# SAFETY DATA SHEET



## 1. IDENTIFICATION OF THE SUBSTANCE/MIXTURE AND THE COMPANY/UNDERTAKING

**Product Identifier**

**Material Name: Sunitinib Malate Capsules**

**Trade Name:** Sutent®
**Chemical Family:** Mixture

**Relevant Identified Uses of the Substance or Mixture and Uses Advised Against**

**Intended Use:** Pharmaceutical product used as Antineoplastic.

**Details of the Supplier of the Safety Data Sheet**

**Pfizer Inc**
**Pfizer Pharmaceuticals Group**
**235 East 42nd Street**
**New York, New York 10017**
**1-800-879-3477**

**Emergency telephone number:**
**CHEMTREC (24 hours): 1-800-424-9300**
**Contact E-Mail:** pfizer-MSDS@pfizer.com

**Pfizer Ltd**
**Ramsgate Road**
**Sandwich, Kent**
**CT13 9NJ**
**United Kingdom**
**+00 44 (0)1304 616161**
**Emergency telephone number:**
**International CHEMTREC (24 hours): +1-703-527-3887**

## 2. HAZARDS IDENTIFICATION

**Classification of the Substance or Mixture**

**GHS - Classification**

Reproductive Toxicity: Category 1B
Specific target organ systemic toxicity (repeated exposure): Category 1
Acute aquatic toxicity: Category 1
Chronic aquatic toxicity: Category 1

**EU Classification:**

EU Indication of danger: Toxic to reproduction, Category 2
T - Toxic
Dangerous for the Environment

EU Risk Phrases:

R48/25 - Toxic: danger of serious damage to health by prolonged exposure if swallowed.
R61 - May cause harm to the unborn child.
R50/53 - Very toxic to aquatic organisms, may cause long-term adverse effects in the aquatic environment.

**Label Elements**

**Signal Word:** Danger
**Hazard Statements:** H360D - May damage the unborn child
H372 - Causes damage to organs through prolonged or repeated exposure
H400 - Very toxic to aquatic life
H410 - Very toxic to aquatic life with long lasting effects

**Precautionary Statements:**

P201 - Obtain special instructions before use
P202 - Do not handle until all safety precautions have been read and understood
P281 - Use personal protective equipment as required
P308 + P313 - IF exposed or concerned: Get medical attention/advice
P260 - Do not breathe dust/fume/gas/mist/vapors/spray
P264 - Wash hands thoroughly after handling
P270 - Do not eat, drink or smoke when using this product
P314 - Get medical attention/advice if you feel unwell
P405 - Store locked up
P501 - Dispose of contents/container in accordance with all local and national regulations

**Other Hazards** No data available
**Australian Hazard Classification (NOHSC):** Hazardous Substance. Dangerous Goods.

**Note:** This document has been prepared in accordance with standards for workplace safety, which requires the inclusion of all known hazards of the product or its ingredients regardless of the potential risk. The precautionary statements and warning included may not apply in all cases. Your needs may vary depending upon the potential for exposure in your workplace.

## 3. COMPOSITION / INFORMATION ON INGREDIENTS

**Hazardous**

| Ingredient | CAS Number | EU EINECS/ELINCS List | EU Classification | GHS Classification | % |
|---|---|---|---|---|---|
| Sunitinib malate | 341031-54-7 | Not Listed | T;R48/25<br>N;R50/53<br>Repr.Cat2;R61 | STOT RE. 1 (H372)<br>Repr 1B (H360D)<br>Aquatic Acute 1 (H400)<br>Acute Chronic 1 (H410) | 15-40 |
| Magnesium stearate | 557-04-0 | 209-150-3 | Not Listed | Not Listed | * |

| Ingredient | CAS Number | EU EINECS/ELINCS List | EU Classification | GHS Classification | % |
|---|---|---|---|---|---|
| Mannitol | 69-65-8 | 200-711-8 | Not Listed | Not Listed | * |
| Croscarmellose sodium | 74811-65-7 | Not Listed | Not Listed | Not Listed | * |
| Povidone | 9003-39-8 | Not Listed | Not Listed | Not Listed | * |
| Hard gelatin capsules | MIXTURE | Not Listed | Not Listed | Not Listed | * |

**Additional Information:** * Proprietary
Ingredient(s) indicated as hazardous have been assessed under standards for workplace safety.
In accordance with 29 CFR 1910.1200, the exact percentage composition of this mixture has been withheld as a trade secret.

**For the full text of the R phrases and CLP/GHS abbreviations mentioned in this Section, see Section 16**

## 4. FIRST AID MEASURES

**Description of First Aid Measures**

**Eye Contact:** Flush with water while holding eyelids open for at least 15 minutes. Seek medical attention immediately.

**Skin Contact:** Remove contaminated clothing. Flush area with large amounts of water. Use soap. Seek medical attention.

**Ingestion:** Never give anything by mouth to an unconscious person. Wash out mouth with water. Do not induce vomiting unless directed by medical personnel. Seek medical attention immediately.

**Inhalation:** Remove to fresh air and keep patient at rest. Seek medical attention immediately.

**Most Important Symptoms and Effects, Both Acute and Delayed**

**Symptoms and Effects of Exposure:** For information on potential signs and symptoms of exposure, See Section 2 - Hazards Identification and/or Section 11 - Toxicological Information.

**Medical Conditions Aggravated by Exposure:** None known

**Indication of the Immediate Medical Attention and Special Treatment Needed**

**Notes to Physician:** None

## 5. FIRE FIGHTING MEASURES

**Extinguishing Media:** Extinguish fires with CO2, extinguishing powder, foam, or water.

**Special Hazards Arising from the Substance or Mixture**

**Hazardous Combustion Products:** Formation of toxic gases is possible during heating or fire.

**Fire / Explosion Hazards:** Not applicable

**Advice for Fire-Fighters**

During all fire fighting activities, wear appropriate protective equipment, including self-contained breathing apparatus.

## 6. ACCIDENTAL RELEASE MEASURES

**Personal Precautions, Protective Equipment and Emergency Procedures**

Personnel involved in clean-up should wear appropriate personal protective equipment (see Section 8). Minimize exposure.

**Environmental Precautions**

Place waste in an appropriately labeled, sealed container for disposal. Care should be taken to avoid environmental release.

**Methods and Material for Containment and Cleaning Up**

**Measures for Cleaning / Collecting:** Contain the source of spill if it is safe to do so. Collect spilled material by a method that controls dust generation. A damp cloth or a filtered vacuum should be used to clean spills of dry solids. Clean spill area thoroughly.

**Additional Consideration for Large Spills:** Non-essential personnel should be evacuated from affected area. Report emergency situations immediately. Clean up operations should only be undertaken by trained personnel.

## 7. HANDLING AND STORAGE

**Precautions for Safe Handling**

Minimize dust generation and accumulation. If tablets or capsules are crushed and/or broken, avoid breathing dust and avoid contact with eyes, skin, and clothing. When handling, use appropriate personal protective equipment (see Section 8). Wash hands and any exposed skin after removal of PPE. Refer to Section 12 - Ecological Information, for information on potential effects on the environment. Releases to the environment should be avoided. Review and implement appropriate technical and procedural waste water and waste disposal measures to prevent occupational exposure or environmental releases. Potential points of process emissions of this material to the atmosphere should be controlled with dust collectors, HEPA filtration systems or other equivalent controls.

**Conditions for Safe Storage, Including any Incompatibilities**

**Storage Conditions:** Store as directed by product packaging.
**Specific end use(s):** Pharmaceutical drug product

## 8. EXPOSURE CONTROLS / PERSONAL PROTECTION

**Control Parameters**

**Refer to available public information for specific member state Occupational Exposure Limits.**

**Sunitinib malate**

**Pfizer OEL TWA-8 Hr:** 10 µg/m$^3$

**Magnesium stearate**

**ACGIH Threshold Limit Value (TWA)** 10 mg/m$^3$
**Lithuania OEL - TWA** 5 mg/m$^3$
**Sweden OEL - TWAs** 5 mg/m$^3$

**Exposure Controls**

**Engineering Controls:** Engineering controls should be used as the primary means to control exposures. General room ventilation is adequate unless the process generates dust, mist or fumes. Keep airborne contamination levels below the exposure limits listed above in this section.

**Personal Protective Equipment:** Refer to applicable national standards and regulations in the selection and use of personal protective equipment (PPE).

**Hands:** Impervious gloves are recommended if skin contact with drug product is possible and for bulk processing operations.

**Eyes:** Wear safety glasses or goggles if eye contact is possible.

**Skin:** Impervious protective clothing is recommended if skin contact with drug product is possible and for bulk processing operations.

**Respiratory protection:** If the applicable Occupational Exposure Limit (OEL) is exceeded, wear an appropriate respirator with a protection factor sufficient to control exposures to below the OEL.

## 9. PHYSICAL AND CHEMICAL PROPERTIES

| | | | |
|---|---|---|---|
| **Physical State:** | Capsule | **Color:** | Orange; opaque brown. |
| **Odor:** | No data available. | **Odor Threshold:** | No data available. |
| **Molecular Formula:** | Mixture | **Molecular Weight:** | Mixture |

**Solvent Solubility:** No data available
**Water Solubility:** No data available

## 9. PHYSICAL AND CHEMICAL PROPERTIES

**pH:** No data available.
**Melting/Freezing Point (°C):** No data available
**Boiling Point (°C):** No data available.
**Partition Coefficient: (Method, pH, Endpoint, Value)**
**Mannitol**
No data available
**Croscarmellose sodium**
No data available
**Povidone**
No data available
**Hard gelatin capsules**
No data available
**Magnesium stearate**
No data available
**Sunitinib malate**
No data available
**Decomposition Temperature (°C):** No data available.

**Evaporation Rate (Gram/s):** No data available
**Vapor Pressure (kPa):** No data available
**Vapor Density (g/ml):** No data available
**Relative Density:** No data available
**Viscosity:** No data available

**Flammablity:**
- **Autoignition Temperature (Solid) (°C):** No data available
- **Flammability (Solids):** No data available
- **Flash Point (Liquid) (°C):** No data available
- **Upper Explosive Limits (Liquid) (% by Vol.):** No data available
- **Lower Explosive Limits (Liquid) (% by Vol.):** No data available

## 10. STABILITY AND REACTIVITY

**Reactivity:** No data available
**Chemical Stability:** Stable under normal conditions of use.
**Possibility of Hazardous Reactions**
- **Oxidizing Properties:** No data available
- **Conditions to Avoid:** None known
- **Incompatible Materials:** As a precautionary measure, keep away from strong oxidizers
- **Hazardous Decomposition Products:** No data available

## 11. TOXICOLOGICAL INFORMATION

**Information on Toxicological Effects**
**General Information:** The information included in this section describes the potential hazards of the individual ingredients.
**Short Term:** May cause mild eye irritation. (based on components) .
**Long Term:** Repeat-dose studies in animals have shown a potential to cause adverse effects on the hematological and reproductive systems.
**Known Clinical Effects:** Common adverse effects include fatigue, gastrointestinal disturbances, hematological effects, and skin effects. Other, more serious, effects include changes in liver function, liver failure

## 11. TOXICOLOGICAL INFORMATION

**Acute Toxicity: (Species, Route, End Point, Dose)**

**Mannitol**
Rat Oral LD 50 13500 mg/kg
Mouse Oral LD 50 22 g/kg

**Povidone**
Rat Oral LD50 100 g/kg

**Magnesium stearate**
Rat Oral LD50 > 2000 mg/kg
Rat Inhalation LC50 > 2000 mg/m$^3$

**Sunitinib malate**
Rat Oral Maximally Tolerated Dose >500 mg/kg
Mouse Oral Maximally Tolerated Dose >500mg/kg
Dog Oral Maximally Tolerated Dose >500mg/kg
Non-human Primate Oral Maximally Tolerated Dose >1200mg/kg
**Acute Toxicity Comments:** A greater than symbol (>) indicates that the toxicity endpoint being tested was not achievable at the highest dose used in the test.

**Irritation / Sensitization: (Study Type, Species, Severity)**

**Sunitinib malate**
Skin Irritation Rabbit Non-irritating
Eye Irritation Rabbit Mild

**Repeated Dose Toxicity: (Duration, Species, Route, Dose, End Point, Target Organ)**

**Sunitinib malate**
4 Week(s) Rat Oral 5 mg/kg/day NOAEL Bone marrow, Blood forming organs
28/56 Day(s) Monkey Oral 6.0 mg/kg/day LOAEL Bone Marrow, Blood forming organs
13 Week(s) Non-human Primate Oral 2.0 mg/kg/day LOAEL Bone Marrow, Blood forming organs
3 Month(s) Rat Oral 1.5 mg/kg/day NOAEL Bone Marrow, Blood forming organs
6 Month(s) Rat Oral 0.3 mg/kg/day NOAEL Bone Marrow

**Reproduction & Developmental Toxicity: (Study Type, Species, Route, Dose, End Point, Effect(s))**

**Sunitinib malate**
Fertility & Early Embryonic Development-Females Rat Oral 1.5 mg/kg/day NOAEL Fetotoxicity
Embryo / Fetal Development Rabbit Oral 0.5 mg/kg/day NOAEL Fetotoxicity
Embryo / Fetal Development Rabbit Oral 1.0 mg/kg/day NOAEL Maternal Toxicity
Embryo / Fetal Development Rat Oral 3 mg/kg/day NOAEL Fetotoxicity
Embryo / Fetal Development Rat Oral 5 mg/kg/day NOAEL Maternal Toxicity

**Genetic Toxicity: (Study Type, Cell Type/Organism, Result)**

**Sunitinib malate**
Bacterial Mutagenicity (Ames) *Salmonella* , *E. coli* Negative
Mammalian Cell Mutagenicity Negative
*In Vitro* Chromosome Aberration Human Lymphocytes Negative

## 11. TOXICOLOGICAL INFORMATION

*In Vivo* Micronucleus Rat Negative

**Sunitinib malate**
6 Month(s) Mouse Female Oral 8 mg/kg/day NOEL Gastrointestinal system
6 Month(s) Mouse Male Oral 25 mg/kg/day NOEL Gastrointestinal system

**Carcinogen Status:** None of the components of this formulation are listed as a carcinogen by IARC, NTP or OSHA.

**Povidone**
**IARC:** Group 3 (Not Classifiable)

## 12. ECOLOGICAL INFORMATION

**Environmental Overview:** Very toxic to aquatic life with long lasting effects. Releases to the environment should be avoided. See aquatic toxicity data, below:

**Toxicity:**
**Aquatic Toxicity: (Species, Method, End Point, Duration, Result)**

**Sunitinib malate**
*Daphnia magna* (Water Flea) OECD EC50 48 Hours 3.1 mg/L
*Oncorhynchus mykiss* (Rainbow Trout) OECD LC50 96 Hours 7.8 mg/L
*Pseudokirchneriella subcapitata* (Green Alga) OECD EC50 72 Hours 0.32 mg/L
*Daphnia magna* (Water Flea) OECD NOEC 21 Days 0.053 mg/L
*Ceriodaphnia dubia* (Daphnids) EPA NOEC 7 Days 0.32 mg/L
*Pimephales promelas* (Fathead Minnow) OECD NOEC 32 Days 0.00027 mg/L

**Bacterial Inhibition: (Inoculum, Method, End Point, Result)**

**Sunitinib malate**
Activated sludge OECD EC50 574 mg/L
*Clostridium perfingens* FDA MIC 80 mg/L
*Bacillus subtilis* (Bacterium) FDA MIC 80 mg/L
*Nostoc sp.* (Freshwater Cyanobacteria) FDA MIC 5.0 mg/L

**Persistence and Degradability:** No data available
**Sunitinib malate**
OECD Soil (various) Ready 8.8% After 28 Day(s)

**Bio-accumulative Potential:** No data available

**Mobility in Soil:** No data available

## 13. DISPOSAL CONSIDERATIONS

**Waste Treatment Methods:** Dispose of waste in accordance with all applicable laws and regulations. Member State specific and Community specific provisions must be considered. Considering the relevant known environmental and human health hazards of the material, review and implement appropriate technical and procedural waste water and waste disposal measures to prevent occupational exposure and environmental release. It is recommended that waste minimization be practiced. The best available technology should be utilized to prevent environmental releases. This may include destructive techniques for waste and wastewater.

**The following refers to all modes of transportation unless specified below.**

This material is regulated for transportation as a hazardous material/dangerous good.

| | |
|---|---|
| **UN number:** | UN 3077 |
| **UN proper shipping name:** | Environmentally Hazardous Substance, Solid, n.o.s (sunitinib malate) |
| **Transport hazard class(es):** | 9 |
| **Packing group:** | III |
| **Environmental Hazard(s):** | Marine Pollutant |

## 15. REGULATORY INFORMATION

**Safety, Health and Environmental Regulations/Legislation Specific for the Substance or Mixture**

**Canada - WHMIS: Classifications**
**WHMIS hazard class:**
Class D, Division 2, Subdivision A

**Sunitinib malate**

| | |
|---|---|
| **CERCLA/SARA 313 Emission reporting** | Not Listed |
| **California Proposition 65** | Not Listed |
| **EU EINECS/ELINCS List** | Not Listed |

**Mannitol**

| | |
|---|---|
| **CERCLA/SARA 313 Emission reporting** | Not Listed |
| **California Proposition 65** | Not Listed |
| **Inventory - United States TSCA - Sect. 8(b)** | Present |
| **Australia (AICS):** | Present |
| **REACH - Annex IV - Exemptions from the obligations of Register:** | Present |
| **EU EINECS/ELINCS List** | 200-711-8 |

**Croscarmellose sodium**

## 15. REGULATORY INFORMATION

| | |
|---|---|
| **CERCLA/SARA 313 Emission reporting** | Not Listed |
| **California Proposition 65** | Not Listed |
| **Australia (AICS):** | Present |
| **EU EINECS/ELINCS List** | Not Listed |

**Povidone**

| | |
|---|---|
| **CERCLA/SARA 313 Emission reporting** | Not Listed |
| **California Proposition 65** | Not Listed |
| **Inventory - United States TSCA - Sect. 8(b)** | Present |
| **Australia (AICS):** | Present |
| **EU EINECS/ELINCS List** | Not Listed |

**Magnesium stearate**

| | |
|---|---|
| **CERCLA/SARA 313 Emission reporting** | Not Listed |
| **California Proposition 65** | Not Listed |
| **Inventory - United States TSCA - Sect. 8(b)** | Present |
| **Australia (AICS):** | Present |
| **EU EINECS/ELINCS List** | 209-150-3 |

**Hard gelatin capsules**

| | |
|---|---|
| **CERCLA/SARA 313 Emission reporting** | Not Listed |
| **California Proposition 65** | Not Listed |
| **EU EINECS/ELINCS List** | Not Listed |

## 16. OTHER INFORMATION

**Text of R phrases and GHS Classification abbreviations mentioned in Section 3**

Reproductive toxicity-Cat.1B; H360D - May damage the unborn child
Specific target organ toxicity, repeated exposure-Cat.1; H373 - May cause damage to organs through prolonged or repeated exposure
Hazardous to the aquatic environment, acute toxicity-Cat.1; H400 - Very toxic to aquatic life
Hazardous to the aquatic environment, chronic toxicity-Cat.1; H410 - Very toxic to aquatic life with long lasting effects

T - Toxic
N - Dangerous for the environment
Toxic to reproduction: Category 1

R61 - May cause harm to the unborn child.
R48/25 - Toxic: danger of serious damage to health by prolonged exposure if swallowed.
R50/53 - Very toxic to aquatic organisms, may cause long-term adverse effects in the aquatic environment.

**Data Sources:** Pfizer proprietary drug development information. Safety data sheets for individual ingredients.

**Reasons for Revision:** Updated Section 2 - Hazard Identification. Updated Section 3 - Composition / Information on Ingredients. Updated Section 14 - Transport Information. Updated Section 15 - Regulatory Information.

**Revision date:** 27-Mar-2014

Product Stewardship Hazard Communication

**Prepared by:** Pfizer Global Environment, Health, and Safety Operations

Pfizer Inc believes that the information contained in this Material Safety Data Sheet is accurate, and while it is provided in good faith, it is without warranty of any kind, expressed or implied. If data for a hazard are not included in this document there is no known information at this time.

**End of Safety Data Sheet**