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

1.1. Product identifier

Product name: ROZLYTREK™ Capsules (100 mg)
Product code: RO710-2122-F09
Synonyms: Entrectinib Capsules (hard) 100 mg

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

Use: pharmaceutical active substance

1.3. Details of the supplier of the safety data sheet

Company information: Genentech, Inc.
1 DNA Way
South San Francisco
USA-CA 94080
United States of America

Phone: 001-(650) 225-1000
E-Mail: info.sds@roche.com
US Chemtrec phone: (800)-424-9300

1.4. Emergency telephone number

Emergency telephone number: US Chemtrec phone: (800)-424-9300

*1 referring to: Entrectinib
SECTION 2: Hazards identification

Classification of the substance or mixture / Label elements

GHS Classification

Health Hazards:
3.3 Serious eye damage/eye irritation (Category 2A)
H319 Causes serious eye irritation.

Signalword: Warning
Label:

Precautionary statements:
- P280 Wear protective gloves/ protective clothing / eye protection / face protection.
- P312 Call a POISON CENTER/doctor if you feel unwell.
- P273 Avoid release to the environment.

Other hazards

Note
- no information available

SECTION 3: Composition/information on ingredients

Characterization
Entrectinib with other inactive ingredients

Ingredients

<table>
<thead>
<tr>
<th>Ingredients</th>
<th>Concentration</th>
<th>GHS-Classification (pure ingredient)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Entrectinib 1108743-60-7</td>
<td>35.1 %</td>
<td>- Serious eye damage/eye irritation (Category 2A), H319</td>
</tr>
<tr>
<td>Lactose anhydrous 63-42-3</td>
<td>22.8 %</td>
<td></td>
</tr>
<tr>
<td>Tartaric acid fine cryst. 87-69-4</td>
<td>10.4 %</td>
<td>- Serious eye damage/eye irritation (Category 2A), H319</td>
</tr>
<tr>
<td>Polyvinylpyrrolidone, cross-linked, micronised 9003-39-8</td>
<td>4.2 %</td>
<td></td>
</tr>
<tr>
<td>2-Hydroxypropyl methyl ether cellulose 9004-65-3</td>
<td>3.2 %</td>
<td></td>
</tr>
<tr>
<td>Microcrystalline cellulose 9004-34-6</td>
<td>2.4 %</td>
<td></td>
</tr>
</tbody>
</table>

For the full text of the ‘Hazard statements’ mentioned in this Section, see Section 16.
### SECTION 4: First aid measures

**4.1. Description of first aid measures**

- **Eye contact**
  - rinse with tap water for 20 minutes - open eyelids forcibly
- **Skin contact**
  - when in contact with the skin, clean with soap and water
- **Inhalation**
  - in the event of symptoms get medical treatment

**4.2. Most important symptoms and effects, both acute and delayed**

- **Note**
  - no information available

**4.3. Indication of any immediate medical attention and special treatment needed**

- **Note to physician**
  - treat symptomatically

### SECTION 5: Firefighting measures

**5.1. Extinguishing media**

- **Suitable extinguishing media**
  - adapt extinguishing media to surrounding fire conditions, water spray jet, dry powder, foam, carbon dioxide
- **Flash point (liquid)**
  - not applicable

**5.2. Special hazards arising from the substance or mixture**

- **Specific hazards**
  - substance is hazardous for water: contain fire-fighting wastewater

**5.3. Advice for firefighters**

- **Protection of fire-fighters**
  - precipitate gases/vapours/mists with water spray

### SECTION 6: Accidental release measures

**6.1. Personal precautions, protective equipment and emergency procedures**

- **Personal precautions**
  - no special precautions required

**6.2. Environmental precautions**

- **Environmental protection**
  - the solvent should be held back due to environmental protection
  - if the substance reaches waters or the sewer system, inform the competent authority
6.3. Methods and material for containment and cleaning up

Methods for cleaning up - take up mechanically and dispose of

SECTION 7: Handling and storage

7.1. Precautions for safe handling

Technical measures - avoid dust formation
  - processing in closed systems, if possible superposed by inert gas (e.g. nitrogen)

7.2. Conditions for safe storage, including any incompatibilities

Storage conditions - room temperature
Validity - see expiry date on the label
Packaging materials - tightly closing

SECTION 8: Exposure controls/personal protection

8.1. Control parameters

Threshold value (Roche) air - IOEL (Internal Occupational Exposure Limit): 0.015 mg/m³
PNEC - 0.3 µg/l, surface freshwaters, based on chronic data

8.2. Exposure controls

Respiratory protection - Respiratory protection is recommended as a precaution to minimize exposure. Effective engineering controls are considered to be the primary means to control worker exposure. Respiratory protection should not substitute for feasible engineering controls.
  - respiratory protection not necessary during normal operations
Hand protection - protective gloves (e.g. made of neoprene, nitrile or butyl rubber)
Eye protection - safety glasses

*1 referring to: Entrectinib

SECTION 9: Physical and chemical properties

9.1. Information on basic physical and chemical properties

Color yellow
Form hard gelatin capsules
### Solubility

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<thead>
<tr>
<th>Condition</th>
<th>Solubility</th>
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<tr>
<td>≥0.964 mg/l, water (20 °C, pH 7.3, OECD No. 105)</td>
<td>≥0.964 mg/l, water (20 °C, pH 7.3, OECD No. 105)</td>
</tr>
<tr>
<td>≤2.94 mg/l, water (20 °C, pH 8.7, OECD No. 105)</td>
<td>≤2.94 mg/l, water (20 °C, pH 8.7, OECD No. 105)</td>
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### Partition coefficient

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<th>Condition</th>
<th>Partition Coefficient</th>
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<tr>
<td>log (P_{ow}) 2.7 (octanol/water 20 °C) pH 5.0</td>
<td>log (P_{ow}) 2.7 (octanol/water 20 °C) pH 5.0</td>
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<tr>
<td>(Shake Flask Method, OECD No. 107)</td>
<td>(Shake Flask Method, OECD No. 107)</td>
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<tr>
<td>log (P_{ow}) 4.3 (octanol/water 20 °C) pH 7.0</td>
<td>log (P_{ow}) 4.3 (octanol/water 20 °C) pH 7.0</td>
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<tr>
<td>(Shake Flask Method, OECD No. 107)</td>
<td>(Shake Flask Method, OECD No. 107)</td>
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<tr>
<td>log (P_{ow}) 5.1 (octanol/water 20 °C) pH 9.0</td>
<td>log (P_{ow}) 5.1 (octanol/water 20 °C) pH 9.0</td>
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<td>(Shake Flask Method, OECD No. 107)</td>
<td>(Shake Flask Method, OECD No. 107)</td>
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<td>(OECD No. 123)</td>
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### 9.2. Other information

- **Note** - no information available

*1 referring to: Entrectinib

### SECTION 10: Stability and reactivity

#### 10.1. Reactivity

- **Note** - no information available

#### 10.2. Chemical stability

- **Stability** - stable under the conditions mentioned in chapter 7

#### 10.3. Possibility of hazardous reactions

- **Note** - no information available

#### 10.4. Conditions to avoid

- **Note** - no information available

#### 10.5. Incompatible materials

- **Note** - no information available

#### 10.6. Hazardous decomposition products

- **Note** - no information available

### SECTION 11: Toxicological information

#### 11.1. Information on toxicological effects

- **Acute toxicity** - LD\(_{50}\) > 2'000 mg/kg (oral, rat)
Local effects
- skin: non-irritant (rabbit)  
- eye: slightly irritating (rabbit)  
- phototoxic potential (In Vitro)  

Sensitization
- no information available  

Mutagenicity
- negative (various in vivo and in vitro test systems)  

Carcinogenicity
- no information available  

Reproductive toxicity
- may lead to malformations at dose levels that cause maternal toxicity (rat)  

STOT-single exposure
- no information available  

STOT-repeated exposure
- no information available  

Aspiration hazard
- no information available  

Potential Health Effects
- Exposure: Inhalation, Ingestion, Skin contact, Eye contact  
- Carcinogenicity: not listed by NTP, IARC or OSHA  

*1 referring to: Entrectinib

SECTION 12: Ecological information

12.1. Toxicity

Ecotoxicity
- Desmodesmus subspicatus
  NOEC (72 h) 0.197 mg/l  
  ErC50 (72 h) > 100 mg/l (nominal concentration)  
  EyC50 (72 h) > 100 mg/l (nominal concentration)  
  (OECD No. 201)  
- Daphnia magna
  NOEC (48 h) 0.266 mg/l  
  EC50 (48 h) > 100 mg/l (nominal concentration)  
  (OECD No. 202)  
- Daphnia magna
  EC10 (21 d) 0.0645 mg/l (average measured concentration)  
  (OECD No. 211 (semi-static))  
- zebrafish
  EC10 (35 d) 0.00311 mg/l (average measured concentration)  
  (OECD No. 210)  
- activated sludge
  NOEC (3 h) 1000 mg/l (nominal concentration)  
  (OECD No. 209)  
- no adverse influence on substrate biodegradation (activated sludge)
  concentration (14 d) 38.0 mg/l  
  (Manometric Respirometry Test, OECD No. 301 F)  

*1
12.2. Persistence and degradability

Ready biodegradability - not readily biodegradable
< 10 %, 28 d
(Manometric Respirometry Test, OECD No. 301 F) *1

Abiotic degradation - not abiotically degradable (Manometric respirometry test, OECD no. 301 F, abiotic control) *1

12.3. Bioaccumulative potential

Note - no information available

12.4. Mobility in soil

Note - no information available

12.5. Results of PBT and vPvB assessment

Note - no information available

12.6. Other adverse effects

Note - no information available

*1 referring to: Entrectinib

SECTION 13: Disposal considerations

13.1. Waste treatment methods

Waste from residues - return to supplier or hand over to authorized disposal company
- observe local/national regulations regarding waste disposal
- incinerate in qualified installation with flue gas scrubbing
- DO NOT FLUSH unused medications or POUR them down a sink or drain. If available in your area, use takeback programs run by household hazardous waste collection programs or community pharmacies to dispose of unused and expired medicines. If you don’t have access to a takeback program, dispose of these medicines in the household trash by removing them from their original containers and mixing them with an undesirable substance, such as used coffee grounds or kitty litter.

SECTION 14: Transport information

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ROZLYTREK™ Capsules (100 mg)

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</table>

Proper shipping name  ENVIRONMENTALLY HAZARDOUS SUBSTANCE, SOLID, N.O.S.

Technical name  Entrectinib

SECTION 15: Regulatory information

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

TSCA Status  - FDA Exemption - not on inventory

Reporting Requirements  - The United States Environmental Protection Agency (USEPA) has not established a Reportable Quantity (RQ) for releases of this material.
- In New Jersey, report all releases which enter a waterway or into soil, or which are likely to endanger the public health, harm the environment or cause a complaint to the NJDEPE Hotline and to local officials.
- State and local regulations vary and may impose additional reporting requirements.

SECTION 16: Other information

Safety-lab number  - BS11425
- BS11450

Full text of H-Statements referred to under section 3
H319  Causes serious eye irritation.

Note  - none

Edition documentation  - changes from previous version in sections 2, 3, 14

*1 referring to:  Entrectinib

The information in this safety data sheet is based on current scientific knowledge. It should not be taken as expressing or implying any warranty concerning product characteristics.