmetic formulation.
- pH:** The optimal pH range for the formulation with PEPHA[®]-TIGHT is between 5.0 and 7.0.
- Ethanol:** PEPHA[®]-TIGHT is compatible with ethanol at concentrations up to 10%.
- Storage:** between 4 and 8°C in the original sealed container, protected from light and in a clean place.



TEST PROCEDURE

The different raw materials were tested with 5% PEPHA[®]-TIGHT in aqueous solutions.

The raw materials to be tested were used in concentrations according to the manufacturer's indications. **Should these raw materials be used at other concentrations, we recommend to eventually re-test their stability with PEPHA[®]-TIGHT.**

The samples were stored at 4°C, room temperature and 40°C. For the evaluation of the stability, we focused on the results at room temperature and at 4°C, since PEPHA[®]-TIGHT itself is not stable at temperatures > 50°C.

The samples were judged after 1 month and again after 3 months according to the following criteria: pH value, odour, colour, appearance, viscosity.

PEPHA[®]-TIGHT is an opalescent to turbid viscous liquid. Slight turbidity or clouding in aqueous solutions is therefore not a sign of incompatibility for this product. That's why the results of these compatibility tests are split into three groups:

- **Recommended substances:** without or with very slight turbidity, stable pH value, no odour or colour development, stable viscosity.
- **Compatible substances:** with slight turbidity, stable pH value, no odour or colour development, stable viscosity.
- **Substances with limited compatibility:** high turbidity, and/or unstable pH value, and/or odour or colour development, and/or unstable viscosity.

However, if these substances with limited compatibility are used with PEPHA[®]-TIGHT in an emulsion or/and in lower concentration, probably no visible incompatibilities can be recognized.



PRESERVATIVES

RECOMMENDED PRESERVATIVES

Product Name / Supplier	% Wt.	INCI Name
Phenoxetol / Nipa	1.0	Phenoxyethanol
Nipaguard PDU / Nipa	0.5	Diazolidinyl Urea
Glydant 2000 / Lonza	0.4	DMDM Hydantoin
Glycacil 2000 / Lonza	0.8	Iodopropynyl Butylcarbamate
Potassium Sorbate / CFS-Schweizerhall	0.5	Potassium Sorbate
Benzyl Alcohol / VWR	1.0	Benzyl Alcohol
Benzoic Acid / VWR	0.2	Benzoic Acid
Geogard 221 / Lonza	0.8	Dehydroacetic Acid (and) Benzyl Alcohol

With PEPHA[®]-TIGHT, these preservatives formed stable aqueous solutions without or with very slight turbidity.

COMPATIBLE PRESERVATIVES

Product Name / Supplier	% Wt.	INCI Name	Observations
Nipastat Sodium / Nipa	0.1	Methylparaben, Butylparaben, Ethylparaben, Propylparaben	Slight clouding
Unicide U-13 / Lipo	0.6	Imidazolidinyl Urea	Slight pH decrease
Isocil PC / Lonza	0.1	Methylchloroisothiazolinone, Methylisothiazolinone	Discoloration at 40°C

PRESERVATIVES WITH LIMITED COMPATIBILITY

Product Name / Supplier	% Wt.	INCI Name	Observations
Spectradyne / Lonza	0.3	Chlorhexidine Digluconate	Precipitate
Irgasan DP300 / Ciba	0.3	Triclosan	Precipitate



HYDROCOLLOIDS

RECOMMENDED HYDROCOLLOIDS

Product Name / Supplier	% Wt.	INCI Name
Pemulen TR-1 / Noveon	1.0	Acrylates / C10-30 Alkyl Acrylate Crosspolymer
Carbopol ETD 2001 / Noveon	0.5	Carbomer
Natrosol 250 HHX / Hercules	2.5	Hydroxyethyl Cellulose
Sepigel 305 / Seppic	3.0	Polyacrylamide (and) C13-14 Isoparaffin (and) Laureth-7
Structure XL / National Starch	5.0	Hydroxypropyl Starch Phosphate

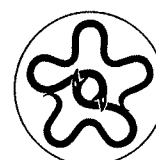
These hydrocolloids formed stable gels with PEPHA[®]-TIGHT.

COMPATIBLE HYDROCOLLOIDS

Product Name / Supplier	% Wt.	INCI Name	Observations
Keltrol RD / Kelco	3.0	Xanthan Gum	Slight pH decrease at 40°C

HYDROCOLLOIDS WITH LIMITED COMPATIBILITY

Product Name / Supplier	% Wt.	INCI Name	Observations
Stabileze QM / ISP	0.5	PVM/MA Decadiene Crosspolymer	Viscosity loss
Aristoflex AVC / Clariant	1.0	Ammonium Acryloyldimethyltaurate / VP Copolymer	Viscosity loss



UV FILTERS

RECOMMENDED UV FILTERS

Product Name / Supplier	% Wt.	INCI Name
Tinosorb M / Ciba	5.0	Methylene Bis-Benzotriazolyl Tetramethylbutylphenol
Tioveil AQ-P / Uniqema	5.0	Aqua (and) Titanium Dioxide (and) Alumina (and) Silica (and) Sodium Polyacrylate (and) Propylene Glycol

These UV filters formed stable solutions with PEPHA[®]-TIGHT.

UV FILTERS WITH LIMITED COMPATIBILITY

Product Name / Supplier	% Wt.	INCI Name	Observations
Parsol HS / DSM	5.0	Phenylbenzimidazole Sulfonic Acid	Precipitate at RT
Uvinul MS 40 / BASF	5.0	Benzophenone-4	Precipitate

DETERGENTS

RECOMMENDED DETERGENTS

Product Name / Supplier	% Wt.	INCI Name
Genapol LRO Liquid / Clariant	20.0	Sodium Laureth Sulfate
Tego Betain F50 / Degussa	20.0	Cocamidopropyl Betain
Rewopol SB FA 30 / Rewo	20.0	Disodium Laureth Sulfosuccinate
Rewoteric AM2CNM / Rewo	20.0	Disodium Cocoamphodiacetate

These detergents formed stable, clear solutions with PEPHA[®]-TIGHT.

COMPATIBLE DETERGENTS

Product Name / Supplier	% Wt.	INCI Name	Observations
Rewoteric AMCAS / Rewo	20.0	Cocamidopropyl Hydroxysultaine	Slight precipitate at RT
Plantacare 2000 UP / Cognis	20.0	Decyl Glucoside	Slight precipitate at 4°C



MISCELLANEOUS SUBSTANCES

RECOMMENDED MISCELLANEOUS SUBSTANCES

Product Name / Supplier	% Wt.	INCI Name
D-Panthenol 50P	2.0	Panthenol
EDTA	0.1	EDTA
Glycerin 86.5%	20.0	Glycerin
Propylene Glycol	20.0	Propylene Glycol
1,3-Butandiol	20.0	Butylene Glycol
Cremophor RH 410	5.0	PEG-40 Hydrogenated Castor Oil
Glycine	2.0	Glycine
L-Lysine	2.0	Lysine
L-Glutamic Acid	2.0	Glutamic Acid
Citric Acid	2.0	Citric acid
Glycolic Acid	2.0	Glycolic Acid

These miscellaneous substances formed stable, clear to very slightly turbid solutions with PEPHA[®]-TIGHT.

MISCELLANEOUS SUBSTANCES WITH LIMITED COMPATIBILITY

Product Name / Supplier	% Wt.	INCI Name	Observations
Sodium Metabisulfite	0.1	Sodium Metabisulfite	pH decrease at 40°C & slight clouding
EDG Plus	5.0	Ethoxydiglycol	pH decrease & discoloration at 40°C
Lactic Acid	2.0	Lactic Acid	pH decrease

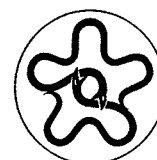


PENTAPHARM ACTIVES

RECOMMENDED PENTAPHARM ACTIVES

Product Name	Product Code	% Wt.	INCI Name
ALPAFLOR [®] EDELWEISS GC	900-01	5.0	Leontopodium Alpinum Extract
COLHIBIN	801057	5.0	Hydrolyzed Rice Protein
EASHAVE	316-01	5.0	Triticum Vulgare (Wheat) Germ Extract, Saccharomyces Cerevisiae Extract, Sodium Hyaluronate
ELHIBIN [®]	301-02	5.0	Glycine Soja (Soybean) Protein
HYASOL-BT	296-04	5.0	Sodium Hyaluronate
IMMUCELL [®]	302-01	5.0	Glycoproteins
IRICALMIN	316-02	5.0	Triticum Vulgare (Wheat) Germ Extract, Saccharomyces Cerevisiae Extract, Sodium Hyaluronate
MELAWHITE [®]	282-01	5.0	Water, Leukocyte Extract
PENTACARE-HP	305-02	5.0	Water, Ceratonia Siliqua Gum, Hydrolyzed Casein
PENTAGLYCAN	181-01	5.0	Glycosaminoglycans
PENTAVITIN [®]	180-01	5.0	Saccharide Isomerate
PHYTALURONATE [®]	298-01	5.0	Ceratonia Siliqua Gum
PREREGEN [®]	315-01	5.0	Glycine Soja (Soybean) Protein, Oxido Reductases
REVITALIN [®] -BT	291-01	5.0	Glycoproteins
SERICIN	303-02	5.0	Sericin
SETAKOL [®]	320-01	5.0	Hydrolyzed Sericin

These water-soluble PENTAPHARM actives formed stable, clear to very slightly turbid solutions with PEPHA[®]-TIGHT.



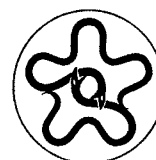
COMPATIBLE PENTAPHARM ACTIVES

Product Name	Product Code	% Wt.	INCI Name	Observations
ALPAFLOR [®] NECTAPURE	911-02	5.0	Glycerin, Water, Buddleja davidii Extract, Thymus vulgaris (Thyme) Extract	Slight pH decrease at 40°C
ALPHA-ARBUTIN	336-01	2.0	Alpha-Arbutin	Slight pH decrease at 40°C
DISMUTIN [®] -BT	313-01	1.0	Superoxide Dismutase	Slight precipitate
ERYTHRULOSE	307-01	5.0	Erythrulose	Slight precipitate, coloration*
MELFADE [®] -J	308-02	5.0	Water, Arctostaphylos Uva Ursi Leaf Extract, Glycerin, Magnesium Ascorbyl Phosphate	Slight precipitate
PENTACARE-NA	319-01	5.0	Hydrolyzed Wheat Gluten, Ceratonia Siliqua Gum	Slight precipitate

* As usual with ERYTHRULOSE, the solution developed an orange coloration at 40°C and pH decreased up to approx. 3.0-3.5 then stabilised.

PENTAPHARM ACTIVES WITH LIMITED COMPATIBILITY

Product Name	Product Code	% Wt.	INCI Name	Observations
ALPAFLOR [®] GIGAWHITE [™]	910-02	5.0	Water, Glycerin, Malva Sylvestris (Mallow) Extract, Mentha Piperita (Peppermint) Leaf Extract, Primula Veris Extract, Alchemilla Vulgaris Extract, Veronica Officinalis Extract, Melissa Officinalis Leaf Extract, Achillea Millefolium Extract	Precipitate
CEPHALIPIN	293-01	5.0	Cephalins, Disodium Cocoamphodiacetate	Precipitate
FITOBROSIDE	292-01	5.0	Triticum Vulgare (Wheat) Germ Extract	Precipitate
LACTOMIDE	309-01	5.0	Milk Lipids, Ceramide 3	Precipitate
PEPHA [®] -NUTRIX	800272	5.0	Colostrum Whey	Precipitate
REGU [®] -AGE	800131	5.0	Oxido Reductases, Glycine Soja (Soybean) Protein, Hydrolyzed Rice Bran Protein	Precipitate



DISCUSSION

PEPHA[®]-TIGHT is very easy to formulate in gels, cream gels and emulsions. As PEPHA[®]-TIGHT itself is an opalescent to turbid liquid, slight turbidity or clouding in aqueous solutions are not regarded as incompatibilities.

We are open to discuss any questions connected with the formulation of PEPHA[®]-TIGHT and any addition to these guidelines would be welcome.

PENTAPHARM LTD.



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SKIN TIGHTENING CREAM - IN VIVO STUDY (PEPHA®-TIGHT)

Description This soft cream was formulated for the in vivo study of PENTAPHARM's new active PEPHA®-TIGHT, which is a unique tightener through its immediate effect on the skin surface and its long-term effect on the collagen-I formation.

Item	Ingredients	INCI Name	% w/w	Suppl.
1	A) Water	Aqua	70.75	
2	Glycerin	Glycerin	3.00	
3	B) Crodafos CES	Cetearyl Alcohol, Dicapryl Phosphate, Ceteth-10 Phosphate	5.00	18
4	Myritol 331	Cocoglycerides	6.00	7
5	Tegosoft TN	C 12-15 Alkyl Benzoate	3.00	6
6	Tegosoft DC	Decyl Cocoate	3.00	6
7	Fitoderme	Squalane	2.00	30
8	C) NaOH (10% solution)	Sodium Hydroxide [q.s.]		
9	D) Dow Corning 345 Fluid	Cyclomethicone	3.00	16
10	Euxyl K-300	Phenoxyethanol, Methylparaben, Butylparaben, Ethylparaben, Propylparaben, Isobutylparaben	0.80	66
11	Aristoflex AVC	Ammonium Acryloyldimethyltaurate, VP Copolymer	0.40	87
12	E) PEPHA®-TIGHT	Algae Extract, Pullulan	3.00	1
13	Fragrance Cotton Flower 0298152	Fragrance	0.05	105

Procedure

Heat phases A) and B) to 75°C separately.
Under stirring add phase B) to phase A) and homogenize.
Adjust pH to approx. 5.5 - 6.5 with phase C).
At approx. 65°C, add ingredients of phase D) one after the other.
Homogenize till Aristoflex AVC is well dispersed.
At approx. 35°C add phase E).
If necessary, adjust pH again at 5.5 - 6.5.

Suppliers

1) PENTAPHARM, CH	18) Croda, GER	105) Symrise, GER
6) Degussa, GER	30) Hispano Quimica, E	
7) Cognis, GER	66) Schülke & Mayr, GER	
16) Dow Corning, USA	87) Clariant, CH	

We recommend to adjust the formulation and manufacture to the local conditions, to the raw materials available and to the current patent situation. Although this formulation has been designed with utmost care, we cannot assume any responsibility.

**QUICK TIGHTENING EFFECT - IN VIVO STUDY (PEPHA®-TIGHT)**

Description This tightening gel was developed for the short term in vivo study of PENTAPHARM's active PEPHA®-TIGHT, which is a unique tightener through its immediate effect on the skin surface and its long-term effect on the collagen-I formation.

Item	Ingredients	INCI Name	% w/w	Suppl.
1	A) Water	Aqua	93.50	
2	Preservative	[q.s.]		
3	B) RapiThix A-100	Sodium Polyacrylate	1.50	49
4	C) PEPHA®-TIGHT	Algae Extract, Pullulan	5.00	1

Procedure

Add component of phase B) to phase A) and let it swell.
Homogenize during 2 minutes.
Add phase C) to phase AB) and mix to obtain a homogeneous gel.

Suppliers

- 1) PENTAPHARM, CH
- 49) ISP, USA

We recommend to adjust the formulation and manufacture to the local conditions, to the raw materials available and to the current patent situation. Although this formulation has been designed with utmost care, we cannot assume any responsibility.



TIGHTENING EYE GEL - IN VIVO STUDY (PEPHA®-TIGHT / REGU®-AGE / REGU®-SHAPE)

Description This light gel was formulated for an in vivo study in the eye area with PENTAPHARM's new active PEPHA®-TIGHT, which has excellent tightening properties. REGU®-SHAPE prevents the accumulation of lipids in the adipocytes and REGU®-AGE strengthens the connective tissue.

Item	Ingredients	INCI Name	% w/w	Suppl.
1	A) Water	Aqua	87.40	
2	Pemulen TR-1	Acrylates/C10-30 Alkyl Acrylate Crosspolymer	0.60	8
3	B) Glycerin	Glycerin	3.50	
4	Preservative	[q.s.]		
5	C) NaOH (10% solution)	Sodium Hydroxide [q.s.]		
6	D) PEPHA®-TIGHT	Algae Extract, Pullulan	4.00	1
7	REGU®-AGE	Hydrolyzed Rice Bran Protein, Glycine Soja (Soybean) Protein, Oxido Reductases	2.00	1
8	REGU®-SHAPE	Isomerized Linoleic Acid, Lecithin, Glycerin, Polysorbate 80	2.00	1
9	Abil B 88183	PEG/PPG-20/6 Dimethicone	0.50	6

Procedure

Disperse item 2 in phase A).
Add phase B) to phase A).
Adjust pH to approx. 5.5-6.0 with phase C).
Add components of phase D one after the other.
Mix to obtain a homogeneous gel.

Suppliers

- 1) PENTAPHARM, CH
- 6) Degussa, GER
- 8) Noveon/B.F. Goodrich

We recommend to adjust the formulation and manufacture to the local conditions, to the raw materials available and to the current patent situation. Although this formulation has been designed with utmost care, we cannot assume any responsibility.



REJUVENATING CREAM (PEPHA[®]-TIGHT / SYN[®]-COLL)

Description This soft cream is formulated with PENTAPHARM's actives SYN[®]-COLL which stimulates the collagen synthesis and PEPHA[®]-TIGHT which has excellent tightening properties.

Item	Ingredients	INCI Name	% w/w	Suppl.
1	A) Tego Care 450	Polyglyceryl-3 Methylglucose Distearate	3.00	6
2	Tego Alkanol 1618	Cetearyl Alcohol	1.00	6
3	Oleic acid	Oleic acid	1.70	
4	Avocado Oil	Persea Gratissima (Avocado) Oil	2.40	91
5	Almond Oil, Sweet	Prunus Amygdalus Dulcis (Sweet Almond) Oil	2.40	91
6	Fitoderme	Squalane	4.00	30
7	Dragoxat EH	Ethylhexyl Ethylhexanoate	1.50	105
8	Tegosoft CT	Caprylic/Capric Triglyceride	6.00	6
9	Dow Corning 345 Fluid	Cyclomethicone	2.00	16
10	B) Water	Aqua	67.50	
11	Glycerin	Glycerin	3.00	
12	Keltrol RD	Xanthan Gum	0.20	65
13	Preservative	[q.s.]		
14	C) SYN [®] -COLL	Glycerin, Palmitoyl Tripeptide-5	2.50	1
15	PEPHA [®] -TIGHT	Algae Extract, Pullulan	2.50	1
16	D) Fragrance Organdi 0283320	Fragrance	0.30	105
17	E) Citric Acid (10% Solution)	Citric Acid [q.s.]		

Procedure

Disperse item 12 in phase B).
 Heat phases A) and B) separately to 80°C.
 Under stirring add phase A) to phase B), then homogenise.
 At 40°C add components of phase C) one after the other.
 At 30°C add phase D).
 Adjust pH to 5.0 - 5.5 with phase E).

Suppliers

- | | |
|------------------------|---------------------|
| 1) PENTAPHARM, CH | 65) Kelco, USA |
| 6) Degussa, GER | 91) H. Lamotte, GER |
| 16) Dow Corning, USA | 105) Symrise, GER |
| 30) Hispano Quimica, E | |

We recommend to adjust the formulation and manufacture to the local conditions, to the raw materials available and to the current patent situation. Although this formulation has been designed with utmost care, we cannot assume any responsibility.



TIGHTENING GEL FOR DECOLLETÉ (PENTAVITIN® / PEPHA®-TIGHT / PREREGEN®)

Description This smoothing gel is formulated with PENTAPHARM's new active PEPHA®-TIGHT, which has excellent tightening properties. PENTAVITIN® exerts a long lasting moisturization to the skin and PREREGEN® has radical scavenging and protease inhibiting properties.

Item	Ingredients	INCI Name	% w/w	Suppl.
1	A) Water	Aqua	90.00	
2	Preservative	[q.s.]		
3	B) RapiThix A-100	Sodium Polyacrylate	2.00	49
4	C) PEPHA®-TIGHT	Algae Extract, Pullulan	3.00	1
5	PENTAVITIN®	Saccharide Isomerase	2.00	1
6	PREREGEN®	Glycine Soya (Soybean) Protein, Oxido Reductases	3.00	1

Procedure

Add item 3 to phase A and let it swell.
Homogenise.
Add components of phase C one after the other and mix to obtain an homogeneous gel.

Suppliers

- 1) PENTAPHARM, CH
- 49) ISP, USA

We recommend to adjust the formulation and manufacture to the local conditions, to the raw materials available and to the current patent situation. Although this formulation has been designed with utmost care, we cannot assume any responsibility.



TIGHTENING CONTOURING GEL (PEPHA[®]-TIGHT / REGU[®]-AGE / REGU[®]-SHAPE)

Description This light gel was formulated with PENTAPHARM's new active PEPHA[®]-TIGHT, which has excellent tightening properties. REGU[®]-SHAPE prevents the accumulation of lipids in the adipocytes and REGU[®]-AGE strengthens the connective tissue.

Item	Ingredients	INCI Name	% w/w	Suppl.
1	A) Water	Aqua	88.40	
2	Pemulen TR-1	Acrylates/C10-30 Alkyl Acrylate Crosspolymer	0.60	8
3	B) Glycerin	Glycerin	3.50	
4	Preservative	[q.s.]		
5	C) NaOH (10% solution)	Sodium Hydroxide [q.s.]		
6	D) PEPHA [®] -TIGHT	Algae Extract, Pullulan	3.00	1
7	REGU [®] -SHAPE	Isomerized Linoleic Acid, Lecithin, Glycerin, Polysorbate 80	2.00	1
8	REGU [®] -AGE	Hydrolyzed Rice Bran Protein, Glycine Soja (Soybean) Protein, Oxido Reductases	2.00	1
9	Abil B 88183	PEG/PPG-20/6 Dimethicone	0.50	6

Procedure

Disperse item 2 in phase A).
 Add phase B) to phase A).
 Adjust pH to approx. 5.5-6.0 with phase C).
 Add components of phase D one after the other.
 Mix to obtain a homogeneous gel.

Suppliers

- 1) PENTAPHARM, CH
- 6) Degussa, GER
- 8) Noveon/B.F. Goodrich

We recommend to adjust the formulation and manufacture to the local conditions, to the raw materials available and to the current patent situation. Although this formulation has been designed with utmost care, we cannot assume any responsibility.



SKIN FIRMING GEL FOR MEN (PEPHA®-TIGHT)

Description This smooth gel is formulated with PENTAPHARM's active ingredient PEPHA®-TIGHT, which has excellent tightening properties and strengthens the skin's connective tissue. This formulation provides a dry and silky skin feeling. [formulation n°3994 developed by S.Black]

Item	Ingredients	INCI Name	% w/w	Suppl.
1	A) Water	Aqua	82.00	
2	Preservative	[q.s.]		
3	Softigen 767	PEG-6 Caprylic/Capric Glycerides	5.00	26
4	PEPHA®-TIGHT	Algae Extract, Pullulan	4.00	1
5	Lexfeel 7	Neopentyl Glycol Diheptanoate	6.00	29
6	B) Dow Corning RM 2051 Thickening Agent	Sodium Polyacrylate, Dimethicone, Cyclopentasiloxane, Trideceth-6, PEG/PPG-18/18 Dimethicone	3.00	16

Procedure

Premix phase A).
Add phase B) to phase A) and mix until homogeneous.

Suppliers

- 1) PENTAPHARM, CH
- 16) Dow Corning, USA
- 26) Hüls, GER
- 29) Inolex, USA

We recommend to adjust the formulation and manufacture to the local conditions, to the raw materials available and to the current patent situation. Although this formulation has been designed with utmost care, we cannot assume any responsibility.