# Safety Data Sheet

## BONIVA(R) Injection (3 mg/3 ml pre-filled syringes)

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

#### 1.1. Product identifier

- **Product name**: BONIVA(R) Injection (3 mg/3 ml pre-filled syringes)
- **Product code**: SAP-10067516
- **Synonyms**: BONVIVA Prefilled Syringes 3 mg/3 ml, Boniva

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

- Use: pharmaceutical active substance (postmenopausal osteoporosis) *1

#### 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

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

*1 referring to: ibandronate

### SECTION 2: Hazards identification

#### Classification of the substance or mixture / Label elements

- **GHS Classification**: no classification and labelling according to GHS

#### Other hazards

- **Note**: no information available
SECTION 3: Composition/information on ingredients

Characterization: Ibandronate with other inactive ingredients

<table>
<thead>
<tr>
<th>Ingredients</th>
<th>Concentration</th>
<th>GHS-Classification (pure ingredient)</th>
</tr>
</thead>
</table>
| Ibandronate | 0.1 %         | - Combustible dust (No category), USH003  
| 138926-19-9 |                | - Acute toxicity (Category 4), H302  
|             |               | - Skin corrosion/irritation (Category 1B), H314  
|             |               | - Specific target organ toxicity - Repeated exposure (Category 2), H373  

*1 referring to: Ibandronate

For the full text of the H-phrases mentioned in this Section, see Section 16.

SECTION 4: First aid measures

4.1. Description of first aid measures

Eye contact: rinse immediately with tap water for 10 minutes - open eyelids forcibly

Skin contact: drench affected skin with 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

Flash point (liquid): not applicable

5.2. Special hazards arising from the substance or mixture

Specific hazards: no particular hazards known
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 - no special environmental precautions required

6.3. Methods and material for containment and cleaning up

Methods for cleaning up - mop or flush the contaminated area with water

SECTION 7: Handling and storage

7.1. Precautions for safe handling

Suitable materials - glass

7.2. Conditions for safe storage, including any incompatibilities

Storage conditions - below 30 °C

Validity - after opening the content should be used within a short period, see expiry date on the label

Packaging materials - prefilled syringes

SECTION 8: Exposure controls/personal protection

8.1. Control parameters

Threshold value (Roche) air - IOEL (Internal Occupational Exposure Limit): 0.002 mg/m³ *1
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
- in case of intense formation of aerosols: respirator with independent air supply or particle respectively filter mask (depending on the aerosol composition)

Hand protection - protective gloves (eg made of neoprene, nitrile or butyl rubber)

Eye protection - safety glasses

*1 referring to: ibandronate

SECTION 9: Physical and chemical properties

9.1. Information on basic physical and chemical properties

Color colorless
Form clear solution
sterile liquid
Density 1.004 g/ml
pH value 3.9 to 4.1

9.2. Other information

Note - no information available

SECTION 10: Stability and reactivity

10.1. Reactivity

Note - no information available

10.2. Chemical stability

Stability - does not contain any antimicrobial preservative; therefore, care must be taken to ensure the sterility of the prepared solution

10.3. Possibility of hazardous reactions

Note - no information available
10.4. Conditions to avoid

Conditions to avoid - temperatures above 30 °C

10.5. Incompatible materials

Note - no information available

10.6. Hazardous decomposition products

Note - Calcium containing solutions should not be mixed with Bondronat concentrate for solution for infusion

SECTION 11: Toxicological information

11.1. Information on toxicological effects

<table>
<thead>
<tr>
<th>Effect</th>
<th>Value</th>
<th>Note</th>
</tr>
</thead>
<tbody>
<tr>
<td>Acute toxicity</td>
<td></td>
<td></td>
</tr>
<tr>
<td>LD₅₀ (oral, rat)</td>
<td>811 mg/kg</td>
<td></td>
</tr>
<tr>
<td>LD₅₀ (i.v., rat)</td>
<td>30 mg/kg</td>
<td>*1</td>
</tr>
<tr>
<td>Local effects</td>
<td></td>
<td></td>
</tr>
<tr>
<td>skin, eyes, mucous membranes: corrosive</td>
<td></td>
<td>*1</td>
</tr>
<tr>
<td>Sensitization</td>
<td></td>
<td></td>
</tr>
<tr>
<td>non-sensitizing (guinea pig)</td>
<td></td>
<td>*1</td>
</tr>
<tr>
<td>Mutagenicity</td>
<td></td>
<td></td>
</tr>
<tr>
<td>not mutagenic (various in vivo and in vitro test systems)</td>
<td></td>
<td>*1</td>
</tr>
<tr>
<td>Carcinogenicity</td>
<td></td>
<td></td>
</tr>
<tr>
<td>not carcinogenic (several species)</td>
<td></td>
<td>*1</td>
</tr>
<tr>
<td>Reproductive toxicity</td>
<td></td>
<td></td>
</tr>
<tr>
<td>not teratogenic, not embryotoxic (several species)</td>
<td></td>
<td>*1</td>
</tr>
<tr>
<td>does not lower parental fertility (several species)</td>
<td></td>
<td>*1</td>
</tr>
<tr>
<td>Note</td>
<td></td>
<td></td>
</tr>
<tr>
<td>inhibits mechanisms reducing bone mass by long-term binding to bone tissue</td>
<td></td>
<td>*1</td>
</tr>
<tr>
<td>high doses cause: liver damages, kidney damages</td>
<td></td>
<td>*1</td>
</tr>
<tr>
<td>decrease in serum calcium level possible</td>
<td></td>
<td>*1</td>
</tr>
<tr>
<td>dosage (oral): 2.5 to 50 mg/d</td>
<td></td>
<td>*1</td>
</tr>
<tr>
<td>dosage (i.v.): 0.5 mg/3 months to 2.5 mg/day</td>
<td></td>
<td>*1</td>
</tr>
<tr>
<td>Potential Health Effects</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Exposure: Inhalation, Ingestion, Skin contact, Eye contact</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Carcinogenicity: formulation not listed by NTP, IARC or OSHA</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Additional Health Information</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Conditions Aggravated: Hypersensitivity to this material and other materials in its chemical class. Uncorrected hypocalcemia. Severe renal impairment.</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

*1 referring to: Ibandronate
SECTION 12: Ecological information

12.1. Toxicity

Ecotoxicity
- strongly toxic for algae (Selenastrum capricornutum)
  \( \text{EbC}_{50} \) (72 h) 1.4 mg/l
  \( \text{ErC}_{50} \) (72 h) 4.7 mg/l
  NOEC (72 h) 0.22 mg/l
  (OECD No. 201)

- highly toxic for algae (Scenedesmus (=Desmodesmus) subspicatus)
  \( \text{EbC}_{50} \) (72 h) 0.218 mg/l (nominal concentration)
  \( \text{ErC}_{50} \) (72 h) 0.390 mg/l (nominal concentration)
  NOEC (72 h) < 0.1 mg/l (nominal concentration)
  (OECD No. 201)

- barely toxic for planktonic crustaceans (Daphnia magna)
  NOEC (48 h) 100 mg/l
  \( \text{EC}_{50} \) (48 h) > 180 mg/l
  (OECD No. 202)

- no adverse influence on substrate biodegradation (activated sludge)
  concentration (28 d) 41.5 mg/l
  (OECD No. 301B, Modified Sturm Test)

- barely inhibitory on aerobic bacterial reproduction (activated sludge)
  NOEC (5 h) 1300 mg/l
  (growth test)

- barely toxic for fish (carp)
  \( \text{LC}_{50} \) (96 h) 200 mg/l
  \( \text{LC}_{0} \) (96 h) 86 mg/l
  (OECD No. 203)

- highly toxic for algae (Scenedesmus (=Desmodesmus) subspicatus)
  \( \text{EC}_{50} \) (14 d) 0.5 mg/l (nominal concentration)
  NOEC (14 d) 0.1 mg/l (nominal concentration)
  (OECD No. 201)

- no adverse influence on substrate biodegradation concentration (28 d) 100 mg/l
  (Manometric Respirometry Test, OECD No. 301 F)

12.2. Persistence and degradability

Ready biodegradability
- not readily biodegradable
  ≤3 %, 28 d
  (CO\(_2\) Evolution Test, Modified Sturm Test, OECD No. 301B)

- not readily biodegradable
  0 %, 28 d
  (Manometric Respirometry Test, OECD No. 301 F)

Inherent biodegradability
- not inherently biodegradable
  < 10 %, 1 d
  < 10 %, 15 d
  < 10 %, 28 d
  (Zahn-Wellens test, OECD No. 302 B)

- not inherently biodegradable
  < 10 %, 28 d
  (Zahn-Wellens test, OECD No. 302 B)
Abiotic degradation - stable in water, no photodegradation 200 mg/l, water < 2 %, 14 d, ~ 22 °C, under illumination *1

12.3. Bioaccumulative potential
Note - no information available

12.4. Mobility in soil
Note - no information available

Mobility - no significant adsorption (, 28 d, ~22 °C)
  \( K_d = 1210 \text{ l/kg (activated sludge)} \)
  (Adsorption to activated sludge in biodegradability test) *1

12.5. Results of PBT and vPvB assessment
Note - no information available

12.6. Other adverse effects
Note - no information available

Note - after the regular 28 days in the Zahn-Wellens test, without significant degradation and still 400 mg DOC/l, 200 mg DOC/l benzoate was added as a well degradable substrate; after 5 days, only 150 mg DOC/l was left, showing some cometabolic degradation *1

- biphosphonates form complexes with bivalent cations, in the relatively high concentrations in the algal test they deplete the medium as scavengers; hence, the effect on algae is not toxic in the strict sense *1

*1 referring to: Ibandronate

SECTION 13: Disposal considerations

13.1. Waste treatment methods
Waste from residues - observe local/national regulations regarding waste disposal

SECTION 14: Transport information

Note - not classified by transport regulations, proper shipping name non-regulated
**SECTION 15: Regulatory information**

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

<table>
<thead>
<tr>
<th>TSCA Status</th>
<th>FDA Exemption - not on inventory</th>
</tr>
</thead>
<tbody>
<tr>
<td>Reporting Requirements</td>
<td>The United States Environmental Protection Agency (USEPA) has not established a Reportable Quantity (RQ) for releases of this material.</td>
</tr>
<tr>
<td></td>
<td>In New Jersey, report all releases which are likely to endanger the public health, harm the environment or cause a complaint to the NJDEPE Hotline (1-609-292-5560) and to local officials.</td>
</tr>
<tr>
<td></td>
<td>State and local regulations vary and may impose additional reporting requirements.</td>
</tr>
</tbody>
</table>

**SECTION 16: Other information**

Full text of H-Statements referred to under section 3

<table>
<thead>
<tr>
<th>H302</th>
<th>Harmful if swallowed.</th>
</tr>
</thead>
<tbody>
<tr>
<td>H314</td>
<td>Causes severe skin burns and eye damage.</td>
</tr>
<tr>
<td>H373</td>
<td>May cause damage to organs through prolonged or repeated exposure.</td>
</tr>
<tr>
<td>USH003</td>
<td>May form combustible dust concentrations in the air</td>
</tr>
</tbody>
</table>

Note

- Please note this Safety Data Sheet for the bulk product does not apply for the finished, packaged medicinal product intended for the final user.

Edition documentation

- changes from previous version in sections 2, 3, 7, 16

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.