

Modified Delphi Panel Consensus Guidance and Essential Strategies on the Prevention and Management of Nausea and Vomiting in Patients Treated With Zolbetuximab + Chemotherapy⁵



- An international RAND/UCLA modified Delphi panel included 15 clinicians
- Panelists^a were selected based on their experience in the zolbetuximab clinical trials

Prior to and During First Infusion



Antilulcer medications (e.g., PPIs or H₂ blockers) or antacids are recommended for patients with dyspepsia who have not had a prior total gastrectomy. Administer PPIs or H₂ blockers for a few days to 1 week prior to zolbetuximab treatment for maximal mucosal protection.

Prophylaxis prior to first infusion

Provide one of the following National Comprehensive Cancer Network[®] (NCCN[®])-recommended high-risk antiemetic drug regimens⁷ (oral/IV antiemetics may be appropriate based on individual patient circumstances)

NK-1 receptor antagonist + 5-HT₃ receptor antagonist + dexamethasone + olanzapine

or

NK-1 receptor antagonist + 5-HT₃ receptor antagonist + dexamethasone

or

5-HT₃ receptor antagonist + dexamethasone + olanzapine

Nausea alone

Consider either making no modifications, or **stopping the infusion for 30-60 mins and restarting it at the same rate** as that prior to stoppage if symptoms improve

and

After the first hour (at an infusion rate of 200-265 mg/m²/hr), consider slowing the infusion rate without stopping it (Please see for additional information)

Any vomiting (Please see for additional information)

Consider adding antiemetic treatment (e.g., rescue medications)⁸

Stop the infusion for 30-60 mins and restart it at a slower rate if symptoms improve (Please see for additional information)

KEY

Zolbetuximab infusion rate
Patient symptoms
 Antiemetic regimens



Note: If infusion was running at the PI rate, **slow the rate by 50%; if the infusion rate had already been slowed to 50%, slow by an additional 50% (i.e., 25% of the initial rate)**



Note: For vomiting, IV hydration may be appropriate depending on individual patient circumstances

Second and Subsequent Infusions



- ✓ Adjust based on the patient's symptoms during previous infusions
- ✓ Begin infusions at the rate that was best tolerated during previous infusion
- ✓ The degree of nausea and vomiting is expected to diminish. In these cases, patients may tolerate titration of the infusion rate by increments of 25% (e.g., if the infusion rate was slowed to 50% and the patient remained asymptomatic for 30-60 mins, consider increasing the rate to 75% back to 100% or maximum tolerated rate. Monitor the patient for recurrence of symptoms, and administer antiemetic medications as needed to manage symptoms

Continuation of treatment



When planning for second or subsequent infusions, if the patient in previous infusion experienced

Nausea alone or one episode of vomiting (Please see for additional information)

Rate recommended in PI

and

Make no changes or escalate the antiemetic regimen from previous infusion

Repeated vomiting (Please see for additional information)

Slower than the recommended rate in PI (Please see for additional information)

and

Escalate the antiemetic regimen from previous infusion

Nausea alone

Make no modifications

Any vomiting (Please see for additional information)

Stop the infusion for 30-60 mins and restart it at a slower rate if symptoms improve (Please see for additional information)

Points to Note



- ✓ Consider administering antiemetics not used for prophylaxis in patients experiencing nausea and vomiting during the infusion
- ✓ Consider a scopolamine patch for refractory nausea or vomiting where premedication options have been escalated
- ✓ Clinicians should avoid zolbetuximab discontinuation and retain flexibility for individual patient management

^aExperts from the US, Europe, Japan, and South Korea reviewed 382 scenarios, reaching an agreement in 85% (n=324). ^bLorazepam, olanzapine, or metoclopramide.

5-HT₃ RA, 5-hydroxytryptamine (serotonin) receptor subtype 3 antagonist; **AE**, adverse event; **C**, cycle; **D**, day; **Hrs**, hours; **H₂**, histamine subtype 2 receptor; **HCP**, healthcare provider; **IRR**, infusion-related reaction; **IV**, intravenous; **mFOLFOX₆**, modified folinic acid, fluorouracil, and oxaliplatin regimen; **NCI-CTCAE v5.0**, National Cancer Institute Common Terminology Criteria for Adverse Events Version 5.0; **NK-1 RA**, neurokinin subtype 1 receptor antagonist; **OS**, overall survival; **PFS**, progression-free survival; **PI**, prescribing information; **PPI**, proton pump inhibitor; **RAND/UCLA**, RAND Corporation/University of California, Los Angeles; **TEAE**, treatment-emergent adverse event; **US**, United States.

1. Zolbetuximab placeholder reference for SmPC. 2. Shitara K, et al. ASCO GI 2024. Abstract #372. 3. Shitara K, et al. Lancet. 2023;401(10389):1655-1668. 4. Shah MA, et al. Nat Med. 2023;29(8):2133-2141. 5. Klempner SJ, et al. ESMO Gastrointest Oncol 2025; 7: 100131. 6. Common Terminology Criteria for Adverse Events (CTCAE) Version 5.0. Accessed January 20, 2025. 7. Referenced with permission from the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines®) for Antiemesis V.2.2025. © National Comprehensive Cancer Network, Inc. 2025. All rights reserved. Accessed May 20, 2025. To view the most recent and complete version of the guideline, go online to NCCN.org. NCCN makes no warranties of any kind whatsoever regarding their content, use or application and disclaims any responsibility for their application or use in any way.