HOSPITAL OUTPATIENT CODING AND BILLING INFORMATION SHEET FOR KYPROLIS® (CARFILZOMIB) FOR INJECTION

INDICATIONS

- KYPROLIS® (carfilzomib) is indicated in combination with dexamethasone, or with lenalidomide plus dexamethasone, or with daratumumab plus dexamethasone, or with daratumumab plus hyaluronidase-fihj plus dexamethasone, or with isatuximab plus dexamethasone for the treatment of adult patients with relapsed or refractory multiple myeloma who have received one to three lines of therapy.
- KYPROLIS® is indicated as a single agent for the treatment of patients with relapsed or refractory multiple myeloma who have received one or more lines of therapy.

IMPORTANT SAFETY INFORMATION FOR KYPROLIS

Cardiac Toxicities

- New onset or worsening of pre-existing cardiac failure (e.g., congestive heart failure, pulmonary edema, decreased ejection fraction), cardiomyopathy, myocardial ischemia, and myocardial infarction including fatalities have occurred following administration of KYPROLIS. Some events occurred in patients with normal baseline ventricular function. Death due to cardiac arrest has occurred within one day of administration.
- Monitor patients for signs or symptoms of cardiac failure or ischemia. Evaluate promptly if cardiac toxicity is suspected. Withhold KYPROLIS for Grade 3 or 4 cardiac adverse reactions until recovery, and consider whether to restart at 1 dose level reduction based on a benefit/risk assessment.
- While adequate hydration is required prior to each dose in Cycle 1, monitor all patients for evidence of volume overload, especially patients at risk for cardiac failure. Adjust total fluid intake as clinically appropriate.
- For patients ≥ 75 years, the risk of cardiac failure is increased. Patients with New York Heart Association Class III and IV
 heart failure, recent myocardial infarction, conduction abnormalities, angina, or arrhythmias may be at greater risk for cardiac
 complications and should have a comprehensive medical assessment prior to starting treatment with KYPROLIS and remain under
 close follow-up with fluid management.



HOSPITAL OUTPATIENT CODING AND BILLING INFORMATION SHEET FOR KYPROLIS® (CARFILZOMIB) FOR INJECTION

Item	Revenue Code	Coding Information (HCPCS/CPT/ICD)	Notes
KYPROLIS	Medicare: 0636, drugs requiring detailed coding	J9047, injection, carfilzomib, 1 mg ¹	KYPROLIS VIALS: KYPROLIS is supplied in single-dose vials containing 60 mg, 30 mg, or 10 mg of carfilzomib. ³
	Other Payers: 0250, general pharmacy; OR 0636, if required by a given payer		The NDC numbers for KYPROLIS, in the 11-digit format, are as follows ³ : - 60-mg vial: 76075-0101-01 - 30-mg vial: 76075-0102-01 - 10-mg vial: 76075-0103-01
			MEDICARE MUE FOR KYPROLIS ³⁻⁶ : Under Medicare,* J9047 has a Medically Unlikely Edit (MUE). Based on the approved dosing range, Medicare will deny KYPROLIS claims billed for more than 160 units per date of service For example, at the BSA of up to 2.2 m², the calculated dose for Kd70 is up to 154 mg
			JW MODIFIER7: For unused drug from single use vials, some payers (eg, Medicare*) require providers to report the JW modifier on a claim and to document the discarded amount in the patient's medical record.
Administration	Appropriate revenue code for the cost center in which the service is performed	96413, chemotherapy administration, IV infusion technique; up to 1 hour, single or initial substance/drug² OR 96409, chemotherapy administration, intravenous push technique (including infusions less than 15 minutes), single or initial substance/drug²	KYPROLIS® can be administered as a 10-minute or 30-minute IV infusion, depending on the selected regimen¹: - At the priming dose of 20 mg/m² and at the therapeutic dose of 70 mg/m² once-weekly (Kd or DKd): KYPROLIS® is administered as a 30-minute IV infusion. - At the priming dose of 20 mg/m² and at the therapeutic dose of 56 mg/m² twice-weekly (Kd, DKd, or K): KYPROLIS® is administered as a 30-minute IV infusion. - At the priming dose of 20 mg/m² and at the therapeutic dose of 27 mg/m² twice-weekly (KRd or K): KYPROLIS® is administered as a 10-minute IV infusion.
Diagnosis/ Condition	N/A	Appropriate diagnosis code(s) for patient condition	ICD-10-CM Examples: C90.00, multiple myeloma not having achieved remission C90.02, multiple myeloma in relapse ⁸

^{*}Applies to Medicare fee-for-service; requirements for Medicare Advantage may vary by plan.

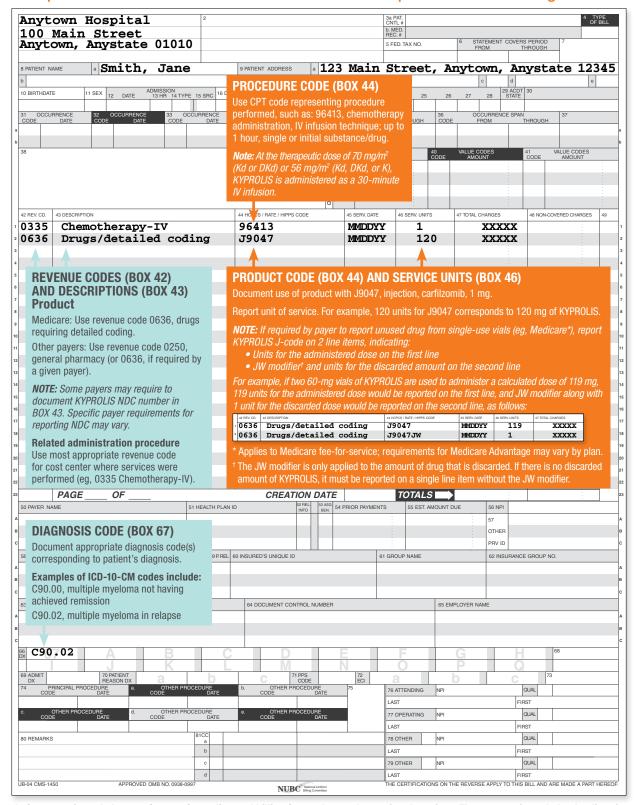
The information provided in this Coding and Billing Information document is of a general nature and for informational purposes only and is not intended to be a comprehensive list nor instructive. Coding and coverage policies change periodically and often without warning. The responsibility to determine coverage and reimbursement parameters, and appropriate coding for a particular patient and/or procedure, is always the responsibility of the provider or physician. Healthcare providers should consult the payer or Medicare contractor to determine which CPT® code(s) are most appropriate for administration of Kyprolis®. The codes provided here are not an exhaustive list of drug administration services. Please refer to the CPT® manual for a complete list of drug administration codes. The information provided in this section should in no way be considered a guarantee of coverage or reimbursement for any product or service.

BSA = body surface area; IV = intravenous; Kd = KYPROLIS® (carfilzomib) and dexamethasone; KRd = KYPROLIS®+lenalidomide and dexamethasone; DKd = KYPROLIS®+daratumumab and dexamethasone



THE SAMPLE UB-04 (CMS 1450) FOR HOSPITAL OUTPATIENT — KYPROLIS AT 70 mg/m² OR 56 mg/m²

Hospital Outpatient Administration of KYPROLIS at the Therapeutic Dose of 70 mg/m² or 56 mg/m²

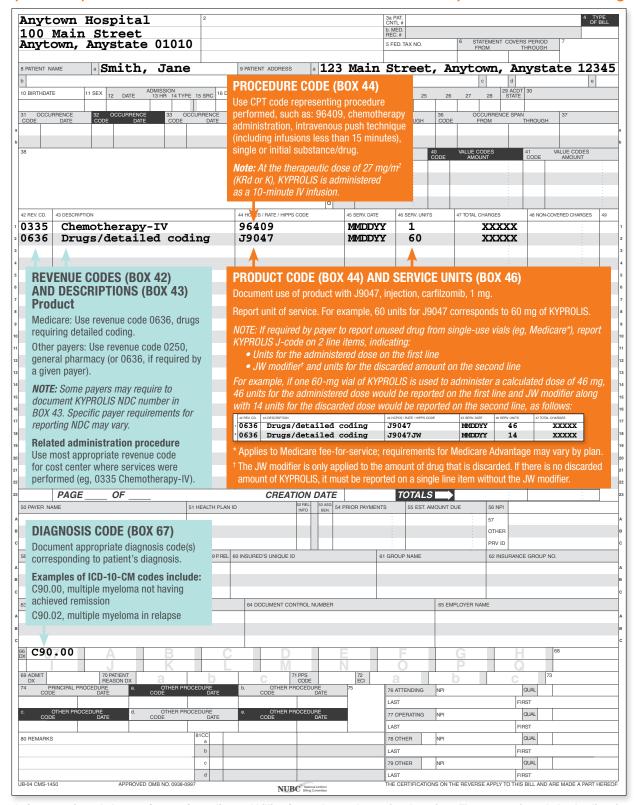


These sample forms are intended as a reference for coding and billing for product and associated services. They are not intended to be directive; the use of the recommended codes does not guarantee reimbursement. Healthcare providers may deem other codes or policies more appropriate and should select the coding options that most accurately reflect their internal system guidelines, payer requirements, practice patterns, and the services rendered. Healthcare providers are responsible for ensuring the accuracy and validity of all billing and claims for appropriate reimbursement.



THE SAMPLE UB-04 (CMS 1450) FOR HOSPITAL OUTPATIENT — KYPROLIS AT 27 mg/m²

Hospital Outpatient Administration of KYPROLIS at the Therapeutic Dose of 27 mg/m²



These sample forms are intended as a reference for coding and billing for product and associated services. They are not intended to be directive; the use of the recommended codes does not guarantee reimbursement. Healthcare providers may deem other codes or policies more appropriate and should select the coding options that most accurately reflect their internal system guidelines, payer requirements, practice patterns, and the services rendered. Healthcare providers are responsible for ensuring the accuracy and validity of all billing and claims for appropriate reimbursement.





Support, Simplified

FINANCIAL SUPPORT OPTIONS FOR ANY INSURANCE TYPE

Whatever type of insurance your patients have—even if they have none—we can help you understand how their Amgen medicine may be covered, and refer them to programs that may be able to help them afford their medication, such as Amgen FIRST STEPTM.*



For Eligible[†] Commercially Insured Patients

The Amgen FIRST STEP™ program can help patients cover their out-of-pocket prescription costs, including deductible, co-insurance, or co-payment.

- \$0 out-of-pocket for first dose or cycle
- As little as \$5* out-of-pocket for subsequent doses or cycles, up to the brand program benefit maximum
- No income eligibility requirement

For Patients On Government Insurance Like Medicare

Amgen Assist 360[™] can refer patients to independent nonprofit patient assistance programs that may be able to help them afford the co-pay costs for their prescribed medicine.§

For Uninsured Patients

Amgen Safety Net Foundation, a nonprofit patient assistance program sponsored by Amgen, helps qualified patients access Amgen medicines at no cost.

Call **1-888-4ASSIST (1-888-427-7478)** Monday—Friday, 9 AM to 8 PM ET or visit **WWW.AMGENASSIST360.COM/ENROLL**

- * Resources include referrals to independent nonprofit patient assistance programs. Eligibility for resources provided by independent nonprofit patient assistance programs is based on the nonprofits' criteria. Amgen has no control over these programs and provides referrals as a courtesy only.
- [†]Terms, conditions, and program maximums apply. This program is not open to patients receiving prescription reimbursement under any federal, state, or government-funded healthcare program. Not valid where prohibited by law. Other restrictions may apply. Please see full Terms and Conditions at www.AmgenFirstStep.com.
- ‡ As little as \$5 out-of-pocket cost for subsequent dose or cycle of KYPROLIS® through Amgen FIRST STEP™.
- § Amgen Assist 360™ can refer patients to independent nonprofit patient assistance programs that may be able to help them afford the co-pay costs for their prescribed medicine.

References: 1. CMS. 2020 Alpha-Numeric HCPCS File. https://www.cms.gov/Medicare/Coding/HCPCSReleaseCodeSets/Alpha-Numeric-HCPCS-Items/2020-Alpha-Numeric-HCPCS-File. Accessed July 25, 2021. 2. American Medical Association. Current Procedural Terminology (CPT®) copyright 2020 American Medical Association. 2021. All rights reserved. 3. KYPROLIS® (carfilzomib) prescribing information. Onyx Pharmaceuticals, Inc., an Amgen Inc. subsidiary. 4. CMS. Medically Unlikely Edits - Facility Outpatient Hospital Services MUE Table - Effective 07-01-2021. https://www.cms.gov/Medicare/Coding/NationalCorrectCodInitEd/MUE.html. Accessed July 25, 2021. 5. CMS. Medically Unlikely Edits - Practitioner Services MUE Table - Effective 07-01-2021. https://www.cms.gov/Medicare/Coding/NationalCorrectCodInitEd/MUE.html. Accessed July 25, 2021. 6. CMS. Medicare NCCI 2021 Coding Policy Manual. Chapter 1. https://www.cms.gov/Medicare/Coding/NationalCorrectCodInitEd. Accessed July 25, 2021. 7. CMS. MLN Matters MM9603. https://www.cms.gov/Outreach-and-Education/Medicare-Learning-Network-MLN/MLNMattersArticles/Downloads/MM9603.pdf. Accessed July 25, 2021. 8. CMS. ICD-10-CM. 2021.



IMPORTANT SAFETY INFORMATION FOR KYPROLIS

Cardiac Toxicities

- New onset or worsening of pre-existing cardiac failure (e.g., congestive heart failure, pulmonary edema, decreased ejection fraction), cardiomyopathy, myocardial ischemia, and myocardial infarction including fatalities have occurred following administration of KYPROLIS. Some events occurred in patients with normal baseline ventricular function. Death due to cardiac arrest has occurred within one day of administration.
- Monitor patients for signs or symptoms of cardiac failure or ischemia. Evaluate promptly if cardiac toxicity is suspected. Withhold KYPROLIS for Grade 3 or 4 cardiac adverse reactions until recovery, and consider whether to restart at 1 dose level reduction based on a benefit/risk assessment.
- While adequate hydration is required prior to each dose in Cycle 1, monitor all patients for evidence of volume overload, especially patients at risk for cardiac failure. Adjust total fluid intake as clinically appropriate.
- For patients ≥ 75 years, the risk of cardiac failure is increased. Patients with New York Heart Association Class III and IV
 heart failure, recent myocardial infarction, conduction abnormalities, angina, or arrhythmias may be at greater risk for cardiac
 complications and should have a comprehensive medical assessment prior to starting treatment with KYPROLIS and remain under
 close follow-up with fluid management.

Acute Renal Failure

Cases of acute renal failure, including some fatal renal failure events, and renal insufficiency (including renal failure) have
occurred. Acute renal failure was reported more frequently in patients with advanced relapsed and refractory multiple myeloma
who received KYPROLIS monotherapy. Monitor renal function with regular measurement of the serum creatinine and/or estimated
creatinine clearance. Reduce or withhold dose as appropriate.

Tumor Lysis Syndrome

Cases of Tumor Lysis Syndrome (TLS), including fatal outcomes, have occurred. Patients with a high tumor burden should be
considered at greater risk for TLS. Adequate hydration is required prior to each dose in Cycle 1, and in subsequent cycles as
needed. Consider uric acid lowering drugs in patients at risk for TLS. Monitor for evidence of TLS during treatment and manage
promptly, and withhold until resolved.

Pulmonary Toxicity

Acute Respiratory Distress Syndrome (ARDS), acute respiratory failure, and acute diffuse infiltrative pulmonary disease such as
pneumonitis and interstitial lung disease have occurred. Some events have been fatal. In the event of drug-induced pulmonary
toxicity, discontinue KYPROLIS.

Pulmonary Hypertension

 Pulmonary arterial hypertension (PAH) was reported. Evaluate with cardiac imaging and/or other tests as indicated. Withhold KYPROLIS for PAH until resolved or returned to baseline and consider whether to restart based on a benefit/risk assessment.

Dyspnea

 Dyspnea was reported in patients treated with KYPROLIS. Evaluate dyspnea to exclude cardiopulmonary conditions including cardiac failure and pulmonary syndromes. Stop KYPROLIS for Grade 3 or 4 dyspnea until resolved or returned to baseline. Consider whether to restart based on a benefit/risk assessment.

Hypertension

Hypertension, including hypertensive crisis and hypertensive emergency, has been observed, some fatal. Control hypertension
prior to starting KYPROLIS. Monitor blood pressure regularly in all patients. If hypertension cannot be adequately controlled,
withhold KYPROLIS and evaluate. Consider whether to restart based on a benefit/risk assessment.

Venous Thrombosis

- Venous thromboembolic events (including deep venous thrombosis and pulmonary embolism) have been observed. Provide
 thromboprophylaxis for patients being treated with the combination of KYPROLIS with dexamethasone or with lenalidomide plus
 dexamethasone or with daratumumab and dexamethasone. The thromboprophylaxis regimen should be based on an assessment
 of the patient's underlying risks.
- For patients using hormonal contraception associated with a risk of thrombosis, consider an alternative method of effective contraception during treatment.



IMPORTANT SAFETY INFORMATION FOR KYPROLIS (cont'd)

Infusion-Related Reactions

Infusion-related reactions, including life-threatening reactions, have occurred. Signs and symptoms include fever, chills, arthralgia, myalgia, facial flushing, facial edema, laryngeal edema, vomiting, weakness, shortness of breath, hypotension, syncope, chest tightness, or angina. These reactions can occur immediately following or up to 24 hours after administration. Premedicate with dexamethasone to reduce the incidence and severity of infusion-related reactions.

Hemorrhage

• Fatal or serious cases of hemorrhage have been reported. Hemorrhagic events have included gastrointestinal, pulmonary, and intracranial hemorrhage and epistaxis. Promptly evaluate signs and symptoms of blood loss. Reduce or withhold dose as appropriate.

Thrombocytopenia

KYPROLIS causes thrombocytopenia with recovery to baseline platelet count usually by the start of the next cycle. Monitor platelet
counts frequently during treatment. Reduce or withhold dose as appropriate.

Hepatic Toxicity and Hepatic Failure

• Cases of hepatic failure, including fatal cases, have occurred. KYPROLIS can cause increased serum transaminases. Monitor liver enzymes regularly regardless of baseline values. Reduce or withhold dose as appropriate.

Thrombotic Microangiopathy

 Cases of thrombotic microangiopathy, including thrombotic thrombocytopenic purpura/hemolytic uremic syndrome (TTP/HUS), including fatal outcome, have occurred. Monitor for signs and symptoms of TTP/HUS. Discontinue if diagnosis is suspected. If the diagnosis of TTP/HUS is excluded, KYPROLIS may be restarted. The safety of reinitiating KYPROLIS is not known.

Posterior Reversible Encephalopathy Syndrome (PRES)

 Cases of PRES have occurred in patients receiving KYPROLIS. If PRES is suspected, discontinue and evaluate with appropriate imaging. The safety of reinitiating KYPROLIS is not known.

Progressive Multifocal Leukoencephalopathy (PML)

Cases of PML, including fatal cases, have occurred. In addition to KYPROLIS, other contributary factors may include prior
or concurrent use of immunosuppressive therapy. Consider PML in any patient with new onset of or changes in pre-existing
neurological signs or symptoms. If PML is suspected, discontinue and initiate evaluation for PML including neurology consultation.

Increased Fatal and Serious Toxicities in Combination with Melphalan and Prednisone in Newly Diagnosed Transplant-ineligible Patients

 In a clinical trial of transplant-ineligible patients with newly diagnosed multiple myeloma comparing KYPROLIS, melphalan, and prednisone (KMP) vs bortezomib, melphalan, and prednisone (VMP), a higher incidence of serious and fatal adverse reactions was observed in patients in the KMP arm. KMP is not indicated for transplant-ineligible patients with newly diagnosed multiple myeloma.

Embryo-fetal Toxicity

- KYPROLIS can cause fetal harm when administered to a pregnant woman.
- Advise pregnant women of the potential risk to a fetus. Females of reproductive potential should use effective contraception
 during treatment with KYPROLIS and for 6 months following the final dose. Males of reproductive potential should use effective
 contraception during treatment with KYPROLIS and for 3 months following the final dose.

Adverse Reactions

- The most common adverse reactions occurring in at least 20% of patients taking KYPROLIS in the combination therapy trials: anemia, diarrhea, hypertension, fatigue, upper respiratory tract infection, thrombocytopenia, pyrexia, cough, dyspnea, and insomnia.
- The most common adverse reactions occurring in at least 20% of patients taking KYPROLIS in monotherapy trials: anemia, fatigue, thrombocytopenia, nausea, pyrexia, dyspnea, diarrhea, headache, cough, edema peripheral.

