SAFETY DATA SHEET

SECTION 1: Identification of the substance/mixture and of the company/undertaking

1.1. Product identifier

Trade name or designation of the mixture	CUTIVATE CREAM
Registration number	-
Synonyms	CUTIVATE CREAM 0.05% W/W * CUTIVAT CREAM 0.05% * FLUTIVATE CREAM 0.05% * FLIXODERM CREAM 0.05% * FLIXOVATE CREAM 0.05% * CUTIVATE CREME * CUTIVATE CREMA * CUTIVATE HYDROFIELE CREME * CUTIVATE KENOCS * CUTIVATE KREM * CUTIVATE KREMA * CUTIVATE KREMS * FLUTICASONE PROPIONATE, FORMULATED PRODUCT
Issue date	16-December-2013
Version number	13
Revision date	16-December-2013

1.2. Relevant identified uses of the substance or mixture and uses advised against

Identified uses	Medicinal Product This safety data sheet is written to provide health, safety and environmental information for people handling this formulated product in the workplace. It is not intended to provide information relevant to medicinal use of the product. In this instance patients should consult prescribing information/package insert/product label or consult their pharmacist or physician. For health and safety information for individual ingredients used during manufacturing, refer to the appropriate safety data sheet for each ingredient.
Uses advised against	No other uses are advised.

1.3. Details of the supplier of the safety data sheet

GlaxoSmithKline UK
980 Great West Road
Brentford, Middlesex TW8 9GS UK
UK General Information (normal business hours): +44-20-8047-5000
Email Address: msds@gsk.com
Website: www.gsk.com

1.4. Emergency telephone number

TRANSPORT EMERGENCIES::
UK In-country toll call: +(44)-870-8200418
International toll call: +1 703 527 3887
available 24 hrs/7 days; multi-language response

SECTION 2: Hazards identification

2.1. Classification of the substance or mixture

The mixture has been assessed and/or tested for its physical, health and environmental hazards and the following classification applies.

Classification according to Directive 67/548/EEC or 1999/45/EC as amended

Exempt from requirements - product regulated as a medicinal product, cosmetic product or medical device.

Classification according to Regulation (EC) No 1272/2008 as amended

Exempt from requirements - product regulated as a medicinal product, cosmetic product or medical device.

2.2. Label elements

Label according to Regulation (EC) No. 1272/2008 as amended

Exempt from requirements - product regulated as a medicinal product, cosmetic product or medical device.

Supplemental label information Not applicable.

2.3. Other hazards Caution - Pharmaceutical agent. See section 11 for additional information on health hazards. This product will support combustion at elevated temperatures.

SECTION 3: Composition/information on ingredients

3.2. Mixtures

General information

Chemical name	%	CAS-No. / EC No.	REACH Registration No.	INDEX No.	Notes
LIGHT MINERAL OIL	40 - < 50	8042-47-5 232-455-8	-	-	
Classification: **DSD:** -					
CLP: -					
Propylene glycol	10 - < 20	57-55-6 200-338-0	-	-	
Classification: **DSD:** -					
CLP: -					
ISOPROPYL MYRISTATE	5 - < 10	110-27-0 203-751-4	-	-	
Classification: **DSD:** Xi;R38					
CLP: Skin Irrit. 2;H315					
FLUTICASONE PROPIONATE	0.05	80474-14-2 -	-	-	
Classification: **DSD:** Repr. Cat. 2;R61, Repr. Cat. 3;R62, Xn;R22-48/20/21					
CLP: Acute Tox. 4;H302, Repr. 1B;H360D, Repr. 2;H361f, STOT RE 2;H373					
HYDROUS CITRIC ACID	< 0.1	5949-29-1 201-069-1	-	-	
Classification: **DSD:** Xi;R37/38-41					
CLP: Skin Irrit. 2;H315, Eye Dam. 1;H318, STOT SE 3;H335					

Other components below reportable levels 40 - < 50

CLP: Regulation No. 1272/2008.
DSD: Directive 67/548/EEC.
M: M-factor
vPvB: very persistent and very bioaccumulative substance.
PBT: persistent, bioaccumulative and toxic substance.
#: This substance has been assigned Community workplace exposure limit(s).

Composition comments The full text for all R- and H-phrases is displayed in section 16.

SECTION 4: First aid measures

General information The need for pre-placement and periodic health surveillance must be determined by risk assessment. Following assessment, if the risk of exposure is considered significant then exposed individuals should receive health surveillance focused on detecting skin conditions.
In the event of overexposure, individuals should receive post-exposure health surveillance focused on detecting skin conditions and adrenal suppression.

4.1. Description of first aid measures

Inhalation In case of accident by inhalation: remove casualty to fresh air and keep at rest. If not breathing, give artificial respiration. If breathing is difficult, trained personnel should give oxygen.

Skin contact Immediately flush with plenty of water for at least 15 minutes while removing contaminated clothing and shoes.

Eye contact Rinse thoroughly with plenty of water for at least 15 minutes and consult a physician.

Ingestion If swallowed, rinse mouth with water (only if the person is conscious).

4.2. Most important symptoms and effects, both acute and delayed Irritation of eyes and mucous membranes. The following adverse effects have been noted with therapeutic use of this material: headache; nosebleed; drying of the nasal passages; Irritation of nose and throat.

4.3. Indication of any immediate medical attention and special treatment needed No specific antidotes are recommended. Treat according to locally accepted protocols. For additional guidance, refer to the local poison control information centre.

SECTION 5: Firefighting measures

General fire hazards This product will support combustion at elevated temperatures.

5.1. Extinguishing media

Suitable extinguishing media Alcohol resistant foam. Dry chemical powder. Carbon dioxide (CO_2).

Unsuitable extinguishing media Water.

5.2. Special hazards arising from the substance or mixture During fire, gases hazardous to health may be formed.

5.3. Advice for firefighters

Special protective equipment for firefighters Self-contained breathing apparatus and full protective clothing must be worn in case of fire.

Special fire fighting procedures Use standard firefighting procedures and consider the hazards of other involved materials. Move containers from fire area if you can do so without risk.

SECTION 6: Accidental release measures

6.1. Personal precautions, protective equipment and emergency procedures

For non-emergency personnel Keep unnecessary personnel away. For personal protection, see section 8.

For emergency responders Keep unnecessary personnel away. Use personal protection recommended in Section 8 of the MSDS.

6.2. Environmental precautions Avoid discharge into drains, water courses or onto the ground.

6.3. Methods and material for containment and cleaning up Large Spills: Stop the flow of material, if this is without risk. Dike the spilled material, where this is possible. Cover with plastic sheet to prevent spreading. Absorb in vermiculite, dry sand or earth and place into containers. Use water spray to reduce vapours or divert vapour cloud drift. Following product recovery, flush area with water.

Small Spills: Wipe up with absorbent material (e.g. cloth, fleece). Clean surface thoroughly to remove residual contamination.

Never return spills in original containers for re-use.

6.4. Reference to other sections For personal protection, see section 8. For waste disposal, see section 13.

SECTION 7: Handling and storage

7.1. Precautions for safe handling No special control measures required for the normal handling of this product. Avoid prolonged or repeated contact with skin. Avoid prolonged exposure. Use only in well-ventilated areas.

7.2. Conditions for safe storage, including any incompatibilities Keep away from heat and sources of ignition. Store in original tightly closed container. Store away from incompatible materials (see Section 10 of the MSDS).

7.3. Specific end use(s) Pharmaceutical Bulk Mixture

SECTION 8: Exposure controls/personal protection

8.1. Control parameters

Occupational exposure limits

GSK

Components	Type	Value	Note
FLUTICASONE PROPIONATE (CAS 80474-14-2)	8 HR TWA	3 mcg/m3	
	OHC	4	Skin
		4	Reproductive hazard
HYDROUS CITRIC ACID (CAS 5949-29-1)	8 HR TWA	5000 mcg/m3	
	OHC	1	

UK. EH40 Workplace Exposure Limits (WELs)

Components	Type	Value	Form
Propylene glycol (CAS 57-55-6)	TWA	474 mg/m3	Total vapour and particulates.
		10 mg/m3	Particulate.
		150 ppm	Total vapour and particulates.

Recommended monitoring procedures Follow standard monitoring procedures.

Derived No Effect Level (DNEL) Not available.

Predicted no effect concentrations (PNECs)	Not available.

8.2. Exposure controls

Appropriate engineering controls	Good general ventilation (typically 10 air changes per hour) should be used. Ventilation rates should be matched to conditions. If applicable, use process enclosures, local exhaust ventilation, or other engineering controls to maintain airborne levels below recommended exposure limits. If exposure limits have not been established, maintain airborne levels to an acceptable level. An Exposure Control Approach (ECA) is established for operations involving this material based upon the OEL/Occupational Hazard Category and the outcome of a site- or operation-specific risk assessment.

Individual protection measures, such as personal protective equipment

General information	Personal protection equipment should be chosen according to the CEN standards and in discussion with the supplier of the personal protective equipment. Follow all local regulations if personal protective equipment (PPE) is used in the workplace.
Eye/face protection	Avoid contact with eyes. Wear approved safety glasses with side shields or cover goggles if eye contact is possible. (eg. EN 166)
Skin protection	
- Hand protection	The choice of an appropriate glove does not only depend on its material but also on other quality features and is different from one producer to the other. Glove selection must take into account any solvents and other hazards present. With respect to the above precautions select suitable chemical resistant protective gloves (EN 374) with a protective index 6 (>480min permeation time).
- Other	Wear suitable protective clothing. (EN 14605 for splashes, EN ISO 13982 for dust)
Respiratory protection	Where breathable aerosols/dust are formed, use suitable combination filter for gases/vapours of organic, inorganic, acid inorganic, alkaline compounds and toxic particles (eg. EN 14387).
Thermal hazards	Wear appropriate thermal protective clothing, when necessary.
Hygiene measures	An occupational/industrial hygiene monitoring method has been developed for this material. For advice on suitable monitoring methods, seek guidance from a qualified environment, health and safety professional. New or expectant mothers are at greater risk if exposed to the active ingredient which is readily absorbed through the skin. They should not handle unpackaged product. Risk assessments must take this into consideration. Female employees anticipating pregnancy or with a confirmed pregnancy must be encouraged to notify an occupational health professional or their line manager. This will act as the trigger for individual re-assessment of the employee's work practices.

Environmental exposure controls

Hazard guidance and control recommendations	Environmental manager must be informed of all major releases.

SECTION 9: Physical and chemical properties

9.1. Information on basic physical and chemical properties

Appearance

Physical state	Liquid.
Form	Cream.
Colour	Not available.
Odour	Not available.
Odour threshold	Not available.
pH	Not available.
Melting point/freezing point	Not available.
Initial boiling point and boiling range	Not available.
Flash point	> 95 °C (> 203 °F) Closed cup (Estimation based on components).
Evaporation rate	Not available.
Flammability (solid, gas)	Not available.

Upper/lower flammability or explosive limits

Flammability limit - lower (%)	Not available.
Flammability limit - upper (%)	Not available.
Vapour pressure	Not available.
Vapour density	Not available.
Relative density	Not available.

Solubility(ies)	Not available.
Partition coefficient (n-octanol/water)	Not available.
Auto-ignition temperature	Not available.
Decomposition temperature	Not available.
Viscosity	Not available.
Explosive properties	Not available.
Oxidizing properties	Not available.

9.2. Other information

Percent volatile	41.2 % estimated

SECTION 10: Stability and reactivity

10.1. Reactivity	Strong oxidising agents.
10.2. Chemical stability	Material is stable under normal conditions.
10.3. Possibility of hazardous reactions	No dangerous reaction known under conditions of normal use.
10.4. Conditions to avoid	Avoid heat, sparks, open flames and other ignition sources. Avoid temperatures exceeding the flash point. Contact with incompatible materials.
10.5. Incompatible materials	Strong oxidising agents.
10.6. Hazardous decomposition products	Irritating and/or toxic fumes and gases may be emitted upon the products decomposition.

SECTION 11: Toxicological information

General information	Caution - Pharmaceutical agent. Occupational exposure to the substance or mixture may cause adverse effects.

Information on likely routes of exposure

Ingestion	May be harmful if swallowed.
Inhalation	Under normal conditions of intended use, this material is not expected to be an inhalation hazard.
Skin contact	Pharmacological effects might occur following direct contact with skin. Repeated contact may increase sensitivity of skin to bruising.
Eye contact	May be irritating to eyes.
Symptoms	The following adverse effects have been noted with therapeutic use of this material: headache; nosebleed; drying of the nasal passages; Irritation of nose and throat..

11.1. Information on toxicological effects

Acute toxicity	May be harmful if swallowed. May be harmful in contact with skin.

Components	Species	Test results
FLUTICASONE PROPIONATE (CAS 80474-14-2)		
Acute		
Oral		
LD50	Rat	> 1000 mg/kg
Subacute		
Inhalation		
NOAEL	Rat	0.2 mcg/L/day, 28 Day
Subchronic		
Inhalation		
LOEL	Rat	3 mcg/kg/day, 26 weeks
NOAEL	Dog	68 mcg/kg/day, 26 weeks
	Rat	14 mcg/kg/day, 26 weeks
LIGHT MINERAL OIL (CAS 8042-47-5)		
Acute		
Oral		
LD50	Rat	> 2000 mg/kg

* Estimates for product may be based on additional component data not shown.

Skin corrosion/irritation	Repeated contact may increase sensitivity of skin to bruising.

Corrosivity

FLUTICASONE PROPIONATE	OECD 404 Result: negative

Irritation Corrosion - Skin: P.I.I. value

FLUTICASONE PROPIONATE	0

Serious eye damage/eye irritation May be irritating to eyes.

Eye

FLUTICASONE PROPIONATE	OECD 405 Result: negative Species: Rabbit

Respiratory sensitisation None known.

Skin sensitisation Allergic skin reactions might occur following repeated contact with this material in susceptible individuals.

Sensitisation

FLUTICASONE PROPIONATE	0 % OECD 406 Result: negative Species: Guinea pig

Germ cell mutagenicity No data available to indicate product or any components present at greater than 0.1% are mutagenic or genotoxic.

Germ cell mutagenicity

Mutagenicity

FLUTICASONE PROPIONATE	Ames Result: negative Bacterial High Throughput Fluctuation Test Result: negative Chinese Hamster Ovarian Cell Test Result: negative Micronucleus Assay Result: negative Species: Mouse Micronucleus Test Result: negative Species: Mouse SOS/umu Assay Result: negative Yeast Result: negative

Carcinogenicity Not classifiable as to carcinogenicity to humans.

FLUTICASONE PROPIONATE	Inhalation Result: negative Species: Rat dermal Result: negative Species: Mouse oral Result: negative Species: Mouse

IARC Monographs. Overall Evaluation of Carcinogenicity

LIGHT MINERAL OIL (CAS 8042-47-5)	3 Not classifiable as to carcinogenicity to humans.

Reproductive toxicity Components in this product have been shown to cause birth defects and reproductive disorders in laboratory animals.

Reproductive toxicity

Reproductivity

FLUTICASONE PROPIONATE	100 mcg/kg/day Embryofetal Development Result: reduced foetal bodyweight, minor skeletal variations Species: Rat 100 mcg/kg/day Female fertility (Segment I) Result: reduced foetal bodyweight, minor skeletal variations Species: Rat 50 mcg/kg/day Pre- and Post-natal development Result: maternal toxicity Species: Rat >= 25.7 mcg/kg/day Embryofetal Development Result: maternal toxicity, reduced foetal body weight; no malformations or other variations Species: Rat

Reproductivity

FLUTICASONE PROPIONATE	>= 45 mcg/kg/day Embryofetal Development Result: cleft palate Species: Mouse >= 50 mcg/kg/day Embryofetal Development Result: maternal toxicity; reduced foetal weight; foetal resorptions Species: Rabbit SAR / QSAR, Glucocorticoid

Specific target organ toxicity - single exposure None known.

Specific target organ toxicity - repeated exposure Adrenal glands. Bone tissue. Immune system. May cause damage to organs through prolonged or repeated exposure.

Aspiration hazard Not established.

Mixture versus substance information No information available.

Other information Caution - Pharmaceutical agent.

SECTION 12: Ecological information

12.1. Toxicity The product contains a substance which may cause long-term adverse effects in the environment. No information is available about the potential of this product to produce adverse environmental effects.

Components		Species	Test results
FLUTICASONE PROPIONATE (CAS 80474-14-2)			
Acute			
	IC50	Activated sludge	> 1000 mg/l, 3 hours
Aquatic			
Acute			
Crustacea	EC50	Water flea (Daphnia magna)	> 0.55 mg/l, 48 hours, Static test
Terrestrial			
Acute			
Earthworm	EC50	Manure worm (Eisenia foetida)	> 1000 mg/kg, 28 days
Propylene glycol (CAS 57-55-6)			
Acute			
	IC50	Activated sludge	> 1000 mg/l, 3 hours
Aquatic			
Acute			
Algae	EC50	Green algae (Selenastrum capricornutum)	19000 mg/l, 14 days
	NOEC	Green algae (Selenastrum capricornutum)	15000 mg/l, 14 days
Crustacea	EC50	Daphnia	43500 mg/l, 48 hours
	NOEC	Daphnia	28500 mg/l, 48 hours
Fish	EC50	Fathead minnow (Adult Pimephales promelas)	51400 mg/l, 96 hours, Static test
		Rainbow trout (Adult Oncorhyncus mykiss)	51600 mg/l, 96 hours, Static test
	NOEC	Fathead minnow (Adult Pimephales promelas)	41000 mg/l, 96 hours, Static test
		Rainbow trout (Adult Oncorhyncus mykiss)	42000 mg/l, 96 hours, Static test
Microtox	EC50	Microtox	51400 mg/l, 30 minutes

* Estimates for product may be based on additional component data not shown.

12.2. Persistence and degradability

Persistence and degradability

Photolysis

Half-life (Photolysis-aqueous)

Propylene glycol	1.3 - 2.3 years Estimated

Half-life (Photolysis-atmospheric)

LIGHT MINERAL OIL	< 1 Days Estimated
Propylene glycol	32 Hours Estimated

Hydrolysis

Half-life (Hydrolysis-neutral)

FLUTICASONE PROPIONATE	> 1 years Measured

Biodegradability

Percent degradation (Aerobic biodegradation-inherent)

Propylene glycol	62 %, 5 days BOD5, Activated sludge 79 %, 20 Days BOD20, Activated sludge

Percent degradation (Aerobic biodegradation-ready)

FLUTICASONE PROPIONATE	< 44 %, 28 days
LIGHT MINERAL OIL	24 %, 28 days Modified Sturm test., Activated sludge

Percent degradation (Aerobic biodegradation-soil)

FLUTICASONE PROPIONATE	9 - 50 %, 64 days

Percent degradation (Anaerobic biodegradation)

Propylene glycol	100 %, 9 days

12.3. Bioaccumulative potential

Partition coefficient n-octanol/water (log Kow)

FLUTICASONE PROPIONATE	2.78
Propylene glycol	-0.92 -1.35

Bioconcentration factor (BCF)

Propylene glycol	< 1 Estimated

12.4. Mobility in soil

Adsorption

Sludge/biomass distribution coefficient - log Kd

FLUTICASONE PROPIONATE	3.13 - 3.55 Estimated

Soil/sediment sorption - log Koc

FLUTICASONE PROPIONATE	3.41 - 3.83 Measured

Mobility in general

Volatility

Henry's law

Propylene glycol	0 atm m^3/mol Estimated

12.5. Results of PBT and vPvB assessment Not available.

12.6. Other adverse effects Not available.

SECTION 13: Disposal considerations

13.1. Waste treatment methods

Residual waste Dispose of in accordance with local regulations. Empty containers or liners may retain some product residues. This material and its container must be disposed of in a safe manner (see: Disposal instructions).

Contaminated packaging Empty containers should be taken to an approved waste handling site for recycling or disposal. Since emptied containers may retain product residue, follow label warnings even after container is emptied.

EU waste code The Waste code should be assigned in discussion between the user, the producer and the waste disposal company.

Disposal methods/information Collect and reclaim or dispose in sealed containers at licensed waste disposal site.

Special precautions Dispose in accordance with all applicable regulations.

SECTION 14: Transport information

ADR

Not regulated as dangerous goods.

IATA

Not regulated as dangerous goods.

IMDG

Not regulated as dangerous goods.

14.7. Transport in bulk according to Annex II of MARPOL73/78 and the IBC Code

MARPOL Annex II applies to liquids used in a ship's operation that pose a threat to the marine environment. These materials may not be transported in bulk.

SECTION 15: Regulatory information

15.1. Safety, health and environmental regulations/legislation specific for the substance or mixture

EU regulations

Regulation (EC) No. 1005/2009 on substances that deplete the ozone layer, Annex I

Not listed.

Regulation (EC) No. 1005/2009 on substances that deplete the ozone layer, Annex II

Not listed.

Regulation (EC) No. 850/2004 On persistent organic pollutants, Annex I as amended

Not listed.

Regulation (EC) No. 689/2008 concerning the export and import of dangerous chemicals, Annex I, part 1 as amended

Not listed.

Regulation (EC) No. 689/2008 concerning the export and import of dangerous chemicals, Annex I, part 2 as amended

Not listed.

Regulation (EC) No. 689/2008 concerning the export and import of dangerous chemicals, Annex I, part 3 as amended

Not listed.

Regulation (EC) No. 689/2008 concerning the export and import of dangerous chemicals, Annex V as amended

Not listed.

Regulation (EC) No. 166/2006 Annex II Pollutant Release and Transfer Registry

Not listed.

Regulation (EC) No. 1907/2006, REACH Article 59(1) Candidate List as currently published by ECHA

Not listed.

Authorisations

Regulation (EC) No. 1907/2006, REACH Annex XIV Substances subject to authorization, as amended

Not listed.

Restrictions on use

Regulation (EC) No. 1907/2006, REACH Annex XVII Substances subject to restriction on marketing and use as amended

Not listed.

Directive 2004/37/EC: on the protection of workers from the risks related to exposure to carcinogens and mutagens at work

Not listed.

Directive 92/85/EEC: on the safety and health of pregnant workers and workers who have recently given birth or are breastfeeding

Not listed.

Other EU regulations

Directive 96/82/EC (Seveso II) on the control of major-accident hazards involving dangerous substances

Not listed.

Directive 98/24/EC on the protection of the health and safety of workers from the risks related to chemical agents at work

Not listed.

Directive 94/33/EC on the protection of young people at work

Not listed.

Other regulations

The product is classified and labelled in accordance with EC directives or respective national laws. This Safety Data Sheet complies with the requirements of Regulation (EC) No 1907/2006.

National regulations

Follow national regulation for work with chemical agents.

15.2. Chemical safety assessment

No Chemical Safety Assessment has been carried out.

SECTION 16: Other information

List of abbreviations

Not available.

References

GSK Hazard Determination

Information on evaluation method leading to the classification of mixture

The classification for health and environmental hazards is derived by a combination of calculation methods and test data, if available.

Full text of any statements or R-phrases and H-statements under Sections 2 to 15	R22 Harmful if swallowed. R37/38 Irritating to respiratory system and skin. R38 Irritating to skin. R41 Risk of serious damage to eyes. R48/20/21 Harmful: danger of serious damage to health by prolonged exposure through inhalation and in contact with skin. R61 May cause harm to the unborn child. R62 Possible risk of impaired fertility. H302 Harmful if swallowed. H315 Causes skin irritation. H318 Causes serious eye damage. H335 May cause respiratory irritation. H360D May damage the unborn child. H361f Suspected of damaging fertility.
Revision information	Product and Company Identification: Product and Company Identification Composition / Information on Ingredients: Ingredients Physical & Chemical Properties: TRANSPORT INFORMATION: Regulatory Information: Risk Phrases - Class. GHS: Classification
Training information	Follow training instructions when handling this material.
Disclaimer	The information and recommendations in this safety data sheet are, to the best of our knowledge, accurate as of the date of issue. Nothing herein shall be deemed to create any warranty, express or implied. It is the responsibility of the user to determine the applicability of this information and the suitability of the material or product for any particular purpose.