

# Bortezomib

MADE IN GERMANY

Solution for Injection



2.5 mg  
1 ml

3.5 mg  
1.4 ml

Bortezomib EVER Pharma is provided as a ready to use formulation for subcutaneous injection (SC), and after dilution for intravenous injection.

Bortezomib is a protease inhibitor indicated for the treatment of progressive or untreated multiple myeloma, and mantle cell lymphoma.

- Available as a “ready to use” liquid formulation – with in use stability superior to powder formulations
- Two fill sizes providing dosing flexibility, reducing wastage and saving costs – the 2.5 mg/1 ml vial covers the most common patient doses for SC injection
- Both vial presentations come in CytoWrap® – for safer handling and transportation



[www.everpharma.com](http://www.everpharma.com)

 **EVER**  
PHARMA

# IN-USE STABILITY OF 28 DAYS AT 25°C (protected from light)

## Indications

As monotherapy or in combination with pegylated liposomal doxorubicin or dexamethasone is indicated for the treatment of adult patients with progressive multiple myeloma who have received at least 1 prior therapy and who have already undergone or are unsuitable for haematopoietic stem cell transplantation.

In combination with melphalan and prednisone is indicated for the treatment of adult patients with previously untreated multiple myeloma who are not eligible for high-dose chemotherapy with haematopoietic stem cell transplantation.

In combination with dexamethasone, or with dexamethasone and thalidomide, is indicated for the induction treatment of adult patients with previously untreated multiple myeloma who are eligible for high-dose chemotherapy with haematopoietic stem cell transplantation.

In combination with rituximab, cyclo-phosphamide, doxorubicin and prednisone is indicated for the treatment of adult patients with previously untreated mantle cell lymphoma who are unsuitable for haematopoietic stem cell transplantation.

## Active Ingredient

Bortezomib (as a mannitol boronic ester)

## Excipients

Mannitol (E421), Sodium chloride, Sodium hydroxide, Hydrochloric acid

## Presentations

- **2.5 mg/1 ml** solution for injection
- **3.5 mg/1.4 ml** solution for injection

## Strength

Each 1 ml of concentrate contains 2.5 mg Bortezomib.

## Stability

- **Unopened:** 24 months at 2-8°C (protect from light)
- **In-use stability:** Chemical and physical in-use stability after first opening and/or dilution has been demonstrated for 28 days when stored at 25°C (protected from light) and at 2-8°C (protected from light), or for 24 hours when stored at 25°C and normal indoor lighting conditions.

## Primary Packaging

Clear, glass vial closed with a grey bromobutyl rubber stopper sealed with an aluminium cap covered with a plastic flip-off cap.

## Pack sizes

1 or 5 vials per pack. Vials may or may not be sheathed in a protective sleeve.



Plastic sleeving for safer handling

# NO RECONSTITUTION NECESSARY

No reconstitution from powder required prior to preparing patient specific doses saving time and costs



No dilution needed prior to SC injection - most patient doses are given SC as this route of administration has become a standard of care for patients with multiple myeloma.

- Smaller 2.5 mg/ml presentation reducing wastage on most common patient doses<sup>1,2</sup>, or for dose reductions
- 3.5 mg/1.4 ml for larger patients. Can be used directly for SC injection or after dilution for intravenous injection

Common patient doses

BSA m <sup>2</sup>	1.3	1.4	1.5	1.6	1.7	1.8	1.9	2	2.1
Std Dose mg/m <sup>2</sup>	1.3	1.3	1.3	1.3	1.3	1.3	1.3	1.3	1.3
Patient Dose mg	1.69	1.82	1.95	2.08	2.21	2.34	2.47	2.6	2.73
Strength mg/ml	2.5	2.5	2.5	2.5	2.5	2.5	2.5	2.5	2.5
SC Injection volume ml	0.7	0.7	0.8	0.8	0.9	0.9	1.0	1.0	1.1

1. Clark L, et al. Ideal vial size for bortezomib: real-world data on waste and cost reduction in treatment of multiple myeloma in Brazil. Value Health. 2011 Jul-Aug;14(5 Suppl 1):S82-4. doi: 10.1016/j.jval.2011.05.013.

2. Overspending driven by oversized single dose vials of cancer drugs. BMJ 2016;352:i788

# Bortezomib

## Solution for Injection

MADE IN GERMANY



Plastic sleeving for safer handling

### Bortezomib EVER Pharma 2.5 mg/ml solution for injection

Composition: One ml of solution contains 2.5 mg bortezomib (as mannitol boronic ester). List of excipients: Mannitol (E 421), Sodium chloride, Sodium hydroxide (for pH adjustment), Hydrochloric acid (for pH adjustment), Water for injections. Therapeutic indications: 1. as monotherapy or in combination with pegylated liposomal doxorubicin or dexamethasone is indicated for the treatment of adult patients with progressive multiple myeloma who have received at least 1 prior therapy and who have already undergone or are unsuitable for haematopoietic stem cell transplantation. 2. in combination with melphalan and prednisone is indicated for the treatment of adult patients with previously untreated multiple myeloma who are not eligible for high-dose chemotherapy with haematopoietic stem cell transplantation. 3. in combination with dexamethasone, or with dexamethasone and thalidomide, is indicated for the induction treatment of adult patients with previously untreated multiple myeloma who are eligible for high-dose chemotherapy with haematopoietic stem cell transplantation. 4. in combination with rituximab, cyclophosphamide, doxorubicin and prednisone is indicated for the treatment of adult patients with previously untreated mantle cell lymphoma who are unsuitable for haematopoietic stem cell transplantation. Contraindications: Hypersensitivity to the active substance, to boron or to any of the excipients listed in section 6.1. Acute diffuse infiltrative pulmonary and pericardial disease Side effects: Multiple Myeloma: very common: Thrombocytopenia, Neutropenia, Anaemia, Decreased appetite, Neuropathies, Peripheral sensory neuropathy, Dysaesthesia, Neuralgia, Nausea and vomiting symptoms, Diarrhoea, Constipation, Musculoskeletal pain, Pyrexia, Fatigue, Asthenia, common: Herpes zoster (incl. disseminated & ophthalmic), Pneumonia, Herpes simplex, Fungal infection, Leukopenia, Lymphopenia, Dehydration, Hypokalaemia, Hyponatraemia, Blood glucose abnormal, Hypocalcaemia, Enzyme abnormality, Mood disorders and disturbances, Anxiety disorder, Sleep disorders and disturbances, Motor neuropathy, Loss of consciousness (inc syncope), Dizziness, Dysgeusia, Lethargy, Headache, Eye swelling, Vision abnormal, Conjunctivitis, Vertigo, Hypotension, Orthostatic hypotension, Hypertension, Dyspnoea, Epistaxis, Upper/lower respiratory tract infection, Cough, Gastrointestinal haemorrhage (inc mucosal), Dyspepsia, Stomatitis, Abdominal distension, Oropharyngeal pain, Abdominal pain (inc gastrointestinal and splenic pain), Oral disorder, Flatulence, Hepatic enzyme abnormality, Rash, Pruritus, Erythema, Dry skin, Muscle spasms, Pain in extremity, Muscular weakness, Renal impairment, Oedema (inc peripheral), Chills, Pain, Malaise, Weight decreased, uncommon: Infection, Bacterial infections, Viral infections, Sepsis (incl. septic shock), Bronchopneumonia, Herpes virus infection, Meningoencephalitis herpetic, Bacteraemia (incl. staphylococcal), Hordeolum, Influenza, Cellulitis, Device related infection, Skin infection, Ear infection, Staphylococcal infection, Tooth infection, Pancytopenia, Febrile neutropenia, Coagulopathy, Leukocytosis, Lymphadenopathy, Haemolytic anaemia, Angioedema, Hypersensitivity, Cushing's syndrome, Hyperthyroidism, Inappropriate antidiuretic hormone secretion, Tumour lysis syndrome, Failure to thrive, Hypomagnesaemia, Hypophosphataemia, Hyperkalaemia, Hypercalcaemia, Hypernatraemia, Uric acid abnormal, Diabetes mellitus, Fluid retention, Mental disorder, Hallucination, Psychotic disorder, Confusion, Restlessness, Tremor, Peripheral sensorimotor neuropathy, Dyskinesia, Cerebellar coordination and balance disturbances, Memory loss (exc dementia), Encephalopathy, Posterior Reversible Encephalopathy Syndrome, Neurotoxicity, Seizure disorders, Post herpetic neuralgia, Speech disorder, Restless legs syndrome, Migraine, Sciatica, Disturbance in attention, Reflexes abnormal, Parosmia, Eye haemorrhage, Eyelid infection, Chalazion, Blepharitis, Eye inflammation, Diplopia, Dry eye, Eye irritation, Eye pain, Lacrimation increased, Eye discharge, Dysacusis (inc tinnitus), Hearing impaired (up to and inc deafness), Ear discomfort, Cardiac tamponade, Cardio-pulmonary arrest, Cardiac fibrillation (inc atrial), Cardiac failure (inc left and right ventricular), Arrhythmia, Tachycardia, Palpitations, Angina pectoris, Pericarditis (inc pericardial effusion), Cardiomyopathy, Ventricular dysfunction, Bradycardia, Cerebrovascular accident, Deep vein thrombosis, Haemorrhage, Thrombophlebitis (inc superficial), Circulatory collapse (inc hypovolaemic shock), Phlebitis, Flushing, Haematoma (inc perirenal), Poor peripheral circulation, Vasculitis, Hyperaemia (inc ocular), Pulmonary embolism, Pleural effusion, Pulmonary oedema (inc acute), Pulmonary alveolar haemorrhage, Bronchospasm, Chronic obstructive pulmonary disease, Hypoxaemia, Respiratory tract congestion, Hypoxia, Pleurisy, Hiccups, Rhinorrhoea, Dysphonia, Wheezing, Pancreatitis (inc chronic), Haematemesis, Lip swelling, Gastrointestinal obstruction (inc small intestinal obstruction, ileus), Abdominal discomfort, Oral ulceration, Enteritis, Gastritis, Gingival bleeding, Gastroesophageal reflux disease, Colitis (inc clostridium difficile), Colitis ischaemic, Gastrointestinal inflammation, Dysphagia, Irritable bowel syndrome, Gastrointestinal disorder, Tongue coated, Gastrointestinal motility disorder, Salivary gland disorder, Hepatotoxicity (inc liver disorder), Hepatitis, Cholestasis, Erythema multiforme, Urticaria, Acute febrile neutrophilic dermatosis, Toxic skin eruption, Toxic epidermal necrolysis, Stevens-Johnson syndrome, Dermatitis, Hair disorder, Petechiae, Ecchymosis, Skin lesion, Purpura, Skin mass, Psoriasis, Hyperhidrosis, Night sweats, Decubitus ulcer, Acne, Blister, Pigmentation disorder, Muscle twitching, Joint swelling, Arthritis, Joint stiffness, Myopathies, Sensation of heaviness, Renal failure acute, Renal failure chronic, Urinary tract infection, Urinary tract signs and symptoms, Haematuria, Urinary retention, Micturition disorder, Proteinuria, Azotaemia, Oliguria, Pollakisuria, Vaginal haemorrhage, Genital pain, Erectile dysfunction, General physical health deterioration, Face oedema, Injection site reaction, Mucosal disorder, Chest pain, Gait disturbance, Feeling cold, Extravasation, Catheter related complication, Change in thirst, Chest discomfort, Feeling of body temperature change, Injection site pain, Hyperbilirubinaemia, Protein analyses abnormal, Weight increased, Blood test abnormal, C-reactive protein increased, Fall, Contusion, rare: Meningitis (inc bacterial), Epstein-Barr virus infection, Genital herpes, Tonsillitis, Mastoiditis, Post viral fatigue syndrome, Neoplasm malignant, Leukaemia plasmacytic, Renal cell carcinoma, Mass, Mycosis fungoides, Neoplasm benign, Disseminated intravascular coagulation, Thrombocytosis, Hypersensitivity syndrome, Platelet disorder, Thrombotic microangiopathy (including Thrombotic purpura), Blood disorder, Haemorrhagic diathesis, Lymphocytic infiltration, Anaphylactic shock, Amyloidosis, Type III immune complex mediated reaction, Hypothyroidism, Hypermagnesaemia, Acidosis, Electrolyte imbalance, Fluid overload, Hypochloroemia, Hypovolaemia, Hyperchloroemia, Hyperphosphataemia, Metabolic disorder, Vitamin B complex deficiency, Vitamin B12 deficiency, Gout, Increased appetite, Alcohol intolerance, Suicidal ideation, Adjustment disorder, Delirium, Libido decreased, Cerebral haemorrhage, Haemorrhage intracranial (inc subarachnoid), Brain oedema, Transient ischaemic attack, Coma, Autonomic nervous system imbalance, Autonomic neuropathy, Cranial palsy, Paralysis, Paresis, Presyncope, Brain stem syndrome, Cerebrovascular disorder, Nerve root lesion, Psychomotor hyperactivity, Spinal cord compression, Cognitive disorder, Motor dysfunction, Nervous system disorder, Radiculitis, Drooling, Hypotonia, Guillain-Barré syndrome, Demyelinating polyneuropathy, Corneal lesion, Exophthalmos, Peritinitis, Scotoma, Eye disorder (inc. eyelid), Dacryoadenitis acquired, Photophobia, Photopsia, Optic neuropathy, Different degrees of visual impairment (up to blindness), Ear haemorrhage, Vestibular neuronitis, Ear disorder, Atrial flutter, Myocardial infarction, Atrioventricular block, Cardiovascular disorder (inc cardiogenic shock), Torsade de pointes, Angina unstable, Cardiac valve disorders, Coronary artery insufficiency, Sinus arrest, Peripheral embolism, Lymphoedema, Pallor, Erythromelalgia, Vasodilatation, Vein discoloration, Venous insufficiency, Respiratory failure, Acute respiratory distress syndrome, Apnoea, Pneumothorax, Atelectasis, Pulmonary hypertension, Haemoptysis, Hyperventilation, Orthopnoea, Pneumonitis, Respiratory alkalosis, Tachypnoea, Pulmonary fibrosis, Bronchial disorder, Hypocapnia, Interstitial lung disease, Lung infiltration, Throat tightness, Dry throat, Increased upper airway secretion, Throat irritation, Upper-airway cough syndrome, Pancreatitis acute, Peritonitis, Tongue oedema, Ascites, Oesophagitis, Chellitis, Faecal incontinence, Anal sphincter atony, Faecaloma, Gastrointestinal ulceration and perforation, Gingival hypertrophy, Megacolon, Rectal discharge, Oropharyngeal blistering, Lip pain, Periodontitis, Anal fissure, Change of bowel habit, Proctalgia, Abnormal faeces, Hepatic failure, Hepatomegaly, Budd-Chiari syndrome, Cytomegalovirus hepatitis, Hepatic haemorrhage, Cholelithiasis, Skin reaction, Jessner's lymphocytic infiltration, Palmar-plantar erythrodysesthesia syndrome, Haemorrhage subcutaneous, Livedo reticularis, Skin induration, Papule, Photosensitivity reaction, Seborrhoea, Cold sweat, Skin disorder, Erythrosis, Skin ulcer, Nail disorder, Rhabdomyolysis, Temporomandibular joint syndrome, Fistula, Joint effusion, Pain in jaw, Bone disorder, Musculoskeletal and connective tissue infections and inflammations, Synovial cyst, Bladder irritation, Testicular disorder, Prostatitis, Breast disorder female, Epididymal tenderness, Epididymitis, Pelvic pain, Vulval ulceration, Aplasia, Gastrointestinal malformation, Ichthyosis, Death (inc sudden), Multi-organ failure, Injection site haemorrhage, Hernia (inc hiatus), Impaired healing, Inflammation, Injection site phlebitis, Tenderness, Ulcer, Irritability, Non-cardiac chest pain, Catheter site pain, Sensation of foreign body, Blood gases abnormal, Electrocardiogram abnormalities (inc QT prolongation), International normalised ratio abnormal, Gastric pH decreased, Platelet aggregation increased, Troponin I increased, Virus identification and serology, Urine analysis abnormal, Transfusion reaction, Fractures, Rigors, Face injury, Joint injury, Burns, Laceration, Procedural pain, Radiation injuries, Macrophage activation, Mantle Cell Lymphoma: very common: Pneumonia, Thrombocytopenia, Febrile neutropenia, Neutropenia, Leukopenia, Anaemia, Lymphopenia, Decreased appetite, Peripheral sensory neuropathy, Dysaesthesia, Neuralgia, Nausea and vomiting symptoms, Diarrhoea, Stomatitis, Constipation, Hair disorder, Pyrexia, Fatigue, Asthenia, common: Sepsis (incl. septic shock), Herpes zoster (incl. disseminated & ophthalmic), Herpes virus infection, Bacterial infections, Upper/lower respiratory tract infection, Fungal infection, Herpes simplex, Hypersensitivity, Hypokalaemia, Blood glucose abnormal, Hyponatraemia, Diabetes mellitus, Fluid retention, Sleep disorders and disturbances, Neuropathies, Motor neuropathy, Loss of consciousness (incl. syncope), Encephalopathy, Peripheral sensorimotor neuropathy, Dizziness, Dysgeusia, Autonomic neuropathy, Vision abnormal, Dysacusis (incl. tinnitus), Cardiac fibrillation (incl. atrial), Arrhythmia, Cardiac failure (incl. left and right ventricular), Myocardial ischaemia, Ventricular dysfunction, Hypertension, Hypotension, Orthostatic hypotension, Dyspnoea, Cough, Hiccups, Gastrointestinal haemorrhage (incl. mucosal), Abdominal distension, Dyspepsia, Oropharyngeal pain, Gastritis, Oral ulceration, Abdominal discomfort, Dysphagia, Gastrointestinal inflammation, Abdominal pain (incl. gastrointestinal and splenic pain), Oral disorder, Hepatotoxicity (inc liver disorder), Pruritus, Dermatitis, Rash, Muscle spasms, Musculoskeletal pain, Pain in extremity, Urinary tract infection, Oedema (incl. peripheral), Chills, Injection site reaction, Malaise, Hyperbilirubinaemia, Protein analyses abnormal, Weight decreased, Weight increased, uncommon: Hepatitis B, Infection, Bronchopneumonia, Pancytopenia, Anaphylactic reaction, Tumour lysis syndrome, Autonomic nervous system imbalance, Vertigo, Hearing impaired (up to and incl. deafness), Cardiovascular disorder (incl. cardiogenic shock), Acute respiratory distress syndrome, Pulmonary embolism, Pneumonitis, Pulmonary hypertension, Pulmonary oedema (incl. acute), Colitis (inc clostridium difficile), Hepatic failure. More information available in the summary of product characteristics. Only available on prescription. Last update: August 2021. Marketing Authorisation Holder: EVER Valinject GmbH, Oberburgau 3, 4866 Unterach am Attersee, Austria.