SAFETY DATA SHEET

COTELLIC(R) Tablets (20 mg)

SECTION 1. IDENTIFICATION

Product name : COTELLIC(R) Tablets (20 mg)
Product code : RO551-4041/F04
Common name(s), synonym(s) of the substance : BS10010, BS10336, BS10337
Cobimetinib F.C. Tablets 20 mg
Cobimetinib Film Coated Tablets 20 mg
Cobimetinib Tablets 20 mg

Manufacturer or supplier's details
Company name of supplier : Genentech, Inc.
Address : 1 DNA Way
South San Francisco, CA 94080
USA
Telephone : 001-(650) 225-1000
E-mail address : info.sds@roche.com
Emergency telephone : US Chemtrec phone (800)-424-9300

Recommended use of the chemical and restrictions on use
Recommended use : Formulated pharmaceutical active substance
Restrictions on use : For professional users only.

SECTION 2. HAZARDS IDENTIFICATION

GHS classification in accordance with the OSHA Hazard Communication Standard (29 CFR 1910.1200)
Acute toxicity (Oral) : Category 4
Carcinogenicity : Category 1A
Reproductive toxicity : Category 1B

GHS label elements
Hazard pictograms : 🚨 🚨

Signal Word : Danger
Hazard Statements : H302 Harmful if swallowed.
H350 May cause cancer.
H360D May damage the unborn child.
Precautionary Statements:

Prevention:
P201 Obtain special instructions before use.
P202 Do not handle until all safety precautions have been read and understood.
P264 Wash skin thoroughly after handling.
P270 Do not eat, drink or smoke when using this product.
P280 Wear protective gloves/ protective clothing/ eye protection/ face protection.

Response:
P301 + P312 + P330 IF SWALLOWED: Call a POISON CENTER/ doctor if you feel unwell. Rinse mouth.
P308 + P313 IF exposed or concerned: Get medical advice/ attention.

Storage:
P405 Store locked up.

Disposal:
P501 Dispose of contents/ container to an approved waste disposal plant.

Other hazards
None known.

SECTION 3. COMPOSITION/INFORMATION ON INGREDIENTS

Substance / Mixture: Mixture

<table>
<thead>
<tr>
<th>Components</th>
<th>Chemical name</th>
<th>CAS-No.</th>
<th>Concentration (% w/w)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Cobimetinib</td>
<td>1369665-02-0</td>
<td>17.8</td>
<td></td>
</tr>
<tr>
<td>Cellulose</td>
<td>9004-34-6</td>
<td>43.9</td>
<td></td>
</tr>
<tr>
<td>Lactose-1-Hydrate</td>
<td>64044-51-5</td>
<td>29.2</td>
<td></td>
</tr>
<tr>
<td>Croscarmellose sodium</td>
<td>74811-65-7</td>
<td>3.9</td>
<td></td>
</tr>
<tr>
<td>Ethenol, homopolymer</td>
<td>9002-89-5</td>
<td>1.5</td>
<td></td>
</tr>
<tr>
<td>Octadecanoic acid, magnesium salt (2:1)</td>
<td>557-04-0</td>
<td>1.4</td>
<td></td>
</tr>
<tr>
<td>Titanium oxide (TiO2)</td>
<td>13463-67-7</td>
<td>&lt; 1.0</td>
<td></td>
</tr>
<tr>
<td>Poly(oxy-1,2-ethanediyl), .alpha.-hydro-.omega.-hydroxy-</td>
<td>25322-68-3</td>
<td>0.8</td>
<td></td>
</tr>
<tr>
<td>Talc (Mg3H2(SiO3)4)</td>
<td>14807-96-6</td>
<td>0.6</td>
<td></td>
</tr>
</tbody>
</table>

SECTION 4. FIRST AID MEASURES

General advice: Move out of dangerous area.
Show this material safety data sheet to the doctor in attendance.
Do not leave the victim unattended.

If inhaled: Move to fresh air.
If unconscious, place in recovery position and seek medical
advice.
If symptoms persist, call a physician.

In case of skin contact: If on skin, rinse well with water.

In case of eye contact:
 Immediately flush eye(s) with plenty of water.
 Remove contact lenses.
 Protect unharmed eye.
 Keep eye wide open while rinsing.
 If eye irritation persists, consult a specialist.

If swallowed:
 Keep respiratory tract clear.
 Do not give milk or alcoholic beverages.
 Never give anything by mouth to an unconscious person.
 If symptoms persist, call a physician.
 Take victim immediately to hospital.
 Rinse mouth with water.

Most important symptoms and effects, both acute and delayed:
 Harmful if swallowed.
 May cause cancer.
 May damage the unborn child.

Notes to physician:
 The first aid procedure should be established in consultation with the doctor responsible for industrial medicine.

SECTION 5. FIRE-FIGHTING MEASURES

Suitable extinguishing media: Use extinguishing measures that are appropriate to local circumstances and the surrounding environment.

Unsuitable extinguishing media: High volume water jet

Specific hazards during fire fighting: Do not allow run-off from fire fighting to enter drains or water courses.

Hazardous combustion products: Carbon oxides
 In case of fire hazardous decomposition products may be produced such as:
 Hydrogen fluoride
 Nitrogen oxides (NOx)

Further information:
 Collect contaminated fire extinguishing water separately. This must not be discharged into drains.
 Fire residues and contaminated fire extinguishing water must be disposed of in accordance with local regulations.

Special protective equipment for fire-fighters:
 Wear self-contained breathing apparatus for firefighting if necessary.

SECTION 6. ACCIDENTAL RELEASE MEASURES

Personal precautions, protective equipment and:
 Avoid exposure
 Use personal protective equipment.
emergency procedures
Avoid dust formation.
Avoid breathing dust.

Environmental precautions
Prevent product from entering drains.
Prevent further leakage or spillage if safe to do so.
If the product contaminates rivers and lakes or drains inform respective authorities.

Methods and materials for containment and cleaning up
Keep in suitable, closed containers for disposal.

SECTION 7. HANDLING AND STORAGE

Advice on protection against fire and explosion
Avoid dust formation.
Provide appropriate exhaust ventilation at places where dust is formed.

Advice on safe handling
Avoid formation of respirable particles.
Do not breathe vapors/dust.
Avoid exposure - obtain special instructions before use.
Avoid contact with skin and eyes.
For personal protection see section 8.
Smoking, eating and drinking should be prohibited in the application area.
Dispose of rinse water in accordance with local and national regulations.

Conditions for safe storage
Keep container tightly closed in a dry and well-ventilated place.
Containers which are opened must be carefully resealed and kept upright to prevent leakage.
Electrical installations / working materials must comply with the technological safety standards.

Further information on storage conditions
See label, package insert or internal guidelines

Storage temperature
Store at room temperature.
Protected from heat and light
Protect from moisture.

Further information on storage stability
No decomposition if stored and applied as directed.

Packaging material
Suitable material: glass, Stainless steel

SECTION 8. EXPOSURE CONTROLS/PERSOAL PROTECTION

Ingredients with workplace control parameters

<table>
<thead>
<tr>
<th>Components</th>
<th>CAS-No.</th>
<th>Value type (Form of)</th>
<th>Control parameters / Basis</th>
</tr>
</thead>
</table>
## Predicted No Effect Concentration (PNEC):

<table>
<thead>
<tr>
<th>Substance name</th>
<th>Environmental Compartment</th>
<th>Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Cobimetinib</td>
<td>Surface waters</td>
<td>8.98 µg/l</td>
</tr>
</tbody>
</table>

Remarks: Based on chronic data

## Engineering measures

- No data available

## Personal protective equipment

- **Respiratory protection**: In the case of dust or aerosol formation use respirator with an
Hand protection

In case of contact through splashing:

- Material: Nitrile rubber
- Break through time: > 30 min
- Glove thickness: > 0.11 mm

In case of full contact:

- Material: butyl-rubber
- Break through time: > 480 min
- Glove thickness: > 0.4 mm

Remarks:
Wear appropriate protective gloves to prevent skin contact.
Replace torn or punctured gloves promptly.

Eye protection:
Eye wash bottle with pure water
Tightly fitting safety goggles

Skin and body protection:
Dust impervious protective suit
Choose body protection according to the amount and concentration of the dangerous substance at the work place.

Protective measures:
Instruction of employees mandatory

Hygiene measures:
When using do not eat or drink.
When using do not smoke.
Wash hands before breaks and at the end of workday.

SECTION 9. PHYSICAL AND CHEMICAL PROPERTIES

Appearance: tablet

Color: white

Odor: Not applicable

Odor Threshold: Not applicable

pH: No data available

Melting point/freezing point: No data available

Boiling point/boiling range: No data available

Evaporation rate: No data available

Self-ignition: No data available
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Date of last issue: 01-22-2020
Date of first issue: 12-07-2015

- Upper explosion limit / Upper flammability limit: No data available
- Lower explosion limit / Lower flammability limit: No data available
- Vapor pressure: No data available
- Relative vapor density: Not applicable
- Relative density: No data available
- Solubility(ies)
  - Water solubility: No data available
  - Solubility in other solvents: No data available
- Partition coefficient: n-octanol/water: No data available
- Autoignition temperature: No data available
- Decomposition temperature: No data available
- Viscosity
  - Viscosity, dynamic: Not applicable
  - Viscosity, kinematic: Not applicable

SECTION 10. STABILITY AND REACTIVITY

- Reactivity: No dangerous reaction known under conditions of normal use.
- Chemical stability: Stable under normal conditions.
- Possibility of hazardous reactions: No decomposition if stored and applied as directed.
- Incompatible materials: No data available
- Hazardous decomposition products: No data available

SECTION 11. TOXICOLOGICAL INFORMATION

Acute toxicity
Harmful if swallowed.

Product:
Acute oral toxicity: Acute toxicity estimate: 560.3 mg/kg
  Method: Calculation method
Acute dermal toxicity: Acute toxicity estimate: > 5,000 mg/kg
  Method: Calculation method
Components:

Cellulose:
Acute oral toxicity : LD50 Oral (Rat): > 5,000 mg/kg
Acute dermal toxicity : LD50 Dermal (Rabbit): > 2,000 mg/kg

Lactose-1-Hydrate:
Acute oral toxicity : LD50 Oral (Rat): > 10,000 mg/kg
Acute inhalation toxicity : Acute toxicity estimate: > 30 mg/l
Test atmosphere: dust/mist
Method: Expert judgment
Acute dermal toxicity : Acute toxicity estimate: > 5,001 mg/kg
Method: Expert judgment

Cobimetinib:
Acute oral toxicity : LD50 Oral (Rat): > 60 - < 75 mg/kg
GLP: yes
The value is given in analogy to the following substances:
Cobimetinib (free base)
NOAEL (No observed adverse effect level) (Rat): 30 mg/kg
GLP: yes
The value is given in analogy to the following substances:
Cobimetinib (free base)

Ethenol, homopolymer:
Acute oral toxicity : LD50 Oral (Rat): > 20,000 mg/kg
Acute inhalation toxicity : Acute toxicity estimate: > 30 mg/l
Test atmosphere: dust/mist
Method: Expert judgment
Acute dermal toxicity : Acute toxicity estimate: > 5,001 mg/kg
Method: Expert judgment

Octadecanoic acid, magnesium salt (2:1):
Acute oral toxicity : LD50 Oral (Rat): > 2,000 mg/kg

Titanium oxide (TiO2):
Acute oral toxicity : LD50 (Rat): > 5,000 mg/kg
Method: OECD Test Guideline 425
Acute inhalation toxicity : LC50 (Rat): > 6.82 mg/l
Exposure time: 4 h
Test atmosphere: dust/mist
Acute dermal toxicity : LD50 (Rabbit): > 5,000 mg/kg
Skin corrosion/irritation
Not classified based on available information.

**Components:**

**Cobimetinib:**
- Method: in silico model
- Result: No skin irritation
The value is given in analogy to the following substances: Cobimetinib (free base)

**Ethenol, homopolymer:**
- Remarks: This information is not available.

**Titanium oxide (TiO2):**
- Species: Rabbit
- Method: OECD Test Guideline 404
- Result: No skin irritation

**Talc (Mg3H2(SiO3)4):**
- Remarks: This information is not available.

**Serious eye damage/eye irritation**
Not classified based on available information.

**Components:**

**Ethenol, homopolymer:**
- Remarks: This information is not available.

**Titanium oxide (TiO2):**
- Species: Rabbit
- Result: No eye irritation
- Method: OECD Test Guideline 405

**Talc (Mg3H2(SiO3)4):**
- Remarks: This information is not available.

**Respiratory or skin sensitization**

**Skin sensitization**
Not classified based on available information.

**Respiratory sensitization**
Not classified based on available information.

**Components:**

**Titanium oxide (TiO2):**
- Species: Guinea pig
- Assessment: Does not cause skin sensitization.
- Method: OECD Test Guideline 406
**SAFETY DATA SHEET**

**COTELLIC(R) Tablets (20 mg)**

<table>
<thead>
<tr>
<th>Version</th>
<th>Revision Date:</th>
<th>Date of last issue:</th>
<th>Date of first issue:</th>
</tr>
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<tbody>
<tr>
<td>1.5</td>
<td>10-27-2021</td>
<td>01-22-2020</td>
<td>12-07-2015</td>
</tr>
</tbody>
</table>

**Germ cell mutagenicity**
Not classified based on available information.

**Components:**

**Cobimetinib:**
- **Genotoxicity in vitro:** Result: negative
  - Remarks: In vitro tests did not show mutagenic effects
  - The value is given in analogy to the following substances: Cobimetinib (free base)

**Carcinogenicity**
May cause cancer.

**Components:**

**Cellulose:**
- **Remarks:** No ingredient of this product present at levels greater than or equal to 0.1% is identified as probable, possible or confirmed human carcinogen by IARC.

**Lactose-1-Hydrate:**
- **Remarks:** No ingredient of this product present at levels greater than or equal to 0.1% is identified as probable, possible or confirmed human carcinogen by IARC.

**IARC**
- Group 2B: Possibly carcinogenic to humans
- Titanium oxide (TiO2) 13463-67-7

**OSHA**
No component of this product present at levels greater than or equal to 0.1% is on OSHA’s list of regulated carcinogens.

**NTP**
- Known to be human carcinogen
- Talc (Mg3H2(SiO3)4) (Silica, Crystalline (Respirable Size)) 14807-96-6

**Reproductive toxicity**
May damage the unborn child.

**Components:**

**Cobimetinib:**
- **Reproductive toxicity - Assessment:** May damage the unborn child., Presumed human reproductive toxicant
  - The value is given in analogy to the following substances: Cobimetinib (free base)

**STOT-single exposure**
Not classified based on available information.
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Components:
Lactose-1-Hydrate:
Assessment: The substance or mixture is not classified as specific target organ toxicant, single exposure.

Ethenol, homopolymer:
Assessment: The substance or mixture is not classified as specific target organ toxicant, single exposure.

Octadecanoic acid, magnesium salt (2:1):
Assessment: The substance or mixture is not classified as specific target organ toxicant, single exposure.

Talc (Mg3H2(SiO3)4):
Assessment: The substance or mixture is not classified as specific target organ toxicant, single exposure.

STOT-repeated exposure
Not classified based on available information.

Components:
Lactose-1-Hydrate:
Assessment: The substance or mixture is not classified as specific target organ toxicant, repeated exposure.

Ethenol, homopolymer:
Assessment: The substance or mixture is not classified as specific target organ toxicant, repeated exposure.

Octadecanoic acid, magnesium salt (2:1):
Assessment: The substance or mixture is not classified as specific target organ toxicant, repeated exposure.

Talc (Mg3H2(SiO3)4):
Assessment: The substance or mixture is not classified as specific target organ toxicant, repeated exposure.

Repeated dose toxicity
Components:
Cobimetinib:
Species: Rat
NOAEL: 1 mg/kg bw/day
Application Route: Oral
Exposure time: 28 Days
Test substance: yes
Remarks: Subacute toxicity
The value is given in analogy to the following substances: Cobimetinib (free base)
SAFETY DATA SHEET

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Species : Rat
LOAEL : >3 mg/kg bw/day
Application Route : Oral
Exposure time : 13 Weeks
Test substance : yes
Remarks : Subchronic toxicity
The value is given in analogy to the following substances: Cobimetinib (free base)

Aspiration toxicity
Not classified based on available information.

Components:

Lactose-1-Hydrate:
No data available

Ethenol, homopolymer:
No data available

Octadecanoic acid, magnesium salt (2:1):
No data available

Talc (Mg3H2(SiO3)4):
No data available

Further information

Components:

Cobimetinib:
Remarks : Not phototoxic
The value is given in analogy to the following substances: Cobimetinib (free base)

SECTION 12. ECOLOGICAL INFORMATION

Ecotoxicity

Components:

Cellulose:

Ecotoxicology Assessment
Acute aquatic toxicity : This product has no known ecotoxicological effects.
Chronic aquatic toxicity : This product has no known ecotoxicological effects.

Lactose-1-Hydrate:
Toxicity to fish : LC50: > 100 mg/l
Exposure time: 96 h

Toxicity to fish (Chronic toxicity): > 1 mg/l

Ecotoxicology Assessment
Acute aquatic toxicity: This product has no known ecotoxicological effects.
Chronic aquatic toxicity: This product has no known ecotoxicological effects.
Toxicity Data on Soil: Not expected to adsorb on soil.
Other organisms relevant to the environment: No data available

Cobimetinib:
Toxicity to fish: LC50 (Danio rerio (zebra fish)): 0.80 mg/l
   End point: mortality
   Exposure time: 96 h
   Method: OECD Test Guideline 203
   GLP: yes
   The value is given in analogy to the following substances:
   Cobimetinib (free base)
   NOEC (Danio rerio (zebra fish)): 0.43 mg/l
   End point: mortality
   Exposure time: 96 h
   Method: OECD Test Guideline 203
   GLP: yes
   The value is given in analogy to the following substances:
   Cobimetinib (free base)

Toxicity to daphnia and other aquatic invertebrates: EC50 (Daphnia magna (Water flea)): 3.54 mg/l
   End point: Immobilization
   Exposure time: 48 h
   Analytical monitoring: yes
   Method: OECD Test Guideline 202
   GLP: yes
   Remarks: average measured concentration
   The value is given in analogy to the following substances:
   Cobimetinib (free base)
   NOEC (Daphnia magna (Water flea)): 2.00 mg/l
   End point: Immobilization
   Exposure time: 48 h
   Analytical monitoring: yes
   Method: OECD Test Guideline 202
   GLP: yes
   Remarks: average measured concentration
   The value is given in analogy to the following substances:
   Cobimetinib (free base)

Toxicity to algae/aquatic plants: ErC50 (Desmodesmus subspicatus (green algae)): 11.8 mg/l
   Exposure time: 72 h
Analytical monitoring: yes
Method: OECD Test Guideline 201
GLP: yes
Remarks: average measured concentration
The value is given in analogy to the following substances:
Cobimetinib (free base)

EyC50 (Desmodesmus subspicatus (green algae)): 2.25 mg/l
Exposure time: 72 h
Analytical monitoring: yes
Method: OECD Test Guideline 201
GLP: yes
Remarks: average measured concentration
The value is given in analogy to the following substances:
Cobimetinib (free base)

NOErC (Desmodesmus subspicatus (green algae)): < 0.62 mg/l
Exposure time: 72 h
Analytical monitoring: yes
Method: OECD Test Guideline 201
GLP: yes
Remarks: average measured concentration
The value is given in analogy to the following substances:
Cobimetinib (free base)

NOEyC (Desmodesmus subspicatus (green algae)): < 0.62 mg/l
Exposure time: 72 h
Analytical monitoring: yes
Method: OECD Test Guideline 201
GLP: yes
Remarks: average measured concentration
The value is given in analogy to the following substances:
Cobimetinib (free base)

Toxicity to fish (Chronic toxicity) : NOEC (Danio rerio (zebra fish)): >= 109 mg/l
Exposure time: 35 d
Method: OECD Test Guideline 210
GLP: yes
The value is given in analogy to the following substances:
Cobimetinib (free base)

Toxicity to daphnia and other aquatic invertebrates (Chronic toxicity) : NOEC (Daphnia magna (Water flea)): 0.0898 mg/l
End point: Immobilization
Exposure time: 21 d
Method: OECD Test Guideline 211
GLP: yes
Remarks: average measured concentration
The value is given in analogy to the following substances:
Cobimetinib (free base)

Toxicity to microorganisms : EC50 (activated sludge): > 1,000 mg/l
Exposure time: 3 h
Test Type: Respiration inhibition
Method: OECD Test Guideline 209
GLP: yes
Remarks: nominal concentration
The value is given in analogy to the following substances:
Cobimetinib (free base)

(activated sludge): 57.5 mg/l
Exposure time: 28 d
Method: OECD Test Guideline 301F
Remarks: no adverse influence on substrate biodegradation measured initial concentration
The value is given in analogy to the following substances:
Cobimetinib (free base)

Ethenol, homopolymer:
Toxicity to fish: LC50: > 100 mg/l
Exposure time: 96 h
Toxicity to fish (Chronic toxicity): > 1 mg/l

Ecotoxicology Assessment
Acute aquatic toxicity: This product has no known ecotoxicological effects.
Chronic aquatic toxicity: This product has no known ecotoxicological effects.
Toxicity Data on Soil: Not expected to adsorb on soil.
Other organisms relevant to the environment: No data available

Octadecanoic acid, magnesium salt (2:1):
Ecotoxicology Assessment
Acute aquatic toxicity: This product has no known ecotoxicological effects.
Chronic aquatic toxicity: This product has no known ecotoxicological effects.
Toxicity Data on Soil: Not expected to adsorb on soil.
Other organisms relevant to the environment: No data available

Titanium oxide (TiO2):
Toxicity to fish: LC50 (Pimephales promelas (fathead minnow)): > 1,000 mg/l
Exposure time: 96 h
Test Type: static test
LC50 (Cyprinodon variegatus (sheepshead minnow)): > 10,000 mg/l
Exposure time: 96 h
Test Type: semi-static test
Method: OECD Test Guideline 203
Toxicity to daphnia and other aquatic invertebrates:
LC50 (Daphnia magna (Water flea)): > 1,000 mg/l
Exposure time: 48 h
Test Type: static test
Method: OECD Test Guideline 202

Toxicity to algae/aquatic plants:
EC50 (Pseudokirchneriella subcapitata (green algae)): > 100 mg/l
Exposure time: 72 h
Test Type: static test
Method: OECD Test Guideline 201

EC50 (Skeletonema costatum (marine diatom)): > 10,000 mg/l
Exposure time: 72 h
Method: ISO 10253

NOEC (Skeletonema costatum (marine diatom)): 5,600 mg/l
Exposure time: 72 h
Method: ISO 10253

Ecotoxicology Assessment:
Toxicity Data on Soil: Not expected to adsorb on soil.

Other organisms relevant to the environment: No data available

Talc (Mg3H2(SiO3)4):
Toxicity to fish:
LC50 (Danio rerio (zebra fish)): > 100,000 mg/l
Exposure time: 24 h

Ecotoxicology Assessment:
Acute aquatic toxicity: This product has no known ecotoxicological effects.
Chronic aquatic toxicity: This product has no known ecotoxicological effects.

Toxicity Data on Soil: Not expected to adsorb on soil.

Other organisms relevant to the environment: No data available

Persistence and degradability:

Components:

Cobimetinib:
Biodegradability: aerobic
Inoculum: activated sludge
Concentration: 60.8 mg/l
Theoretical oxygen demand
Result: Not readily biodegradable.
Biodegradation: 0 %
Exposure time: 28 Days
Method: OECD Test Guideline 301F
GLP: yes
The value is given in analogy to the following substances:
Cobimetinib (free base)

Titanium oxide (TiO2):
Biodegradability : Remarks: Not applicable

Bioaccumulative potential

Components:

Cellulose:
Partition coefficient: n-octanol/water : Remarks: No data available

Lactose-1-Hydrate:
Partition coefficient: n-octanol/water : log Pow: -5.03

Cobimetinib:
Partition coefficient: n-octanol/water : log Pow: 0.32

Polyvinyl alcohol:
Partition coefficient: n-octanol/water : Remarks: No data available

Magnesium stearate:
Partition coefficient: n-octanol/water : log Pow: 0.8
Method: OECD Test Guideline 107

Titanium dioxide:
Partition coefficient: n-octanol/water : Remarks: No data available

Talc:
Partition coefficient: n-octanol/water : Remarks: No data available

Mobility in soil

Components:

Cobimetinib:
Distribution among environmental compartments : Medium: Soil
Method: OECD Test Guideline 106
Remarks: immobile

Medium: Sludge
Method: OECD Test Guideline 106
Remarks: immobile
Other adverse effects

Product:
Ozone-Depletion Potential: Regulation: 40 CFR Protection of Environment; Part 82 Protection of Stratospheric Ozone - CAA Section 602 Class I Substances
Remarks: This product neither contains, nor was manufactured with a Class I or Class II ODS as defined by the U.S. Clean Air Act Section 602 (40 CFR 82, Subpt. A, App.A + B).

Additional ecological information: An environmental hazard cannot be excluded in the event of unprofessional handling or disposal.
Toxic to aquatic life with long lasting effects.

Components:
Ethenol, homopolymer:
Adsorbed organic bound halogens (AOX): Remarks: Not applicable
Additional ecological information: No data available

Talc (Mg3H2(SiO3)4):
Adsorbed organic bound halogens (AOX): Remarks: Not applicable
Additional ecological information: No data available

SECTION 13. DISPOSAL CONSIDERATIONS

Disposal methods
Waste from residues: The product should not be allowed to enter drains, water courses or the soil.
Do not contaminate ponds, waterways or ditches with chemical or used container.
Send to a licensed waste management company.

Contaminated packaging: Empty remaining contents.
Dispose of as unused product.
Empty containers should be taken to an approved waste handling site for recycling or disposal.
Do not re-use empty containers.

SECTION 14. TRANSPORT INFORMATION

International Regulations
SAFETY DATA SHEET

COTELLIC(R) Tablets (20 mg)

Version: 1.5
Revision Date: 10-27-2021
Date of last issue: 01-22-2020
Date of first issue: 12-07-2015

UNRTDG
UN number: UN 3077
Proper shipping name:_ENVIRONMENTALLY HAZARDOUS SUBSTANCE, SOLID, N.O.S._ (Cobimetinib, mixture)

Class: 9
Packing group: III
Labels: 9

IATA-DGR
UN/ID No.: UN 3077
Proper shipping name: Environmentally hazardous substance, solid, n.o.s. (Cobimetinib, mixture)

Class: 9
Packing group: III
Labels: Miscellaneous
Packing instruction (cargo aircraft): 956
Packing instruction (passenger aircraft): 956
Environmentally hazardous: yes

IMDG-Code
UN number: UN 3077
Proper shipping name: _ENVIRONMENTALLY HAZARDOUS SUBSTANCE, SOLID, N.O.S._ (Cobimetinib, mixture)

Class: 9
Packing group: III
Labels: 9
EmS Code: F-A, S-F
Marine pollutant: yes

Transport in bulk according to Annex II of MARPOL 73/78 and the IBC Code
Not applicable

Domestic regulation

49 CFR
UN/ID/NA number: UN 3077
Proper shipping name: Environmentally hazardous substance, solid, n.o.s. (Cobimetinib, mixture)

Class: 9
Packing group: III
Labels: CLASS 9
ERG Code: 171
Marine pollutant: no

Special precautions for user
The transport classification(s) provided herein are for informational purposes only, and solely based upon the properties of the unpackaged material as it is described within this Safety Data Sheet. Transportation classifications may vary by mode of transportation, package sizes, and variations in regional or country regulations.
SECTION 15. REGULATORY INFORMATION

CERCLA Reportable Quantity
This material does not contain any components with a CERCLA RQ.

SARA 304 Extremely Hazardous Substances Reportable Quantity
This material does not contain any components with a section 304 EHS RQ.

SARA 302 Extremely Hazardous Substances Threshold Planning Quantity
This material does not contain any components with a section 302 EHS TPQ.

SARA 311/312 Hazards
- Acute toxicity (any route of exposure)
- Carcinogenicity
- Reproductive toxicity

SARA 313
- This material does not contain any chemical components with known CAS numbers that exceed the threshold (De Minimis) reporting levels established by SARA Title III, Section 313.

Clean Air Act
This product neither contains, nor was manufactured with a Class I or Class II ODS as defined by the U.S. Clean Air Act Section 602 (40 CFR 82, Subpt. A, App.A + B).
This product does not contain any hazardous air pollutants (HAP), as defined by the U.S. Clean Air Act Section 112 (40 CFR 61).
This product does not contain any chemicals listed under the U.S. Clean Air Act Section 112(r) for Accidental Release Prevention (40 CFR 68.130, Subpart F).
This product does not contain any chemicals listed under the U.S. Clean Air Act Section 111 SOCMI Intermediate or Final VOC's (40 CFR 60.489).

Clean Water Act
This product does not contain any Hazardous Substances listed under the U.S. CleanWater Act, Section 311, Table 116.4A.
This product does not contain any Hazardous Chemicals listed under the U.S. CleanWater Act, Section 311, Table 117.3.
This product does not contain any toxic pollutants listed under the U.S. Clean Water Act Section 307.
This product does not contain any priority pollutants related to the U.S. Clean Water Act.

US State Regulations

Massachusetts Right To Know
- Cellulose 9004-34-6

Pennsylvania Right To Know
- Cellulose 9004-34-6
- Lactose-1-Hydrate 64044-51-5
- Cobimetinib 1369665-02-0
- Croscarmellose sodium 74811-65-7

Maine Chemicals of High Concern
- Product does not contain any listed chemicals

Vermont Chemicals of High Concern
- Product does not contain any listed chemicals

Washington Chemicals of High Concern
...
Product does not contain any listed chemicals

**California Prop. 65**
WARNING: This product can expose you to chemicals including Titanium oxide (TiO2), which is/are known to the State of California to cause cancer. For more information go to www.P65Warnings.ca.gov.

**California Permissible Exposure Limits for Chemical Contaminants**
- Cellulose: 9004-34-6
- Octadecanoic acid, magnesium salt (2:1): 557-04-0

**California Regulated Carcinogens**
- Talc (Mg3H2(SiO3)4): 14807-96-6

The ingredients of this product are reported in the following inventories:

- **AIIC**: Not in compliance with the inventory
- **DSL**: This product contains the following components that are not on the Canadian DSL nor NDSL:
  - Cobimetinib
  - Croscarmellose sodium
- **NZIoC**: Not in compliance with the inventory
- **ENCS**: Not in compliance with the inventory
- **ISHL**: Not in compliance with the inventory
- **KECI**: Not in compliance with the inventory
- **PICCS**: Not in compliance with the inventory
- **IECSC**: Not in compliance with the inventory
- **TCSI**: Not in compliance with the inventory
- **TSCA**: Product contains substance(s) not listed on TSCA inventory.
- **TECI**: Not in compliance with the inventory

**TSCA list**
No substances are subject to a Significant New Use Rule.

No substances are subject to TSCA 12(b) export notification requirements.

**SECTION 16. OTHER INFORMATION**
SAFETY DATA SHEET

COTELLIC(R) Tablets (20 mg)

Version 1.5
Revision Date: 10-27-2021
Date of last issue: 01-22-2020
Date of first issue: 12-07-2015

NFPA 704:

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HMIS® IV:

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<tr>
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</table>

HMIS® ratings are based on a 0-4 rating scale, with 0 representing minimal hazards or risks, and 4 representing significant hazards or risks. The "*" represents a chronic hazard, while the "/'" represents the absence of a chronic hazard.

Full text of other abbreviations

ACGIH: USA. ACGIH Threshold Limit Values (TLV)
NIOSH REL: USA. NIOSH Recommended Exposure Limits
OSHA P0: USA. OSHA - TABLE Z-1 Limits for Air Contaminants - 1910.1000
OSHA Z-1: USA. Occupational Exposure Limits (OSHA) - Table Z-1 Limits for Air Contaminants
OSHA Z-3: USA. Occupational Exposure Limits (OSHA) - Table Z-3 Mineral Dusts
ACGIH / TWA: 8-hour, time-weighted average
NIOSH REL / TWA: Time-weighted average concentration for up to a 10-hour workday during a 40-hour workweek
OSHA P0 / TWA: 8-hour time weighted average
OSHA Z-1 / TWA: 8-hour time weighted average
OSHA Z-3 / TWA: 8-hour time weighted average

AIIC - Australian Inventory of Industrial Chemicals; ASTM - American Society for the Testing of Materials; bw - Body weight; CERCLA - Comprehensive Environmental Response, Compensation, and Liability Act; CMR - Carcinogen, Mutagen or Reproductive Toxicant; DIN - Standard of the German Institute for Standardisation; DOT - Department of Transportation; DSL - Domestic Substances List (Canada); ECx - Concentration associated with x% response; EHS - Extremely Hazardous Substance; ELx - Loading rate associated with x% response; EmS - Emergency Schedule; ENCS - Existing and New Chemical Substances (Japan); ErCx - Concentration associated with x% growth rate response; ERG - Emergency Response Guide; GHS - Globally Harmonized System; GLP - Good Laboratory Practice; HMIS - Hazardous Materials Identification System; IARC - International Agency for Research on Cancer; IATA - International Air Transport Association; IBC - International Code for the Construction and Equipment of Ships carrying Dangerous Chemicals in Bulk; IC50 - Half maximal inhibitory concentration; ICAO - International Civil Aviation Organization; IECSC - Inventory of Existing Chemical Substances in China; IMDG - International Maritime Dangerous Goods; IMO - International Maritime Organization; ISHL - Industrial Safety and Health Law (Japan); ISO - International Organisation for Standardization; KECI - Korea Existing Chemicals Inventory; LC50 - Lethal Concentration to 50 % of a test population; LD50 - Lethal Dose to 50% of a test population (Median Lethal Dose); MARPOL - International Convention for the Prevention of Pollution from Ships; MSHA - Mine Safety and Health Administration; OSHA - Occupational Safety and Health Administration; PEL - Permissible Exposure Limit; PPM - Parts Per Million; ppm - Parts per million; ppmv - Parts Per Million Volume; ppmv - Parts per million by volume; safety data sheet; TWA - Time Weighted Average; UL - Underwriters Laboratories; USCS - United States Code of Federal Regulations; WOE - Weight of Evidence; WHO - World Health Organization; X.4 - Hazardous Substance; Z-1 - United States. Code of Federal Regulations - Part 1910 Z-1 - Minimum Requirements for the Protection of Employees Against Hazardous Chemical Substances in the Workplace; Z-3 - United States. Code of Federal Regulations - Part 1910 Z-3 - Minimum Requirements for the Protection of Employees Against Hazardous Dusts.
SAFETY DATA SHEET

COTELLIC(R) Tablets (20 mg)

Version 1.5

Revision Date: 10-27-2021
Date of last issue: 01-22-2020
Date of first issue: 12-07-2015

Health Administration; n.o.s. - Not Otherwise Specified; NFPA - National Fire Protection Association; NO(A)EC - No Observed (Adverse) Effect Concentration; NO(A)EL - No Observed (Adverse) Effect Level; NOELR - No Observable Effect Loading Rate; NTP - National Toxicology Program; NZIoC - New Zealand Inventory of Chemicals; OECD - Organization for Economic Co-operation and Development; OPPTS - Office of Chemical Safety and Pollution Prevention; PBT - Persistent, Bioaccumulative and Toxic substance; PICCS - Philippines Inventory of Chemicals and Chemical Substances; (Q)SAR - (Quantitative) Structure Activity Relationship; RCRA - Resource Conservation and Recovery Act; REACH - Regulation (EC) No 1907/2006 of the European Parliament and of the Council concerning the Registration, Evaluation, Authorisation and Restriction of Chemicals; RQ - Reportable Quantity; SADT - Self-Accelerating Decomposition Temperature; SARA - Superfund Amendments and Reauthorization Act; SDS - Safety Data Sheet; TCSI - Taiwan Chemical Substance Inventory; TECI - Thailand Existing Chemicals Inventory; TSCA - Toxic Substances Control Act (United States); UN - United Nations; UNRTDG - United Nations Recommendations on the Transport of Dangerous Goods; vPvB - Very Persistent and Very Bioaccumulative

Revision Date : 10-27-2021

The information provided in this Safety Data Sheet is correct to the best of our knowledge, information and belief at the date of its publication. The information given is designed only as a guidance for safe handling, use, processing, storage, transportation, disposal and release and is not to be considered a warranty or quality specification. The information relates only to the specific material designated and may not be valid for such material used in combination with any other materials or in any process, unless specified in the text.

US / Z8 / 2010