

Supplementary Table S1. Population characteristics according to low and normal hand grip strength based on the median (exploratory low HGS).

| Characteristics of patients | Level | Overall | Dynapenia ¹ | Normal HGS ¹ | <i>p</i> -value |
|--|--------------------------------|------------------|------------------------|-------------------------|-----------------|
| Total, <i>n</i> (%) | | 244 | 107 (43.9) | 137 (56.1) | |
| Sex, <i>n</i> (%) | Female | 109 (44.7) | 50 (46.7) | 59 (43.1) | 0.659 |
| | Male | 135 (55.3) | 57 (53.3) | 78 (56.9) | |
| Age, median (IQR) | | 69.0 (59.0-74.0) | 72.0 (68.0 - 78.0) | 63.0 (56.0 - 71.0) | <0.001 |
| BMI, median (IQR) | | 24.6 (21.5-28.6) | 24.9 (21.5 - 29.4) | 24.3 (21.6 - 27.5) | 0.522 |
| ECOG PS, <i>n</i> (%) | 0 | 66 (27.0) | 20 (18.7) | 46 (33.6) | 0.006 |
| | 1 | 150 (61.5) | 68 (63.6) | 82 (59.9) | |
| | 2 | 26 (10.7) | 17 (15.9) | 9 (6.6) | |
| | 3 | 2 (0.8) | 2 (1.9) | | |
| Serum albumin level, median (IQR) | | 39.0 (36.0-42.0) | 39.0 (34.0 - 41.5) | 39.0 (37.0 - 42.0) | 0.153 |
| CRP, median (IQR) | | 9.0 (4.0-33.2) | 11.0 (4.0 - 36.5) | 8.0 (3.0-33.0) | 0.121 |
| mGPS, <i>n</i> (%) | 0 | 118 (48.4) | 46 (43.0) | 72 (52.6) | 0.020 |
| | 1 | 87 (35.7) | 36 (33.6) | 51 (37.2) | |
| | 2 | 39 (16.0) | 25 (23.4) | 14 (10.2) | |
| Lymphopenia, <i>n</i> (%) | No | 226 (92.6) | 99 (92.5) | 127 (92.7) | 1.000 |
| | Yes | 18 (7.4) | 8 (7.5) | 10 (7.3) | |
| G8 score, median ² (IQR) | | 12.0 (11.0-15.0) | 12.0 (10.0 - 13.8) | 13.0 (11.0-15.0) | 0.050 |
| Primary tumor location, <i>n</i> (%) | Colon and rectum | 105 (43.2) | 55 (51.4) | 50 (36.8) | 55 (51.4) |
| | Stomach | 26 (10.7) | 6 (5.6) | 20 (14.7) | |
| | Esophagus | 18 (7.4) | 10 (9.3) | 8 (5.9) | |
| | Pancreas | 69 (28.4) | 28 (26.2) | 41 (30.1) | |
| | Others ³ | 25 (10.2) | 8 (7.5) | 17 (12.5) | |
| Stage, <i>n</i> (%) | Localized | 84 (34.4) | 39 (36.4) | 45 (32.8) | 0.707 |
| | Locally-advanced | 57 (23.4) | 26 (24.3) | 31 (22.6) | |
| | Metastatic | 103 (42.2) | 42 (39.3) | 61 (44.5) | |
| Number of metastatic sites, <i>n</i> (%) | 1 | 68 (65.4) | 28 (65.1) | 40 (65.6) | 1.000 |
| | ≥ 2 | 36 (34.6) | 15 (34.9) | 21 (34.4) | |
| Chemotherapy regimen, <i>n</i> (%) | 5FU + Oxaliplatin | 96 (39.3) | 46 (43.0) | 50 (36.5) | 0.072 |
| | 5FU + Irinotecan + Oxaliplatin | 69 (28.3) | 22 (20.6) | 47 (34.3) | |
| | 5FU alone | 24 (9.8) | 15 (14.0) | 9 (6.6) | |
| | Gemcitabine | 18 (7.4) | 9 (8.4) | 9 (6.6) | |
| | Others ⁴ | 37 (15.1) | 15 (14) | 22 (16.1) | |
| Biotherapy, <i>n</i> (%) | None | 204 (83.6) | 89 (83.2) | 115 (83.9) | 0.036 |
| | Bevacizumab | 26 (10.7) | 16 (15.0) | 10 (7.3) | |
| | Others ⁵ | 14 (5.7) | 2 (1.8) | 12 (8.8) | |

| | | | | | |
|--|-----|------------|-----------|------------|-------|
| Concomitant radiotherapy, <i>n</i> (%) | No | 222 (91.0) | 99 (92.5) | 123 (89.8) | 0.605 |
| | Yes | 22 (9.0) | 8 (7.5) | 14 (10.2) | |

Abbreviations: 5FU: 5 Fluorouracil; BMI: Body Mass Index; CRP: C-reactive protein; ECOG PS: Eastern Cooperative Oncology Group Criteria Performance Status; HGS: handgrip strength; IQR: interquartile range; mGPS: modified Glasgow prognosis score.

¹ According HGS cut-off based on the median in the population as HGS <34 for men and <22kg for women; ² Data available for 82 patients; ³ Other localizations: biliary tract (*n*=8), small intestine (*n*=7), ampulla of Vater (*n*=3), neuroendocrine tumor (*n*=4), appendix (*n*=1), anal (*n*=1), unknown primary (*n*=1); ⁴ Other chemotherapy: 5FU + Irinotecan (*n*=8), 5FU + Oxaliplatin + Docetaxel (*n*=9), 5FU + Cisplatin (*n*=1), 5FU + Dacarbazine (*n*=3), Carboplatin-Etoposide (*n*=1), Gemcitabine + Cisplatin (*n*=4), Gemcitabine + Oxaliplatin (*n*=3), Capecitabine + Oxaliplatin (*n*=7), Capecitabine + Mitomycin (*n*=1); ⁵ Other biotherapy: Panitumumab (*n*=10), Trastuzumab (*n*=4).

Supplementary Table S2. Association between low hand grip strength based on the median (exploratory low HGS) and chemotherapy-induced toxicities and DLT (detailed grades and therapeutic modifications).

| | Overall (<i>n</i> = 244) | Dynapenia ¹ (<i>n</i> = 23) | Normal HGS ¹ (<i>n</i> = 221) | <i>p</i> value |
|--------------------------------------|------------------------------|--|--|----------------|
| Neuropathy | | | | |
| Neuropathy (all grade), <i>n</i> (%) | 174 (94.6) | 66 (91.7) | 108 (96.4) | 0.193 |
| DLT, <i>n</i> (%) | 76 (41.3) | 26 (36.1) | 50 (44.6) | 0.285 |
| Toxicity grade 1, <i>n</i> (%) | 103 (58.9) | 43 (64.2) | 60 (55.6) | 0.334 |
| Toxicity grade 2, <i>n</i> (%) | 61 (34.9) | 19 (28.4) | 42 (38.9) | |
| Toxicity grade 3, <i>n</i> (%) | 11 (6.3) | 5 (7.5) | 6 (5.6) | |
| Median appearance, days (IQR) | 28.0 (14.0 - 42.0) | 28.0 (14.0 - 42.0) | 28.0 (14.0 - 42.2) | 0.619 |
| Decrease of dose, <i>n</i> (%) | 26 (14.1) | 12 (16.7) | 14 (12.5) | 0.717 |
| Discontinuation, <i>n</i> (%) | 64 (34.8) | 19 (26.4) | 45 (40.2) | 0.152 |
| Delay of chemotherapy, <i>n</i> (%) | 0 | | | |
| Asthenia | | | | |
| Toxicity (all grade), <i>n</i> (%) | 224 (91.8) | 104 (97.2) | 120 (87.6) | 0.014 |
| DLT, <i>n</i> (%) | 24 (9.8) | 16 (15.0) | 8 (5.8) | 0.029 |
| Toxicity grade 1, <i>n</i> (%) | 145 (64.4) | 59 (56.7) | 86 (71.1) | 0.071 |
| Toxicity grade 2, <i>n</i> (%) | 57 (25.3) | 31 (29.8) | 26 (21.5) | |
| Toxicity grade 3, <i>n</i> (%) | 23 (10.2) | 14 (13.5) | 9 (7.4) | |
| Median appearance, days (IQR) | 17.0 (14.0 - 39.0) | 17.0 (14.0 - 29.2) | 17.0 (14.0 - 42.0) | 0.687 |
| Decrease of dose, <i>n</i> (%) | 15 (6.1) | 9 (8.4) | 6 (4.4) | 0.002 |
| Discontinuation, <i>n</i> (%) | 7 (2.9) | 5 (4.7) | 2 (1.5) | 0.001 |
| Delay of chemotherapy, <i>n</i> (%) | 13 (5.3) | 9 (8.4) | 4 (2.9) | 0.001 |
| Diarrhea | | | | |
| Toxicity (all grade), <i>n</i> (%) | 139 (57.0) | 56 (52.3) | 83 (60.6) | 0.214 |
| DLT, <i>n</i> (%) | 20 (8.2) | 9 (8.4) | 11 (8.0) | 1.000 |
| Toxicity grade 1, <i>n</i> (%) | 82 (59.4) | 25 (44.6) | 57 (69.5) | 0.015 |
| Toxicity grade 2, <i>n</i> (%) | 40 (29.0) | 22 (39.3) | 18 (22.0) | |
| Toxicity grade 3, <i>n</i> (%) | 15 (10.9) | 8 (14.3) | 7 (8.5) | |
| Toxicity grade 4, <i>n</i> (%) | 1 (0.7) | 1 (1.8) | 0 | |
| Median appearance, days (IQR) | 16.0 (14.0 - 42.0) | 27.0 (14.0 - 42.0) | 15.0 (14.0 - 40.0) | 0.293 |
| Decrease of dose, <i>n</i> (%) | 13 (5.3) | 5 (4.7) | 8 (5.8) | 0.550 |
| Discontinuation, <i>n</i> (%) | 6 (2.5) | 4 (3.7) | 2 (1.5) | 0.242 |
| Delay of chemotherapy, <i>n</i> (%) | 7 (2.9) | 4 (3.7) | 3 (2.2) | 0.324 |
| Nausea³ | | | | |
| Toxicity (all grade), <i>n</i> (%) | 115 (56.9) | 41 (49.4) | 74 (62.2) | 0.084 |

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|-------------------------------------|--------------------|--------------------|--------------------|-------|
| DLT, <i>n</i> (%) | 4 (2.0) | 2 (2.4) | 2 (1.7) | 1.000 |
| Toxicity grade 1, <i>n</i> (%) | 76 (66.1) | 24 (58.5) | 52 (70.3) | 0.437 |
| Toxicity grade 2, <i>n</i> (%) | 35 (30.4) | 15 (36.6) | 20 (27.0) | |
| Toxicity grade 3, <i>n</i> (%) | 4 (3.5) | 2 (4.9) | 2 (2.7) | |
| Median appearance, days (IQR) | 15.0 (14.0 - 33.0) | 28.0 (14.0 - 42.0) | 14.5 (14.0 - 28.0) | 0.115 |
| Decrease of dose, <i>n</i> (%) | 4 (2.0) | 2 (2.4) | 2 (1.7) | 0.135 |
| Discontinuation, <i>n</i> (%) | 1 (0.5) | 1 (1.2) | 0 | 0.043 |
| Delay of chemotherapy, <i>n</i> (%) | 1 (0.5) | 0 | 1 (0.8) | 0.110 |
| Vomiting³ | | | | |
| Toxicity (all grade), <i>n</i> (%) | 51 (25.2) | 15 (18.1) | 36 (30.3) | 0.047 |
| DLT, <i>n</i> (%) | 4 (2.0) | 1 (1.2) | 3 (2.5) | 0.645 |
| Toxicity grade 1, <i>n</i> (%) | 35 (67.3) | 9 (60.0) | 26 (70.3) | 0.775 |
| Toxicity grade 2, <i>n</i> (%) | 14 (26.9) | 5 (33.3) | 9 (24.3) | |
| Toxicity grade 3, <i>n</i> (%) | 3 (5.8) | 1 (6.7) | 2 (5.4) | |
| Median appearance, days (IQR) | 38.0 (15.0 - 77.2) | 42.0 (15.0 - 74.0) | 28.0 (15.0 - 77.0) | 0.871 |
| Decrease of dose, <i>n</i> (%) | 2 (1.0) | 0 | 2 (1.7) | 0.034 |
| Discontinuation, <i>n</i> (%) | 2 (1.0) | 0 | 2 (1.7) | 0.034 |
| Delay of chemotherapy, <i>n</i> (%) | 2 (1.0) | 1 (1.2) | 1 (0.8) | 0.030 |
| Neutropenia | | | | |
| Toxicity (all grade), <i>n</i> (%) | 60 (24.6) | 26 (24.3) | 34 (24.8) | 1.000 |
| DLT, <i>n</i> (%) | 28 (11.5) | 10 (9.3) | 18 (13.1) | 0.172 |
| Toxicity grade 1, <i>n</i> (%) | 11 (18.0) | 7 (26.9) | 4 (11.4) | 0.086 |
| Toxicity grade 2, <i>n</i> (%) | 21 (34.4) | 6 (23.1) | 15 (42.9) | |
| Toxicity grade 3, <i>n</i> (%) | 24 (39.3) | 9 (34.6) | 15 (42.9) | |
| Toxicity grade 4, <i>n</i> (%) | 5 (8.2) | 4 (15.4) | 1 (2.9) | |
| Median appearance, days (IQR) | 37.5 (23.8 - 68.0) | 30.0 (19.2 - 51.0) | 41.5 (27.0 - 90.5) | 0.108 |
| Discontinuation, <i>n</i> (%) | 1 (0.4) | 0 | 1 (0.7) | 1.000 |
| Delay of chemotherapy, <i>n</i> (%) | 20 (8.2) | 8 (7.5) | 12 (8.8) | 0.901 |
| Anemia | | | | |
| Toxicity (all grade), <i>n</i> (%) | 165 (67.6) | 83 (77.6) | 82 (59.9) | 0.006 |
| DLT, <i>n</i> (%) | 6 (2.5) | 4 (3.7) | 2 (1.5) | 0.409 |
| Toxicity grade 1, <i>n</i> (%) | 102 (61.8) | 48 (57.8) | 54 (65.9) | 0.345 |
| Toxicity grade 2, <i>n</i> (%) | 48 (29.1) | 25 (30.1) | 23 (28.0) | |
| Toxicity grade 3, <i>n</i> (%) | 15 (9.1) | 10 (12.0) | 5 (6.1) | |
| Median appearance, days (IQR) | 15.0 (13.0 - 41.0) | 14.0 (13.0 - 29.5) | 26.5 (14.0 - 54.5) | 0.005 |
| Decrease of dose, <i>n</i> (%) | 6 (2.5) | 4 (3.7) | 2 (1.5) | 0.012 |

| | | | | |
|---------------------------------------|--------------------|--------------------|--------------------|-------|
| Discontinuation, <i>n</i> (%) | 0 | | | |
| Delay of chemotherapy, <i>n</i> (%) | 0 | | | |
| Thrombopenia | | | | |
| Toxicity (all grade), <i>n</i> (%) | 73 (29.9) | 32 (29.9) | 41 (29.9) | 1.000 |
| DLT, <i>n</i> (%) | 13 (5.3) | 5 (4.7) | 8 (5.8) | 0.779 |
| Toxicity grade 1, <i>n</i> (%) | 58 (79.5) | 27 (84.4) | 31 (75.6) | 0.288 |
| Toxicity grade 2, <i>n</i> (%) | 12 (16.4) | 3 (9.4) | 9 (22.0) | |
| Toxicity grade 3, <i>n</i> (%) | 3 (4.1) | 2 (6.2) | 1 (2.4) | |
| Median appearance, days (IQR) | 53.0 (27.0 - 90.0) | 43.0 (25.2 - 71.5) | 60.0 (28.0 - 98.0) | 0.161 |
| Decrease of dose, <i>n</i> (%) | 4 (1.6) | 1 (0.9) | 3 (2.2) | 0.805 |
| Discontinuation, <i>n</i> (%) | 1 (0.4) | 4 (3.7) | 1 (0.7) | 0.937 |
| Delay of chemotherapy, <i>n</i> (%) | 8 (3.3) | 4 (3.7) | 4 (2.9) | 0.969 |
| Oral Mucositis⁴ | | | | |
| Toxicity (all grade), <i>n</i> (%) | 29 (11.9) | 15 (14.0) | 14 (10.2) | 0.455 |
| DLT, <i>n</i> (%) | 3 (1.2) | 1 (0.9) | 2 (1.5) | 1.000 |
| Toxicity grade 1, <i>n</i> (%) | 22 (78.6) | 13 (86.7) | 9 (69.2) | 0.533 |
| Toxicity grade 2, <i>n</i> (%) | 3 (10.7) | 1 (6.7) | 2 (15.4) | |
| Toxicity grade 3, <i>n</i> (%) | 3 (10.7) | 1 (6.7) | 2 (15.4) | |
| Decrease of dose, <i>n</i> (%) | 3 (1.2) | 1 (0.9) | 2 (1.5) | 0.525 |
| Discontinuation, <i>n</i> (%) | 0 | | | |
| Delay of chemotherapy, <i>n</i> (%) | 0 | | | |
| Hand Foot Syndrome⁴ | | | | |
| Toxicity (all grade), <i>n</i> (%) | 10 (4.8) | 5 (5.4) | 5 (4.3) | 0.629 |
| DLT, <i>n</i> (%) | 4 (1.9) | 2 (2.2) | 2 (1.7) | 1.000 |
| Toxicity grade 1, <i>n</i> (%) | 3 (30.0) | 2 (40.0) | 1 (20.0) | 0.282 |
| Toxicity grade 2, <i>n</i> (%) | 5 (50.0) | 3 (60.0) | 2 (40.0) | |
| Toxicity grade 3, <i>n</i> (%) | 2 (20.0) | 0 | 2 (40.0) | |
| Median appearance, days (IQR) | 53.0 (40.0 - 68.8) | 54.0 (37.0 - 70.0) | 52.0 (43.0 - 65.0) | 0.917 |
| Decrease of dose, <i>n</i> (%) | 3 (1.4) | 1 (1.1) | 2 (1.7) | 0.733 |
| Discontinuation, <i>n</i> (%) | 1 (0.5) | 1 (1.1) | 0 | 0.531 |
| Delay of chemotherapy, <i>n</i> (%) | 0 | | | |

Abbreviation: DLT: Dose Limiting Toxicity; HGS: Hand Grip Strength; IQR: Interquartile

¹ HGS cut-off based on the median in the population as HGS <34 for men and <22kg for women (exploratory low HGS) ; ² Only patients receiving neurotoxic chemotherapy (*n*=184), and graded according Levi scale ; ³ Patients receiving 5FU and gemcitabine alone were not analyzed for this adverse effect (*n*=202) ; ⁴ Only patients receiving 5FU- or capecitabine-based chemotherapy regimen (*n*=210).

Supplementary Table S3. Association between dynapenia based on EWSGOP2 criteria and chemotherapy-induced toxicities and DLT (detailed grades and therapeutic modifications).

| | Overall (<i>n</i> = 244) | Dynapenia ¹ (<i>n</i> = 23) | Normal HGS ¹ (<i>n</i> = 221) | <i>p</i> value |
|--------------------------------------|------------------------------|--|--|----------------|
| Neuropathy² | | | | |
| Neuropathy (all grade), <i>n</i> (%) | 174 (94.6) | 14 (93.3) | 160 (94.7) | 0.582 |
| DLT, <i>n</i> (%) | 76 (41.3) | 7 (46.7) | 69 (40.8) | 0.786 |
| Toxicity grade 1, <i>n</i> (%) | 103 (58.9) | 8 (53.3) | 95 (59.4) | 0.116 |
| Toxicity grade 2, <i>n</i> (%) | 61 (34.9) | 4 (26.7) | 57 (35.6) | |
| Toxicity grade 3, <i>n</i> (%) | 11 (6.3) | 3 (20.0) | 8 (5.0) | |
| Median appearance, days (IQR) | 28.0 (14.0 - 42.0) | 18.0 (14.0 - 42.0) | 28.0 (14.0 - 42.0) | 0.274 |
| Decrease of dose, <i>n</i> (%) | 26 (14.1) | 6 (40.0) | 20 (11.8) | 0.025 |
| Discontinuation, <i>n</i> (%) | 64 (34.8) | 3 (20.0) | 61 (36.1) | 0.335 |
| Delay of chemotherapy, <i>n</i> (%) | 0 | | | |
| Asthenia | | | | |
| Toxicity (all grade), <i>n</i> (%) | 224 (91.8) | 23 (100.0) | 201 (91.0) | 0.303 |
| DLT, <i>n</i> (%) | 24 (9.8) | 5 (21.7) | 19 (8.6) | 0.059 |
| Toxicity grade 1, <i>n</i> (%) | 145 (64.4) | 11 (47.8) | 134 (66.3) | 0.156 |
| Toxicity grade 2, <i>n</i> (%) | 57 (25.3) | 8 (34.8) | 49 (24.3) | |
| Toxicity grade 3, <i>n</i> (%) | 23 (10.2) | 4 (17.4) | 19 (9.4) | |
| Median appearance, days (IQR) | 17.0 (14.0 - 39.0) | 15.0 (14.0 - 28.0) | 19.0 (14.0 - 40.8) | 0.469 |
| Decrease of dose, <i>n</i> (%) | 15 (6.1) | 3 (13.0) | 12 (5.4) | 0.093 |
| Discontinuation, <i>n</i> (%) | 7 (2.9) | 2 (8.7) | 5 (2.3) | 0.062 |
| Delay of chemotherapy, <i>n</i> (%) | 13 (5.3) | 3 (13.0) | 10 (4.5) | 0.071 |
| Diarrhea | | | | |
| Toxicity (all grade), <i>n</i> (%) | 139 (57.0) | 12 (52.2) | 127 (57.5) | 0.693 |
| DLT, <i>n</i> (%) | 20 (8.2) | 2 (8.7) | 18 (8.1) | 1.000 |
| Toxicity grade 1, <i>n</i> (%) | 82 (59.4) | 5 (41.7) | 77 (61.1) | 0.071 |
| Toxicity grade 2, <i>n</i> (%) | 40 (29.0) | 4 (33.3) | 36 (28.6) | |
| Toxicity grade 3, <i>n</i> (%) | 15 (10.9) | 2 (16.7) | 13 (10.3) | |
| Toxicity grade 4, <i>n</i> (%) | 1 (0.7) | 1 (8.3) | | |
| Median appearance, days (IQR) | 16.0 (14.0 - 42.0) | 20.0 (14.0 - 29.0) | 15.0 (14.0 - 42.0) | 0.915 |
| Decrease of dose, <i>n</i> (%) | 13 (5.3) | 0 | 13 (5.9) | 0.760 |
| Discontinuation, <i>n</i> (%) | 6 (2.5) | 1 (4.3) | 5 (2.3) | 0.482 |
| Delay of chemotherapy, <i>n</i> (%) | 7 (2.9) | 1 (4.3) | 6 (2.7) | 0.367 |
| Nausea³ | | | | |
| Toxicity (all grade), <i>n</i> (%) | 115 (56.9) | 9 (52.9) | 106 (57.3) | 0.801 |

| | | | | |
|-------------------------------------|--------------------|--------------------|--------------------|-------|
| DLT, <i>n</i> (%) | 4 (2.0) | 1 (5.9) | 3 (1.6) | 0.298 |
| Toxicity grade 1, <i>n</i> (%) | 76 (66.1) | 6 (66.7) | 70 (66.0) | 0.380 |
| Toxicity grade 2, <i>n</i> (%) | 35 (30.4) | 2 (22.2) | 33 (31.1) | |
| Toxicity grade 3, <i>n</i> (%) | 4 (3.5) | 1 (11.1) | 3 (2.8) | |
| Median appearance, days (IQR) | 15.0 (14.0 - 33.0) | 16.0 (15.0 - 42.0) | 15.0 (14.0 - 29.0) | 0.561 |
| Decrease of dose, <i>n</i> (%) | 4 (2.0) | 1 (5.9) | 3 (1.6) | 0.283 |
| Discontinuation, <i>n</i> (%) | 1 (0.5) | 0 | 1 (0.5) | 0.818 |
| Delay of chemotherapy, <i>n</i> (%) | 1 (0.5) | 0 | 1 (0.5) | 0.818 |
| Vomiting³ | | | | |
| Toxicity (all grade), <i>n</i> (%) | 51 (25.2) | 3 (17.6) | 48 (25.9) | 0.605 |
| DLT, <i>n</i> (%) | 4 (2.0) | 0 | 4 (2.2) | 1.000 |
| Toxicity grade 1, <i>n</i> (%) | 35 (67.3) | 1 (33.3) | 34 (69.4) | 0.102 |
| Toxicity grade 2, <i>n</i> (%) | 14 (26.9) | 1 (33.3) | 13 (26.5) | |
| Toxicity grade 3, <i>n</i> (%) | 3 (5.8) | 1 (33.3) | 2 (4.1) | |
| Median appearance, days (IQR) | 38.0 (15.0 - 77.2) | 39.0 (27.0 - 48.5) | 37.0 (15.0 - 78.0) | 0.921 |
| Decrease of dose, <i>n</i> (%) | 2 (1.0) | 0 | 2 (1.1) | 0.637 |
| Discontinuation, <i>n</i> (%) | 2 (1.0) | 0 | 2 (1.1) | 0.637 |
| Delay of chemotherapy, <i>n</i> (%) | 2 (1.0) | 0 | 2 (1.1) | 0.637 |
| Neutropenia | | | | |
| Toxicity (all grade), <i>n</i> (%) | 60 (24.6) | 3 (13.0) | 57 (25.8) | 0.286 |
| DLT, <i>n</i> (%) | 28 (11.5) | 0 | 28 (12.7) | 0.086 |
| Toxicity grade 1, <i>n</i> (%) | 11 (18.0) | 2 (66.7) | 9 (15.5) | 0.028 |
| Toxicity grade 2, <i>n</i> (%) | 21 (34.4) | 0 | 21 (36.2) | |
| Toxicity grade 3, <i>n</i> (%) | 24 (39.3) | 0 | 24 (41.4) | |
| Toxicity grade 4, <i>n</i> (%) | 5 (8.2) | 1 (33.3) | 4 (6.9) | |
| Median appearance, days (IQR) | 37.5 (23.8 - 68.0) | 26.0 (23.0 - 28.5) | 39.0 (25.0 - 68.0) | 0.277 |
| Discontinuation, <i>n</i> (%) | 11 (4.5) | 0 | 11 (5.0) | 0.522 |
| Delay of chemotherapy, <i>n</i> (%) | 20 (8.2) | 0 | 20 (9.0) | 0.373 |
| Delay of chemotherapy, <i>n</i> (%) | 1 (0.4) | 0 | 1 (0.5) | 0.369 |
| Anemia | | | | |
| Toxicity (all grade), <i>n</i> (%) | 165 (67.6) | 19 (82.6) | 146 (66.1) | 0.238 |
| DLT, <i>n</i> (%) | 6 (2.5) | 1 (4.3) | 5 (2.3) | 0.451 |
| Toxicity grade 1, <i>n</i> (%) | 102 (61.8) | 13 (68.4) | 89 (61.0) | 0.695 |
| Toxicity grade 2, <i>n</i> (%) | 48 (29.1) | 4 (21.1) | 44 (30.1) | |
| Toxicity grade 3, <i>n</i> (%) | 15 (9.1) | 2 (10.5) | 13 (8.9) | |
| Median appearance, days (IQR) | 15.0 (13.0 - 41.0) | 14.0 (14.0 - 41.0) | 15.5 (13.0 - 41.0) | 0.912 |

| | | | | |
|---------------------------------------|--------------------|--------------------|--------------------|-------|
| Decrease of dose, <i>n</i> (%) | 6 (2.5) | 1 (4.3) | 5 (2.3) | 0.189 |
| Discontinuation, <i>n</i> (%) | 0 | | | |
| Delay of chemotherapy, <i>n</i> (%) | 0 | | | |
| Thrombopenia | | | | |
| Toxicity (all grade), <i>n</i> (%) | 73 (29.9) | 10 (43.5) | 63 (28.5) | 0.235 |
| DLT, <i>n</i> (%) | 13 (5.3) | 2 (8.7) | 11 (5.0) | 0.352 |
| Toxicity grade 1, <i>n</i> (%) | 58 (79.5) | 8 (80.0) | 50 (79.4) | 0.591 |
| Toxicity grade 2, <i>n</i> (%) | 12 (16.4) | 1 (10.0) | 11 (17.5) | |
| Toxicity grade 3, <i>n</i> (%) | 3 (4.1) | 1 (10.0) | 2 (3.2) | |
| Median appearance, days (IQR) | 53.0 (27.0 - 90.0) | 55.5 (31.2 - 85.8) | 52.0 (27.0 - 88.0) | 0.810 |
| Decrease of dose, <i>n</i> (%) | 4 (1.6) | 0 | 4 (1.8) | 0.201 |
| Discontinuation, <i>n</i> (%) | 1 (0.4) | 0 | 1 (0.5) | 0.227 |
| Delay of chemotherapy, <i>n</i> (%) | 8 (3.3) | 2 (8.7) | 6 (2.7) | 0.121 |
| Oral Mucositis⁴ | | | | |
| Toxicity (all grade), <i>n</i> (%) | 29 (11.9) | 2 (8.7) | 27 (12.2) | 0.836 |
| DLT, <i>n</i> (%) | 3 (1.2) | 0 | 3 (1.4) | 1.000 |
| Toxicity grade 1, <i>n</i> (%) | 22 (78.6) | 2 (100.0) | 20 (76.9) | 0.745 |
| Toxicity grade 2, <i>n</i> (%) | 3 (10.7) | 0 | 3 (11.5) | |
| Toxicity grade 3, <i>n</i> (%) | 3 (10.7) | 0 | 3 (11.5) | |
| Decrease of dose, <i>n</i> (%) | 3 (1.2) | 0 | 3 (1.4) | 0.805 |
| Discontinuation, <i>n</i> (%) | 0 | | | |
| Delay of chemotherapy, <i>n</i> (%) | 0 | | | |
| Hand Foot Syndrome⁴ | | | | |
| Toxicity (all grade), <i>n</i> (%) | 10 (4.8) | 4 (18.2) | 6 (3.2) | 0.007 |
| DLT, <i>n</i> (%) | 4 (1.9) | 2 (9.1) | 2 (1.1) | 0.075 |
| Toxicity grade 1, <i>n</i> (%) | 3 (30.0) | 1 (25.0) | 2 (33.3) | 0.329 |
| Toxicity grade 2, <i>n</i> (%) | 5 (50.0) | 3 (75.0) | 2 (33.3) | |
| Toxicity grade 3, <i>n</i> (%) | 2 (20.0) | 0 | 2 (33.3) | |
| Median appearance, days (IQR) | 53.0 (40.0 - 68.8) | 62.0 (49.8 - 96.2) | 47.5 (40.0 - 61.8) | 0.394 |
| Decrease of dose, <i>n</i> (%) | 3 (1.4) | 1 (4.5) | 2 (1.1) | 0.007 |
| Discontinuation, <i>n</i> (%) | 1 (0.5) | 1 (4.5) | 0 | 0.001 |
| Delay of chemotherapy, <i>n</i> (%) | 0 | | | |

Abbreviation: DLT: Dose Limiting Toxicity; HGS: Hand Grip Strength; IQR: Interquartile

¹ Dynapenia was defined according the EWGSOP2 as HGS <27kg for men and <16kg for women; ² Only patients receiving neurotoxic chemotherapy (*n*=184), and graded according to Levi scale; ³ Patients receiving 5FU and gemcitabine alone were not analyzed for this adverse effect (*n*=202); ⁴ Only patients receiving 5FU- or capecitabine-based chemotherapy regimen (*n*=210).