# Survey of Implementation of Antiemetic Prescription Standards in Indian Oncology Practices and Its Adherence to the American Society of Clinical Oncology Antiemetic Clinical Guideline

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Purpose Adherence to international antiemetic prophylaxis guidelines like those of ASCO can result in better control of chemotherapy-induced nausea and vomiting; however, the extent of implementation of such guidelines in India is unknown. Therefore, this survey was planned.

Methods This study was an anonymized cross-sectional survey approved by the ethics committee. Survey items were generated from the clinical questions given in the ASCO guidelines. The survey was disseminated through personal contacts at an oncology conference and via e-mail to various community oncology centers across India. The B1, B2, and B3 domains included questions regarding the optimal antiemetic prophylaxis for high, moderate, and low-minimal emetogenic regimens.

Results Sixty-six (62.9%) of 105 responded and 65 centers (98.5%) were aware of the published guidelines. The partial, full, and no implementation scores were 92.5%, 4.5%, and 3.0%, respectively. Full implementation was better for the low-minimal emetogenic regimens (34.8%) than the highly emetogenic regimens (6.1%). The three most frequent reasons for hampered implementation of ASCO guidelines in routine chemotherapy practice cited by centers were a lack of sensitization (26 centers; 39.4%), lack of national guidelines (12 centers; 18.2%), and lack of administrative support (10 centers; 15.2%).

Conclusion Awareness regarding ASCO antiemetic guidelines is satisfactory in Indian oncology practices; however, there is a need for sensitization of oncologists toward complete implementation of these guidelines in their clinical practice.

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# INTRODUCTION

Chemotherapy-induced nausea and vomiting (CINV) is one of the most common and most distressing complications of chemotherapy. <sup>1</sup> It has a detrimental effect on the quality of life of patients and their functional well-being. <sup>2-4</sup> Uncontrolled nausea or vomiting is also associated with frequent hospital and emergency department visits, is resource consuming, <sup>5-8</sup> and can lead to impaired compliance with chemotherapy. <sup>1,9,10</sup>

There has been significant progress in the development of newer antiemetics and the use of combinations of antiemetics. The optimal combination that is required depends on the emetogenic potential of the specific chemotherapy regimen

used. Clinical guidelines have been published by professional bodies, such as ASCO and the European Society for Medical Oncology, for the same. 11,12 These guidelines have been shown to control the rate and severity of CINV<sup>13-16</sup>; however, adherence rates internationally are variable (29% to 57.3%). 13,14 In India, there are no national guidelines, and oncologists rely most commonly on ASCO clinical updates and the National Comprehensive Cancer Network recommendations, with the extent of implementation largely unknown. To address this gap, the present survey was planned with the primary objective of evaluating the proportion of cancer centers that have fully implemented the ASCO antiemetics clinical practice guideline<sup>11</sup> standards in routine chemotherapy practice. Secondary objectives were to determine the proportion of centers that implement these guidelines partially, to determine the standards that are most commonly and least commonly implemented in each category of chemotherapy drugs according to emetogenicity, and to determine the difficulties that are faced in implementing these standards.

#### **METHODS**

#### **Survey Instrument**

A written survey on the basis of the standards according to the ASCO antiemetics clinical practice guideline was designed. 11 We had conducted a similar survey previously and based the present study on the earlier experience. 17 Survey items were generated from the clinical questions in the guidelines and in the same order. The first and second clinical questions, which dealt with antiemetic prophylaxis in high and moderate emetogenic chemotherapy regimens, respectively, were further broken down into six subquestions each. These questions enquired about the use of an NK receptor antagonist, 5HT3 antagonist, dexamethasone, and their schedules. The remaining survey questions were based on statements in the guideline. Because our intent was to focus on implementation of the ASCO guidelines in a relatively homogenous population of adults with solid tumors, the survey did not include questions related to pediatric patients, hematologic malignancies, bone marrow transplant centers, patients being treated with radiation, and breakthrough emesis.

The survey included questions regarding institutional antiemetic policy (domain A); optimal antiemetic prophylactic regimen for highly emetogenic antineoplastic drugs (domain B1); optimal antiemetic prophylactic regimen for moderately emetogenic antineoplastic drugs (domain B2); optimal antiemetic prophylactic regimen for low-minimal emetogenic antineoplastic drugs (domain B3); and antiemetic use in special situations (domain C). Response options in the survey included a four-item Likert scale (always, usually, rarely, and never), a binary scale (yes or no), or a multiplechoice format, depending on the type of question. In addition, we inquired about the important factors that prevented the center from fully implementing the standards. This response was in the form of a multiple-choice item along with a free-text option. The survey instrument is shown in the Appendix.

In addition, some questions regarding the nature of oncology practice were also added. These questions addressed the following items: state in which

the center was located, setting of practice (urban or rural), teaching status (yes or no), funding source (public or private), and approximate number of patients seen daily.

# **Survey Distribution**

This was an anonymized cross-sectional survey. The survey was designed on Google forms (Google, Mountain View, CA). Oncologists that administer chemotherapy were identified from the ICON (Indian Cooperative Oncology Network) database and invited to participate in this survey. Individual emails with a link to the survey form were sent to recognized cancer center chemotherapy units, and oncologists in these units were requested to complete the survey between October 22, 2015, and January 10, 2016. The invitation for the survey was restricted to a single oncologist from each unit. If a center had a single team, only one of the oncologists was contacted. The center's team was considered a single unit. If a center had multiple chemotherapy units that functioned independently of each other and had policies independent of each other, then one oncologist from the unit was invited and was considered an independent entity in the survey

In addition, the survey instrument in PDF format was distributed through personal contacts in a national biennial joint conference of the ISMPO (Indian Society of Medical and Pediatric Oncology) and the Indian Society of Oncology that was held from November 6 to 8, 2015, at Hotel Grand Hyatt, Mumbai, India. Only members from units that were not invited (for lack of a valid e-mail address), or who were invited but had not completed the survey online, were given the option of completing the survey at the conference.

Electronic responses were automatically captured in a Google spreadsheet that was linked to the online form, and responses collected on the PDF version were manually entered into the same sheet.

# **Survey Population**

The survey population consisted of adult oncology practices that administer chemotherapy on a regular basis. This included regional cancer centers, dedicated corporate cancer centers, cancer wings of medical colleges, hospitals, and private oncology day care centers. As much as possible, only a single oncologist was contacted from each center. If multiple oncologists from one center participated, they were asked to collaborate and submit a single response.

#### **Ethics**

The protocol was approved by the ICON ethics committee. ICON is an autonomous body of ISMPO, with a primary mandate for research.

#### Sample Size

The exact number of oncology centers in which chemotherapy is administered in the country is unknown. As convenience sampling was used for this survey, and formal sample size calculations were not performed.

# **Statistical Analysis**

To calculate the completeness of the implementation of guidelines, we counted the number of correct responses for each major question domain. The correct responses to these questions were decided before the start of the survey by the investigators (V.P. and K.P.) in accordance with the target guidelines. The correct response for each of the survey instrument questions is documented in the Appendix. A domain standard was considered to be fully implemented if > 90% of the items had correct responses for the given standard. It was considered partially implemented if between 50% and 90% of the items had correct responses for the given standard. When < 50% had correct responses for the given standard, it was considered not implemented. Thus, the formula for calculating the percentage implementation rate for each domain was:

% Implementation rate in given domain = (number of items with correct reponse/number of items for a given domain)  $\times$  100

The detailed scoring system and calculation of the percentage implementation rate is shown in the Appendix. A facility was considered to have fully implemented ASCO guidelines if it scored a percent implementation rate that exceeded 90% in the B (B1, B2, and B3 combine) domains. The percentage of 90% was decided by consensus among the investigators.

Descriptive data regarding frequencies of implementation for a given standard as well as the domain are presented. We have calculated the number and proportion of oncology centers that have fully implemented each standard as well as the full domain. Frequency of major reasons for nonimplementation of a given standard are presented. Linear regression analysis was performed to identify factors that predicted low implementation scores in B1, B2, and B3 domains.

#### **RESULTS**

#### **Baseline Details of Participating Centers**

Sixty-six (62.9%) of 105 centers participated in the survey. Details about these centers are listed in Table 1. The majority of these centers (60; 90.9%) were located in urban areas, were dedicated cancer centers (55; 83.3%), and were teaching institutes (45; 68.2%). The median number of patients seen per physician was 40 (interquartile range, 30 to 50 patients). The average number of patients seen per physician was 54.5 in the government sector, whereas it was 33.8 in the private sector (P= .009). Sixty-five (98.5%) of 66 centers were aware of the presence of international antiemetic guidelines.

# Implementation of Standards

The target of partial, full, and no implementation of standards was seen in 92.5% (95% CI, 83.0% to 97.0%), 4.5% (95% CI, 1.1% to 13.2%), and 3.0% (95% CI, 0.3% to 11.2%) of centers, respectively, as shown in Figure 1. Only two centers had all standards implemented fully, whereas one center had > 90% standards implemented. Full implementation was better for the low-minimal emetogenic regimens (34.8% of centers; 95% CI, 24.5% to 46.9%) than the highly emetogenic regimens (6.1% of centers; 95% CI, 2% to 15.1%).

Details about the implementation of each individual standard in each domain are listed in the Appendix (Appendix Tables A1 to A6). In the B1 domain (high antiemetic prophylaxis), the recommendations with lowest compliance were the use of olanzapine when aprepitant is not used (nine centers; 13.6%), appropriate use of 5HT3 antagonist on days 2 and 3 (13 centers; 19.7%), and use of dexamethasone on days 2 and 3 (44 centers; 66.7%). Similarly, in B2 domain, the appropriate use of 5HT3 and dexamethasone on days 2 and 3 was significantly lacking (Appendix Tables A2 and A3).

#### **Factors Adversely Impacting Implementation**

The three most frequently cited reasons for hampered implementation of ASCO guidelines in routine chemotherapy practice were a lack of sensitization (26 centers; 39.4%), lack of national guidelines (12 centers; 18.2%), and lack of administrative support (10 centers; 15.2%). (Appendix Table A7). None of the following factors—place of practice, funding source, presence of dedicated cancer center, and patient load—were independently associated with low implementation scores in B1, B2, or B3 domains (Appendix Tables A8 to A10).

**Table 1.** Baseline Details of Participating Centers

Characteristic	No. (%)
Location of practice	
Rural	6 (9.1)
Urban	60 (90.9)
Source of funding	
Government (public)	29 (43.9)
Private	37 (56.1)
Teaching institute	
Yes	45 (68.2)
No	21 (31.8)
Dedicated cancer center	
Yes	55 (83.3)
No	11 (16.7)
Chemotherapy facilities	
Day care only	5 (07.6)
Both inpatient and daycare	61 (92.4)
Awareness of international antiemetic prophylaxis guidelines	
Yes	65 (98.5)
No	1 (1.5)
Workload	
Median number of daily patient consultations per practitioner	40 (IQR, 30-50)

Abbreviation: IQR, interquartile range.

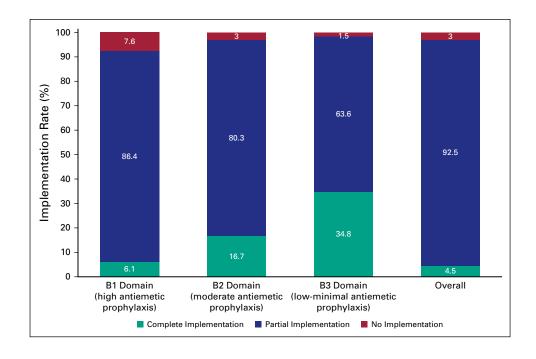
As a post hoc linear regression analysis failed to identify any single predictive factor, a composite regression tree analysis was performed using R

version 3.1.2 (R Foundation for Statistical Computing, Vienna, Austria) for B1 (high antiemetic prophylaxis); B2 (moderate antiemetic prophylaxis); and B3 (low-minimal antiemetic prophylaxis) domains independently with respect to the dependent variables (funding source [government or private]; center [academic or not]; location [urban or rural]; and type of cancer center [dedicated or nondedicated]). The lowest implementation rates were observed in the high antiemetic prophylaxis recommendations (B1 domain) in private rural centers. The lowest implementation rates were observed in the moderate antiemetic prophylaxis recommendations (B2 domain) in noncancer, dedicated government centers. The lowest implementation rates were observed in the low-minimal antiemetic prophylaxis recommendations (B3 domain) in noncancer, dedicated government centers and in private urban centers.

# **Institutional Antiemetic Policy Details**

A written institutional antiemetic policy was present in 27 participating centers (40.9%). Recommendations regarding high and moderate antiemetic prophylaxis were included in > 90% of institutional antiemetic policies. Recommendations regarding management of anticipatory and refractory CINV were present in 16 (59.3%) and 15 (55.6%) centers, respectively. The primary reasons for hampered implementation of an institutional antiemetic policy are listed in Table 2.

Fig 1. Implementation rate of ASCO guidelines in each of the domains. B1, B2, and B3 domains had question dealing with high, moderate, and low-minimal emetogenic agents respectively. Although overall calculation suggested ≥ 90% score in three centers, only two centers had consistent ≥ 90% scores in each domain. These two centers had 100% scores in all domains. The third center had 100% scores in B2 and B3 domains but had a score of 83.33% in the B1 domain.



**Knowledge and Practice in Special Situations** 

Details of the responses to special situations are listed in Appendix Tables A11 and A12). In situations that pertained to multiday regimens, 31.8% (21) of centers started antiemetics 1 day before the start of chemotherapy, 74.2% (49) of centers selected antiemetic protocol for each day on the basis of the emetogenic risk class of chemotherapy administered, and 63.6% (42) of centers continued antiemetic therapy for 2 days after the chemotherapy was completed. In protocols that pertained to chemoradiation, 82.8% (53) of centers selected antiemetics, taking into account the risk of emesis of both radiation and chemotherapy. Participating centers were divided in their protocols regarding the emetogenic risk of weekly cisplatin (30 to 40 mg/m<sup>2</sup>) administered concurrently with radiation. Of centers, 53.1% (34) considered it as highly emetogenic and the remaining considered this protocol moderately emetogenic.

#### **DISCUSSION**

The profile and distribution of cancer centers in India presents several unique challenges in managing the complications of chemotherapy, including

No (%)

Table 2. Details of Domain A: Institutional Antiemetic Policy

Characteristic

Characteristic	No. (%)
Number of centers having institutional antiemetic policy	
Yes	27 (40.9)
No	39 (59.1)
Details included in institutional policy	27
Recommendations for highly emetogenic agents	26 (96.3)
Recommendations for moderately emetogenic agents	25 (92.6)
Recommendations for low emetogenic agents	22 (81.5)
Recommendations for minimal emetogenic agents	14 (51.9)
Recommendations for anticipatory CINV	16 (59.3)
Recommendations for breakthrough CINV	18 (66.7)
Recommendations for refractory CINV	15 (55.6)
Factors hampering the development of institutional antiemetic policy	39*
Lack of administrative support	9 (23.1)
Lack of sensitization	21 (53.8)
Lack of national guidelines	3 (7.7)
Lack of consensus among practitioners	4 (10.3)
Practicing international guidelines	6 (15.4)
Funding constraints	5 (12.8)
High number of patients	7 (17.9)

Abbreviation: CINV, chemotherapy-induced nausea and vomiting.

CINV. Although 75% to 80% of the population stays in rural areas, cancer centers are predominantly located in major cities. <sup>18</sup> Thus, a majority of patients do not have ready access to medical care, and this adds to the challenge in deciding the appropriate antiemetic regimen. Compounding the issue is the average number of patients seen by individual oncologists, which has been reported to be much higher than in the West. <sup>19-22</sup> In fact, this factor was mentioned as one of the factors that hindered appropriate antiemetic prophylaxis (12.3% of centers).

Selection of the optimal antiemetic regimen consists of gauging the emetogenic potential of chemotherapy regimens and then deciding the appropriate antiemetic prophylaxis considering the factors that are unique to each country. Whereas international evidence-based guidelines have been formulated for the selection of appropriate antiemetic prophylaxis, there are minor variations depending on local oncologic practice. In general, treatment guidelines formulated in developed countries are difficult to implement in developing countries. 17 Unfortunately, many developing countries, such as India, do not have their own guidelines for antiemetic prophylaxis. Hence, most oncologists in India use international guidelines, such as the ASCO antiemetic guidelines. This is also reflected in the current study, where 98.5% of the responding centers were aware and had knowledge of these guidelines.

Overall, an encouraging finding in our study was the fact that 97.0% of centers had > 50% of ASCO antiemetic clinical guideline standards implemented in routine practice; however, only three centers implemented > 90% of standards and only two centers implemented all standards fully. Guidelines regarding high emetogenic prophylaxis were the least implemented (only four centers; 6.1%). Our survey identified three major areas of concern relating to the ASCO antiemetic guidelines: the absence of olanzapine when aprepitant is not used (86.4%), overuse of 5HT3 antagonist for delayed emesis (80.3%), and absence of dexamethasone for delayed emesis (33.3%). Whereas it may be argued that the ASCO antiemetic guidelines did not offer olanazapine as an option, a reference was made about its role in a scenario precluding aprepitant. 11 On this basis, the investigators decided that olanzapine is an essential component of an antiemetic regimen when aprepitant cannot be used. Overuse of 5HT3 antagonist for delayed emesis (71.1% in aprepitant users and 57.1% in nonusers) and inappropriate use of dexamethasone on days 2 and 3 postchemotherapy

<sup>\*</sup>Participants would choose more than a single option in response to this question and hence the total of responses is more than 39.

Fig 2. Evidence-based algorithm for quick selection of appropriate antiemetic regimen. (\*) NK-1 antagonist schedule: aprepitant 125 mg day 1 and 80 mg days 2 and 3 orally or fosaprepitant 150 mg IV day 1 only. (†) 5HT3 antagonist: granisetron 1 mg IV/oral or ondansetron 8 mg IV/oral or palonosetron 0.25 mg IV day 1 only. Emetogenic potential: High: AC/EC, carmustine  $(> 250 \text{ mg/m}^2)$ , cisplatin (any dose), cyclophosphamide  $(> 1.5 \text{ g/m}^2)$ , dacarbazine, doxorubicin ( $\geq$  60 mg/m<sup>2</sup>), epirubicin ( $> 90 \text{ mg/m}^2$ ), ifosfamide ( $\geq 2 \text{ g/m}^2$ ), mechlorethamine. Moderate (NK-1 antagonist preferred): carboplatin, carmustine ( $\leq 250 \text{ mg/m}^2$ ), dactinomycin, daunorubicin, doxorubicin (< 60 mg/m<sup>2</sup>), epirubicin ( $\leq$  90 mg/m<sup>2</sup>), ifosfamide  $(< 2 g/m^2)$ , irinotecan, methotrexate  $(\geq 250 \text{ mg/m}^2)$ . Moderate: cyclophosphamide  $(\leq 1.5 \text{ g/m}^2)$ , IFN-alpha  $(\geq 10 \text{ million U/m}^2),$ oxaliplatin, temozolomide. Low: carfilzomib, liposomal doxorubicin, etoposide, eribulin, FU, floxuridine, gemcitabine, INF-alpfa (> 5 to > 10 million units/m<sup>2</sup>),

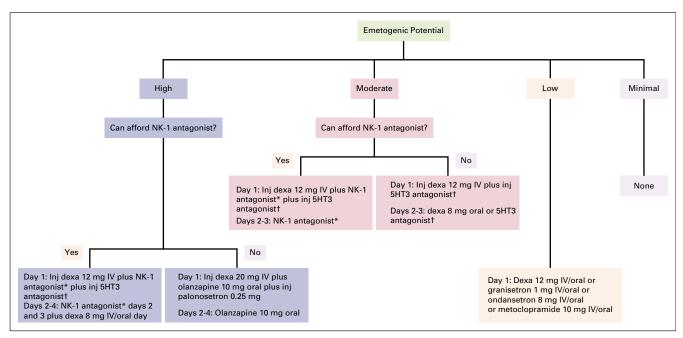
(42.1% overuse in aprepitant users and 67.0% underuse in aprepitant nonusers) were the major deficiencies in implementation of moderate antiemetic prophylaxis. Guidelines were fully implemented in the low and minimal risk setting in 90.9% and 42.4% centers, respectively. Of centers, 57.6% used antiemetics with agents that had minimal risk of emetogenesis. Overuse of 5HT3 antagonist for delayed emesis prophylaxis and underuse of dexamethasone for the same are the main issues in published work from other developed countries. <sup>23-26</sup>

Our survey also highlights the variable practices in oncology centers regarding antiemetics for multidrug chemotherapy and antiemetic prophylaxis for concurrent chemoradiation. One of the factors this survey did not touch upon was patient risk factor adjusted antiemetic regimens. It is a known fact that female patients and patients who had previous episodes of intractable vomiting are at a high risk of emesis and that modification in selection of antiemetic regimens might be warranted. <sup>13,27,28</sup>

As previously noted, the major factors that hindered wider implementation of ASCO antiemetic guidelines is the lack of sensitization, despite a majority of centers being aware of the existence of such guidelines. In this context, lack of sensitization means lack of concern, or apathy, regarding chemotherapy-induced nausea and vomiting. Awareness and knowledge unfortunately do not always translate into action, and emetic prophylaxis seems to be one example. The authors

therefore decided to organize a biannual continuous medical education program for practicing oncologists and oncology trainees on antiemetic prophylaxis under the aegis of ICON. The program would stress the recommendations which were minimally implemented as per our survey. We hope to improve the antiemetic prophylaxis for patients who receive chemotherapy in the country.

Another factor that impaired implementation of ASCO antiemetic guidelines was lack of national antiemetic prophylaxis guidelines. Presence of national guidelines or better institutional policy mandates physicians adhere to such guidelines or policies. These guidelines and policies are medico-legally and ethically binding. Therefore, it was decided by the authors to provide a simple, single-page algorithm for appropriate selection of antiemetic prophylaxis. Cost of antiemetic regimens was also factored in the algorithm. The algorithm was drafted by the authors (V.P. and K.P.) and was debated by the other members before a final algorithm was drafted (Fig 2). The algorithm is primarily for centers that do not have institutional antiemetic guidelines. As per our survey, 50% (33 centers) of responding centers belong to this category. In addition, the algorithm can be an effective supplement even for those centers where the antiemetic guidelines do not have recommendations for all situations as outlined in the algorithm. We plan to design comprehensive antiemetic guidelines for the Indian subcontinent in partnership with such Indian oncology associations as ICON and ISMPO.



ixabepilone, methotrexate  $(> 50 \text{ to} < 250 \text{ mg/m}^2),$ mitomycin, mitoxantrone, pemetrexed, topotecan, taxanes. Minimal: bevacizumab, bleomycin, cetuximab, methotrexate  $(\leq 50 \text{ mg/m}^2)$ , nivolumab, panitumumab, pertuzumab, ramucirumab, temsirolimus, trastuzumab, vinca alkaloids dexa dexamethasone; FU, fluorouracil; IFN, interferon; inj, injection; IV, intravenous; NK-1, neurokinin-1; PO, orally.

In conclusion, awareness regarding the ASCO antiemetic clinical guidelines is satisfactory in Indian oncology practices; however, there is a need for further sensitization of oncologists toward complete implementation of the guidelines in their

clinical practice. Developing national guidelines that are specific for India may help in the standardization of antiemetic regimens.

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# AUTHORS' DISCLOSURES OF POTENTIAL CONFLICTS OF INTEREST

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- 30. Reference deleted.
- 31. Reference deleted.

# **APPENDIX**

Table A1. Response to B1 Domain of Survey

	Response			
Domain B1 Question (n = 66)	Always	Usually	Rarely	Never
Do you routinely use aprepitant or fosaprepitant, unless contraindicated, for highly emetogenic antineoplastic agents?	33 (50.0)	25 (37.9)	7 (10.6)	1 (1.5)
	Correct respo	nse: 58 (87.9)	Incorrect resp	onse: 8 (12.1)
Do you routinely use olanzapine, unless contraindicated, for highly emetogenic antineoplastic agents if aprepitant or fosaprepitant is not used?	3 (4.5)	6 (9.1)	31 (47.0)	26 (39.4)
	Correct respo	onse: 9 (13.6)	Incorrect response	onse: 57 (86.4)
Do you routinely administer 5HT3 antagonist	59 (89.4)	6 (9.1)	0	1 (1.5)
(for example, ondansetron or granisetron), unless contraindicated, for highly emetogenic antineoplastic agents?	Correct response: 65 (98.5)		Incorrect response: 1 (1.5)	
Do you routinely administer dexamethasone, unless contraindicated, for highly emetogenic antineoplastic agents?	57 (86.4)	8 (12.1)	0	1 (1.5)
	Correct respo	nse: 65 (98.5)	Incorrect resp	ponse: 1 (1.5)
Do you prescribe 5HT3 antagonist on days 2 and 3 of single day chemotherapy regimen of high emetogenic potential?	37 (56.1)	16 (24.2)	5 (7.6)	8 (12.1)
	Incorrect response	onse: 53 (80.3)	Correct respo	nse: 13 (19.7)
Do you prescribe dexamethasone on days 2 and 3 of single day chemotherapy regimen of high emetogenic potential?	22 (33.3)	22 (33.3)	18 (27.3)	4 (6.1)
	Correct respo	nse: 44 (66.7)	Incorrect resp	onse:22 (33.3)

NOTE. Data are given as No. (%). The actual responses are shown and, in addition, the scoring of responses as correct and incorrect in accordance with guidelines is also shown.

Table A2. Response to B2 Domain of Survey

	Response				
Domain B2 Question (n = 38)	Always	Usually	Rarely	Never	
Do you routinely use aprepitant or fosaprepitant, unless contraindicated, for moderately emetogenic antineoplastic agents?	32 (84.2)	6 (16.8)	NA	NA	
	Correct respo	nse: 38 (100)	Incorrect re	sponse: NA	
Do you routinely administer 5HT3 antagonist (for example, ondansetron or granisetron), unless contraindicated, for moderately emetogenic antineoplastic agents?	30 (78.9)	8 (21.1)	0 (0)	0 (0)	
	Correct respo	nse: 38 (100)	Incorrect response: 0 (0		
Do you routinely administer dexamethasone, unless contraindicated, for moderately emetogenic antineoplastic agents?	25 (65.8)	11 (28.9)	2 (5.3)	0 (0)	
	Correct respon	nse: 36 (94.7)	Incorrect resp	onse: 2 (5.3)	
Do you prescribe 5HT3 antagonist on days 2 and 3 of single day chemotherapy regimen of moderately emetogenic potential?	12 (31.6)	15 (39.5)	7 (18.4)	4 (10.5)	
	Incorrect response: 27 (71.1)		Correct respon	nse: 11 (28.9)	
Do you prescribe dexamethasone on days 2 and 3 of single day chemotherapy regimen of moderately emetogenic potential?	3 (7.9)	13 (34.2)	17 (44.7)	5 (13.2)	
	Incorrect response	onse: 16 (42.1)	Correct respon	nse: 22 (57.9)	

NOTE. Data are given as No. (%). Table shows data regarding those centers who routinely administer NK-1 receptor antagonist in routine practice. The actual responses are shown, as is the scoring of responses as correct and incorrect in accordance with guidelines.

Abbreviation: NA, not applicable, as only centers that routinely administer NK-1 receptor antagonist are selected.

Table A3. Response to B2 Domain of Survey

	Response			
Domain B2 Question (n = 28)	Always	Usually	Rarely	Never
Do you routinely use aprepitant or fosaprepitant, unless contraindicated, for moderately emetogenic antineoplastic agents?	NA	NA	5 (17.9)	23 (82.1)
	Incorrect response: NA Correct response:		nse: 28 (100)	
Do you routinely administer 5HT3 antagonist (for example, ondansetron or granisetron), unless contraindicated, for moderately emetogenic antineoplastic agents?	22 (78.6)	6 (21.4)	0 (0)	0 (0)
	Correct response: 28 (100)		Incorrect res	sponse: 0 (0)
Do you routinely administer dexamethasone, unless contraindicated, for moderately emetogenic antineoplastic agents?	15 (53.6)	09 (32.1)	3 (10.7)	1 (3.6)
	Correct response: 24 (85.7)		Incorrect response: 4 (14.	
Do you prescribe 5HT3 antagonist on days 2 and 3 of single day chemotherapy regimen of moderately emetogenic potential?	5 (17.9)	11 (39.3)	6 (21.4)	6 (21.4)
	Incorrect response: 16 (57.1)		Correct respo	nse: 12 (42.9)
Do you prescribe dexamethasone on days 2 and 3 of single day chemotherapy regimen of moderately emetogenic potential?	3 (10.7)	6 (21.4)	13 (46.5)	6 (21.4)
	Correct respo	onse: 9 (32.1)	Incorrect response	onse: 19 (67.9)

NOTE. Data are given as No. (%). The table shows data regarding those centers that do not routinely administer NK-1 receptor antagonist in routine practice. The actual responses are shown as is the scoring of responses as correct and incorrect in accordance with guidelines. Abbreviation: NA, not applicable as only centers that routinely do not administer NK-1 receptor antagonist are selected.

Table A4. Guideline Implementation Rate in B2 Domain

		Implementation	
Domain	Complete (score > 90%)	Partial (score 50%-90%)	No Implementation (score 0%-49.9%)
B2 domain (with NK-1 receptor antagonist; n = 38)	10 (26.3)	28 (73.7)	0 (0)
B2 domain (without NK-1 receptor antagonist; n = 28)	1 (3.6)	25 (89.3)	2 (7.1)
B2 domain (total; n = 66)	11 (16.7)	53 (80.3)	2 (3.0)

NOTE. Data are given as No. (%).

Table A5. Choice of 5HT-3 Antagonist in B2 Domain

Choice of 5HT-3 Antagonist	No. (%)
Ondansetron	19 (28.8)
Granisetron	18 (27.3)
Palonosetron	29 (43.9)

Table A6. Response to B3 Domain of Survey

Response
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Domain B3 Question (n = 66)	Dexamethasone	5HT3 Antagonist	Domperidone	Aprepitant or Fosaprepitant	None
Optimal antiemetic regimen to prevent nausea and vomiting from low emetogenic antineoplastic agents	12 (18.2)	41 (62.1)	7 (10.6)	0	6 (9.1)
	Corr	ect response: 60 (90.	9)	Incorrect response: 6 (9	9.1)
Optimal antiemetic regimen to prevent nausea and vomiting from minimal emetogenic antineoplastic agents	6 (9.1)	17 (25.8)	15 (22.7)	0	28 (42.4)

Incorrect response: 38 (57.6) Correct response: 28 (42.4)

NOTE. Data are given as No. (%).

Table A7. Factors That Hamper Implementation of ASCO Guidelines in Routine Chemotherapy Practice

Factors That Hamper Implementation of ASCO Antiemetic Policy	(n = 66)*
Lack of administrative support	10 (15.2)
Lack of sensitization	26 (39.4)
Lack of national guidelines	12 (18.2)
Lack of consensus among practitioners	1 (01.5)
Practicing other international or institutional guidelines	5 (07.6)
Funding constraints	11 (16.7)
High number of patients	8 (12.1)

NOTE. Data are given as No. (%).

 Table A8.
 Impact of Various Factors on Center Ability to Implement Standards for High Emetogenic Prophylaxis

Variable	Mean	P
Location		
Urban	58.33	.327
Rural	64.72	
Source of funding		
Government (public)	66.09	.343
Private	62.61	
Teaching institute		
Yes	63.49	.660
No	64.44	
Dedicated cancer center		
Yes	63.94	.679
No	65.15	
Patient load (daily patients seen per physician)		
≤ 40	62.60	.563
> 40	66.67	

<sup>\*</sup>Participants would choose more than a single option in response to this question and, hence, the total of responses is more than 66.

Table A9. Impact of Various Factors on Center Ability to Implement Standards for Moderate Emetogenic Prophylaxis

Variable	Mean	P
Location		
Urban	70.67	.246
Rural	80.83	
Source of funding		
Government (public)	71.55	.835
Private	71.62	_
Teaching institute		
Yes	69.56	.221
No	75.95	
Dedicated cancer center		
Yes	71.73	.867
No	70.91	
Patient load (daily patients seen per physician)		
≤ 40	72.80	.958
> 40	69.60	

**Table A10.** Impact of Various Factors on Center Ability to Implement Standards for Low and Minimal Emetogenic Prophylaxis

Variable	Mean	P
Location		
Urban	65.00	.305
Rural	83.33	
Source of funding		
Government (public)	74.13	.157
Private	60.81	
Teaching institute		
Yes	70.00	.161
No	59.52	
Dedicated cancer center		
Yes	68.18	.334
No	59.09	
Patient load (daily patients seen per physician)		
≤ 40	68.29	.112
> 40	64.00	

Table A11. Factors That Prompt a Modification in the Prophylactic Antiemetic Regimen or Its Doses in Chemotherapy Practice

#### **Factor**

Situation	Age	Uncontrolled Comorbidities	Moderate Renal Dysfunction	Child Pugh B Liver Dysfunction	QTc Prolongation	No Knowledge
Modification of antiemetic regimen	25 (37.9)	36 (54.5)	21 (31.8)	18 (27.3)	34 (51.5)	8 (12.1)
Modification in doses of antiemetic agents regimen	22 (33.3)	36 (54.5)	27 (40.9)	23 (34.8)	29 (43.9)	9 (13.6)

NOTE. Data are given as No. (%). The participants would choose more than a single option in response to this question and hence the total of responses are more than 66. Abbreviation: QTc, corrected QT interval.

Table A12. Responses of Participants in Special Situations

Special Situation	No. (%)
Multiday chemotherapy, the optimal treatment of nausea and vomiting includes (n = 66)*	
Start antiemetics 1 day before	21 (31.8)
Each day antiemetics are selected on the basis of the emetogenic risk class of chemotherapy administered	49 (74.2)
Continue antiemetic for 2 days after chemotherapy is over	42 (63.6)
For chemoradiation, the selection of antiemetics takes into account which factors (n = $64$ )†	
Risk of emesis with radiation	1 (01.6)
Risk of emesis with chemotherapy	10 (15.6)
Both	53 (82.8)
Emetogenic potential regimen used while using concurrent weekly cisplatin (30-40 mg/m $^2$ ) with radiation (n = 64)†	
High	34 (53.1)
Moderate	30 (46.9)

<sup>\*</sup>Participants would choose more than a single option in response to this question and hence the total of responses are more than 66. †Two participants did not respond to chemoradiation-related survey items, hence the number is 64.