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Title: The relationship between Malnutrition Universal Screening Tool (MUST), CTderived body composition, systemic inflammation and clinical outcomes in patients undergoing surgery for colorectal cancer

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List of abbreviations:

ASA: American Society of Anesthesiologists

BMI: body mass index

mGPS: modified Glasgow Prognostic Score MUST: malnutrition universal screening tool NLR: Neutrophil-to-lymphocyte Ratio SIR: Systemic Inflammatory Response. SMD: skeletal muscle radiodensity. SMI: skeletal muscle index TNM stage: Tumour, Node, Metastasis 1 Abstract

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Background: Nutritional status is an important factor affecting a patient's clinical outcomes.
Early identification of patients who are at risk of malnutrition is important to improve clinical
outcomes and reduce health cost. Malnutrition universal screening tool (MUST) has been
recommended as part of the routine nursing assessment for all patients at hospital admission.

Objective: The aim of this study was to examine the association between nutritional status
(MUST), systemic inflammatory response (SIR), body composition, and clinical outcomes in
patients undergoing surgery for colorectal cancer.

Methods: The malnutrition risk was examined using MUST in patients admitted for surgery for colorectal cancer between March 2013 and June 2016. Pre-operative CT scans were used to define the body composition. The presence of SIR was evidenced by the modified Glasgow prognostic score (mGPS) and neutrophil to lymphocyte ratio (NLR). Post-operative complications, severity of complication, length of hospital stay and mortality were considered as outcome measures.

Result: The study included 363 patients (199 males, 164 females), 21% of the patients 16 17 presented with a medium or high nutritional risk. There were significant associations between 18 MUST and subcutaneous adiposity (p<0.001), visceral obesity (p<0.001) and low SMI 19 (p<0.001). No statistically significant association was identified between MUST score and 20 presence of any complication or severity of complication. On multivariate analysis, MUST 21 remained independently associated with the length of hospital stay [OR=2.17 (95% CI 22 1.45,3.26) p<0.001]. Kaplan–Meier survival curves showed an increased number of deaths 23 for patients at medium or high risk of malnutrition (p<0.001). This association was found to 24 be independent of other confounding factors [HR=1.45 (95% CI 1.06,1.99) p=0.020].

Conclusion: MUST score is an independent marker of risk in those undergoing surgery for
 colorectal cancer surgery and should remain a key part of preoperative assessment.

- 27 Keywords: MUST, nutrition status, body composition, systemic inflammatory response,
- 28 colorectal cancer

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31 Introduction

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Colorectal cancer is the third most common cancer and the fourth leading cause of cancerrelated deaths worldwide, and its burden is expected to increase by 60% to more than 2.2 million new cases and 1.1 million cancer deaths by 2030 (1). In the United Kingdom in 2016, more than 41,000 patients are diagnosed and almost 16,500 patients die from colorectal cancer (2). Postoperative complications occur in up to one-third of patients undergoing colorectal procedures (3). Patients with complications have been shown to be at higher risk for mortality (4).

Nutritional status is an important factor affecting patient's clinical outcomes. Malnutrition
has been shown to be associated with adverse postoperative outcomes of morbidity and
mortality in patients with gastrointestinal cancer (5,6). Indeed, the presence of malnutrition
can alter immune responses and impair wound healing in surgical patients (7).

44 Early identification of patients who are malnourished or at risk of becoming malnourished, 45 and those who would benefit from specific nutritional support, is vital to reduce the risk of 46 surgical complications, improve clinical outcomes and reduce health cost. According to the 47 British Association for Parenteral and Enteral Nutrition (BAPEN) recommendation, the screening method should be: simple and understandable, rapid to implement, and validated 48 49 for hospital use (8). Several nutritional screening tools have been developed for this purpose, 50 including the Malnutrition Universal Screening Tool (MUST). MUST identifies patients who 51 are malnourished or at risk of developing malnutrition based on assessment of body mass 52 index (BMI), unintentional weight loss in the preceding three to six months, and the presence 53 of an acute disease resulting in absence of dietary intake for more than five days (9) 54 Subsequent management guidelines have followed, based on the overall malnutrition risk 55 (MUST) score.

| 56 | Published studies have been conducted to assess nutritional status in oncologic patients using |
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| 57 | different nutritional screening tools such as subjective global assessment (SGA), malnutrition |
| 58 | screening tool (MST) and nutritional risk screening (NRS 2002) (10). MUST is now used in |
| 59 | most of the UK as part of the routine nursing assessment for all patients admitted to hospital |
| 60 | (11), However, to our knowledge no studies have examined the relationship between MUST, |
| 61 | body composition, and systemic inflammatory responses. The aim of the present study was to |
| 62 | examine these associations, and to relate MUST score to clinical outcomes after surgery for |
| 63 | colorectal cancer. |
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81 **Patients and Methods**

82 Patients

83 Consecutive patients who underwent potentially curative resection for colorectal cancer 84 between March 2013 and June 2016 were included. All procedures were performed at Glasgow Royal Infirmary. Those patients with a preoperative recorded MUST score, 85 86 preoperative CT scan, and documentation reporting the presence or absence of post-operative complications were included. Data regarding the nature, severity and management of 87 88 complications were retrospectively classified using the Clavien-Dindo grade, where 0 is no 89 complication, 1 is any deviation from the normal post-operative course without the need for 90 pharmacological treatment or surgical, endoscopic and radiological interventions, 2 is 91 requiring pharmacological treatment with drugs other than such allowed for grade 1 92 complications, 3 is requiring surgical, endoscopic or radiological intervention, 4 is a life 93 threatening complication (including CNS complications) requiring IC/ICU management, 5 is 94 the death of a patient (12). Patient comorbidity was classified using the American Society of 95 Anesthesiologists (ASA) grading system. ASA 1 a patient with normal health, ASA 2 a 96 patient with mild systemic disease, ASA 3 a patient with severe systemic disease and ASA 4 97 a patient with severe systemic disease that is a constant threat to life.

All data were collected prospectively in a database, anonymised, and were subsequently
analysed. Any uncertainties were addressed by review of electronic and/or physical case
notes.

101 Malnutrition risk assessment

102 The MUST scores were recorded in patient notes, using a dedicated pro forma, by clinical 103 nursing staff, within 24 h of admission. MUST incorporates three components to determine 104 the overall risk for malnutrition (Figure1): current weight status using BMI, unintentional 105 weight loss, and acute disease effect that has induced a phase of nil per mouth for > 5 days.

Each parameter can be rated as 0, 1, or 2. Overall risk for malnutrition is established as low (score = 0), medium (score = 1), or high (score \geq 2).

108 Body composition

109 CT images were obtained at the level of the third lumbar vertebra. Patients whose scans were 110 taken 3 months or more prior to their surgery were excluded from the study. The median and range for the interval between CT scanning and operation was 0.91 months (0.03-2.83) with 111 78% of scans carried out within 30 days of operation. Scans with significant movement 112 artefact or missing region of interest were not considered for inclusion. Each image was 113 analysed using a free-ware program [NIH Image J version 1.47 (24)]. Region of interest 114 115 (ROI) measurements were made of visceral adipose tissue, subcutaneous adipose tissue and skeletal muscle areas (cm²) using standard Hounsfield Unit (HU) ranges (adipose tissue -190 116 to -30, and skeletal muscle -29 to +150). These were then normalised for height² to create 117 indices; total adipose tissue index (cm²/m²), subcutaneous adipose tissue index (cm²/m²), 118 visceral adipose tissue index (cm^2/m^2) , and skeletal muscle index (SMI, cm^2/m^2). Skeletal 119 muscle radiodensity (SMD, HU) was measured from the same ROI used to calculate SMI, as 120 its mean HU. Visceral obesity was defined as VFA >160 cm² for male patients and >80 cm² 121 for female patient (13). High subcutaneous adipose tissue index was defined as >50.0 cm² 122 $/m^2$ in males and ≥ 42.0 cm² $/m^2$ in females (14). Sarcopenia was described by Caan and 123 colleagues as an SMI<52.3 cm²/m² if BMI<30kg/m² and SMI<54.3 cm²/m² if BMI>30kg/m² 124 in male patients and an SMI<38.6 cm^2/m^2 if BMI<30kg/m² and an SMI<46.6 cm^2/m^2 if 125 BMI>30kg/m² in female patients (15). Low SMD was defined by SMD <41HU in patients 126 with BMI <25kg/m² and <33HU in patients with BMI >25kg/m² (25). 127

128 Inflammatory markers

129 Serum CRP (mg/L) and albumin (g/L) concentrations were measured by routine laboratory

130 procedures with an automated analyzer (Architect; Abbott Diagnostics, Maidenhead, UK).

131 The limit of detection for CRP was 0.1 mg/L. Intraassay imprecision was <4%. The 132 laboratory participated in external quality assurance/ proficiency testing programs, the A, B 133 and C scores were within the EQA (NEQAS) targets throughout the time period of the study. 134 Performance was acceptable throughout, which indicated that methodologic changes did not result in any bias. The presence of preoperative systemic inflammatory response was 135 136 evidenced by an mGPS and NLR. The mGPS was derived as the following; patients with a normal C-reactive protein (<10mg/L) were allocated a score of 0, those with an elevated C-137 138 reactive protein (>10 mg/L) allocated a score 1, and those with an elevated C-reactive protein 139 (>10 mg/L) and hypoalbuminaemia (<35 g/L) were allocated a score of 2 (16). The 140 neutrophil lymphocyte ratio (NLR) was calculated for each patient for whom preoperative 141 neutrophil and lymphocyte counts were available. Values < than 3 considered normal, 3-5 142 moderate and >5 were considered raised (26).

143 Survival

144 The cause and date of death were confirmed with the Registrar General (Scotland) until 30 145 June 2018, which served as the censor date. Informed consent was obtained from patients 146 prior to surgery. Ethical approval was granted by the West of Scotland Research Ethics 147 Committee, Glasgow.

148 Statistical analysis:

In the present study no formal power calculation was carried out since low SMI has been shown to be associated with overall survival (primary endpoint) in smaller and similar sized cohorts of patients undergoing surgery for colorectal cancer (19, 27). Secondary outcome measures were length of hospital stay and post-operative complications

153 Categorical variables were analysed using Chi-square test for linear-by-linear association.
154 Where there was a significant association on Chi-square analysis, pairwise comparisons were
155 carried out to detect where the differences in proportions were. Missing data were excluded

156 from analysis on a variable by variable basis. Due to the number of statistical comparisons 157 carried out (~40) a P value <0.001 was considered statistically significant. Mortality within 158 30 days of the index procedure or during the index admission were excluded from subsequent survival analysis. The time between the date of surgery and the date of death of any cause 159 160 was used to define overall survival (OS). The Cox regression hazard model was applied to 161 assess the ability of the MUST to predict survival. Those variables associated to a degree of 162 p<0.1 at univariate analysis were entered into a backward conditional multivariate model. 163 Length of hospital stay was classified as binary variable (<7 vs >7 days) and then analysed 164 using univariate and multivariate binary logistic regression.

165 Statistical analysis was performed using SPSS software (Version 21.0. SPSS Inc., Chicago,

166 IL, USA).

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- 172 **Results:**
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174 In total 483 patients were identified as having undergone potentially curative surgery for 175 colorectal cancer. Of these, 120 were excluded due to missing MUST score, or clinical and 176 pathological data. A total of 363 patients were included in the final analysis. There were 199 males and 164 females. The mean (\pm SD) age was 66 \pm 12 y and BMI 27.6 \pm 6.4. The 177 178 majority of patients had ASA grade 1 and 2 comorbidity (69%), colon cancer (62%) and had TNM stage 0-2 disease (64%). The majority of patients had subcutaneous (63%) and visceral 179 180 (72%) adiposity, low SMI (50%) and low SMD (61%). The majority of patients had a mGPS 181 (76%) and NLR (51%) in the normal range. The majority of patients had no complication 182 type (63%) and severity measured by Clavien-Dindo grade (66%), had a hospital stay more 183 than 7 days (54%) and were alive at 3 years (79%).

184 MUST and patients' characteristics

185 A total of 288 patients had a MUST 0 (79%, Low risk), 31 had MUST 1 (9%, medium risk),

186 and 44 had a MUST ≥ 2 (12%, high risk).

187 The associations of nutritional status classified by MUST with patients' characteristics, body composition, systemic inflammatory response and colorectal cancer outcomes are presented 188 189 in Table 1. Since TNM stage 4 disease is commonly associated with malnutrition risk the 190 analysis was repeated after excluding the patients (n=13). The results of the analysis did not 191 change materially. There were significant associations between MUST, age and BMI. 192 Around 39% of patients with MUST score 2 were older than 74y (p<0.001), and BMI: All 193 underweight patients had MUST score ≥ 1 and only 7% of patients with MUST score 2 were 194 obese.

195 There were no association with sex, ASA, tumour site and stage.

196 MUST and body composition

There were significant associations between MUST and subcutaneous adiposity (p<0.001), visceral obesity (p<0.001) and low SMI (p<0.001). Specifically, 42% of patients with a MUST score 2 had subcutaneous adiposity compared with 86% of patients with a MUST score of 0 (p<0.001). Also, 38% of patients with MUST score 2 had visceral obesity compared with 80% of patients with a MUST score 0 (p<0.001). Finally, 76% of patients with MUST score 2 had low SMI compared with 45% of patients with a MUST score 0 (p=0.001).

204 MUST and systemic inflammatory response

There were significant associations between MUST and mGPS (p<0.001). Specifically, 36% of patients with a MUST score 2 had a mGPS of 2 compared with 8% of patients with a MUST score of 0 (p<0.001).

208 MUST and clinical outcome

There were significant associations between MUST and length of hospital stay (p<0.001) and 3 year survival (p=0.002). Specifically, 78% of patients with a MUST score 2 had a length of stay >7 days compared with 49% of patients with a MUST score of 0 (p=0.002). Also, 33% of patients with MUST score 2 were dead at 3 years compared with 17% of patients with a MUST score 0 (p=0.001).

214 Length of hospital stay

The variables associated with the length of hospital stay are presented in **Table 2**. On univariate logistic regression, age (p=0.019), ASA (p=0.026), BMI (p=0.045), MUST (p=0.001),visceral obesity (p=0.094), mGPS (p =0.043), NLR(p=0.002), and complication (p<0.001) were significantly associated with length of hospital stay >7 days. Multivariate analysis showed that MUST [OR=2.17 (95% CI 1.45, 3.26) p<0.001] and complication [OR=11.04 (95% CI 5.96,20.44) p<0.001] were independently associated with length of hospital stay >7 days.

222 Complications

There was no significant association between MUST score and either the presence of complications or their severity.

225 Survival

A total of 239 patients were alive at the censor date. Death due to any cause occurred in 82 patients with 51 being cancer specific. The median survival was 38 months (range 1-122 months). After exclusion of thirty-day postoperative mortality (1%), there was a significant association between MUST score and overall survival (p<0.001). Kaplan–Meier curves (**Figure 2**) showed an increased number of deaths for patients at medium or high risk of malnutrition (log rank test, p<0.001).

On univariate cox regression survival analysis (Table 3), age (p<0.001), sex (p=0.048), ASA

233 (p<0.001), TNM stage (p<0.001), BMI (p=0.005), MUST (p<0.001), subcutaneous adiposity

234 (p=0.033), low SMD (p=0.015), mGPS (p=0.005), and NLR (p=0.026) were significantly

associated with overall survival. Multivariate analysis showed that age [HR=1.45 (95% CI

236 1.05,2.01) p=0.023], TNM stage [HR=2.19(95% CI 1.56,3.08) p<0.001], MUST [HR=1.45

237 (95% CI 1.06,1.99) p=0.020], and NLR [HR=1.39(95% CI 1.01,1.90) p=0.037] were

238 independently associated with overall survival.

240

241 **Discussion**

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243 The results of the present study show for the first time that there was a significant association 244 between pre-operative nutritional status (MUST), subcutaneous adiposity, visceral obesity 245 low SMI, systemic inflammatory response and clinical outcomes after surgery of colorectal 246 cancer. Specifically, the majority of patients with a MUST score of 2 were underweight 247 (54%) but only 19% in those with a score of 1. Similarly, there was low adiposity in 58% 248 and 40% respectively, low SMI in 76% and 60% respectively and a mGPS 2 in 36% and 31% 249 respectively. Also, MUST was independently associated with length of hospital stay and 250 overall survival. Therefore, routine MUST assessment usefully characterises important 251 phenotypes associated with malnutrition including loss of skeletal muscle mass and the 252 presence of a systemic inflammatory response in patients with primary operable colorectal 253 cancer.

254 The present results are consistent with previous studies that have shown a relationship 255 between the presence of a systemic inflammatory response and the loss of skeletal muscle 256 (sarcopenia) (17,18,19&20) in primary operable colorectal cancer. Indeed, the presence of a 257 systemic inflammatory response is now recognised as a key hallmark of progressive 258 nutritional decline in patients with cancer (21). It is now clear that nutritional assessment 259 tools such as MUST may also reflect the systemic inflammatory status in patients with 260 cancer. To date, although other studies have reported that nutritional screening tools are 261 associated with poor post-operative outcomes in patients with gastrointestinal cancer (5,6), 262 few studies have clearly delineated the prognostic value of nutritional screening tools 263 independent of potential confounding factors. The results of the present study show that 264 MUST was independently associated with length of hospital stay and long term survival. In 265 contrast, MUST was not associated with the development and severity of post-operative complications. 266

267 It may be that poor nutritional status per se is a relatively weak determinant of post-operative 268 complications in these patients. While this might appear to be counterintuitive, in terms of the present study cohort MUST clearly identifies the underweight (BMI<18.5kg/m²). 269 270 However, post-operative complications have been consistently associated with obesity $(BMI>30 \text{kg/m}^2)$ (22) and it may be that in the present cohort where approximately 10% were 271 272 underweight and approximately 30% were obese that obesity was the main driver of post-273 operative complications. Indeed, visceral obesity has been reported to be independently 274 associated with post-operative complications (23).

In contrast, MUST was independently associated with length of hospital stay and overall survival. The basis of these associations may be more obvious. Patients identified at nutritional risk (ie underweight) are more likely to receive dietetic input and have delayed discharge. Patients with cancer and underweight are likely to be cachectic and this has long been recognised to compromise long term outcomes (21).

280

281 The implications of the present observations are important. MUST uses three criteria to 282 assess the overall risk for malnutrition and it appears that each of the criteria can independently predict clinical outcome (9). Furthermore, it appears to compare well with 283 284 other nutritional screening tools such as NRI, MST and SGA for defining nutritional status 285 and it has been concluded that MUST and SGA are effective in the outpatient setting (10). 286 However, the interpretation of nutritional screening tools such as MUST is of major 287 importance since if such patients are identified to be at nutritional risk there may be an 288 assumption that they are likely to benefit from a nutritional intervention. If, in patients with 289 cancer, MUST reflects, in part, the cachectic process including the systemic inflammatory 290 response then it may be that down regulation of the systemic inflammatory response would be of more benefit (28). There is also the possibility to treat the malnutrition and down 291

292 regulate the systemic inflammatory response. Indeed, recently updated nutritional strategies 293 for cancer patients now suggest considering nutrition with anti-catabolic and inflammation-294 suppressing ingredients such as arginine, omega-3-polyunsaturated fatty acids, and glutamine 295 (21). A growing number of studies have evaluated the use of such immunonutrients in 296 patients with cancer undergoing surgery. For example, a systematic review and meta-297 analysis of 20 studies with a total of 2005 gastrointestinal cancer patients reported that, 298 compared with a standard feed, an immune enhancing feed was associated with lower 299 infective complication rates and length of hospital stay (29&30) More clinical trials in the 300 context of the systemic inflammatory response are required.

301

The main limitation of the present study is that it was a retrospective study of patients in a single institution and only patients with an available MUST score were included in the analysis. However, this study, is to our knowledge, the first to examine the association between a preoperative nutritional assessment tool (MUST), body composition and systemic inflammation in large number of patients undergoing surgery for primary operable cancer.

308 In summary, there was a significant association between pre-operative nutritional status 309 (MUST), body composition, systemic inflammatory response and clinical outcomes after 310 surgery of colorectal cancer. These observations warrant confirmation in other clinical 311 scenarios where nutritional assessment tools are routinely used.

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- 319
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- 321 the data; AA and DM analyzed data and interpretation; AA wrote the paper; DM and CE
- 322 manuscript editing; AA, CE and DM had responsibility for final content. All authors read and
- 323 approved the final manuscript.
- 324
- **Conflict of interest** 325
- 326 All authors confirm no conflict of interest

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Tables:

Table 1. The relationship between MUST and clinical and pathological characteristics, SIR and clinical outcomes¹

| | | Nutrition risk MUST | | | | |
|----------------|-------------------------|---------------------|-----------------------|------------------------|-----------------------|---------|
| Characteristic | | All n(%) | Low risk n=288(79) | Medium risk n=31(9) | High risk n=44(12) | P value |
| Age | <65 | 141(39) | 124 (43) | 6(19) | 11(25) | <0.001 |
| - | 65 - 74 | 136(37) | 108(38) | 12(39) | 16(36) | |
| | >74 | 86(24) | 56(19) | 13(42) | 17(39) | |
| Sex | Male | 199(55) | 163(57) | 13(42) | 23(52) | 0.334 |
| | Female | 164(45) | 125(43) | 18(58) | 21(48) | |
| BMI (kg/m2) | Underweight (<20) | 30(8) | 0(0) | 6(19) | 24(54) | <0.001 |
| | Normal 20-24.9 | 98(27) | 75(26) | 14(45) | 9(20) | |
| | Overweight 25-29.9 | 124(34) | 109(38) | 7(23) | 8(18) | |
| | Obese (<u>></u> 30) | 111(31) | 104(36) | 4(13) | 3(7) | _ |
| ASA grade | 1 | 89(25) | 72(26) | 6(21) | 11(25) | 0.036 |
| | 2 | 155(44) | 128(46) | 14(48) | 13(30) | |
| - | 3 | 96(27) | 74(26) | 7(24) | 15(34) | |
| | 4 | 13(4) | 6(2) | 2(7) | 5(11) | |
| Tumor site | Colon | 225(62) | 173(60) | 23(74) | 29(66) | 0.264 |
| | Rectum | 137(38) | 114(34) | 8(26) | 15(34) | |

| TNM stage | 0 | 11(3) | 9(3) | 0(0) | 2(5) | 0.459 | |
|-------------------|------|---------|---------|--------|--------|--------|--|
| | 1 | 75(21) | 66(24) | 6(19) | 3(7) | _ | |
| | 2 | 140(40) | 103(37) | 16(52) | 21(49) | _ | |
| | 3 | 115(32) | 89(32) | 9(29) | 17(40) | _ | |
| | 4 | 13(5) | 13(5) | 0(0) | 0(0) | | |
| Body composition | | | | | | | |
| Subcutaneous | No | 112(37) | 38(14) | 12(40) | 22(58) | <0.001 | |
| adiposity | Yes | 188(63) | 242(86) | 18(60) | 16(42) | _ | |
| Visceral | No | 100(28) | 58(20) | 16(52) | 26(62) | <0.001 | |
| obesity | Yes | 258(72) | 227(80) | 15(48) | 16(38) | - | |
| Low SMI | No | 175(50) | 154(55) | 12(40) | 9(24) | <0.001 | |
| | Yes | 173(50) | 126(45) | 18(60) | 29(76) | _ | |
| Low SMD | No | 138(39) | 116(41) | 12(39) | 10(24) | 0.047 | |
| | Yes | 220(61) | 169(59) | 19(61) | 32(76) | _ | |
| Inflammatory resp | onse | I | | | | | |
| mGPS | 0 | 266(76) | 226(80) | 17(59) | 23(55) | <0.001 | |
| | 1 | 39(11) | 32(11) | 3(10) | 4(9) | _ | |
| | 2 | 47(13) | 23(8) | 9(31) | 15(36) | _ | |
| NLR | <3 | 184(51) | 151(53) | 13(42) | 20(45) | 0.038 | |
| | 3-5 | 115(32) | 92(32) | 12(39) | 11(25) | | |
| | >5 | 60(17) | 41(14) | 6(19) | 13(29) | | |

| Post-operative of | outcome | | | | | |
|-------------------|------------------|---------|---------|--------|--------|-------|
| Any | No | 227(63) | 182(64) | 18(58) | 27(61) | 0.682 |
| complication | Yes | 133(37) | 103(36) | 13(42) | 17(39) | |
| Non infective | No | 301(84) | 243(85) | 25(81) | 33(75) | 0.077 |
| complication | Yes | 59(16) | 42(15) | 6(19) | 11(25) | |
| Infective | No | 267(74) | 214(75) | 21(68) | 32(73) | 0.566 |
| complication | Yes | 93(26) | 71(25) | 10(32) | 12(27) | |
| Clavien | 0 | 236(66) | 191(67) | 18(58) | 27(61) | |
| Dindo grade | 1-2 | 102(28) | 78(27) | 12(39) | 12(27) | 0.285 |
| | 3-5 | 22(6) | 16(6) | 1(3) | 5(11) | |
| Length of | \leq 7 days | 165(46) | 145(51) | 11(36) | 9(22) | 0.001 |
| hospital stay | >7 days | 192(54) | 141(49) | 20(64) | 31(78) | |
| Survival for 3 | Alive | 288(79) | 239(83) | 24(77) | 25(67) | 0.002 |
| years | All causes death | 75(21) | 49(17) | 7(23) | 19(33) | |

¹N(%), Chi test for linear by linear association. Significant value p<0.001. mGPS: modified Glasgow Prognostic Score. NLR: Neutrophil-to-lymphocyte Ratio. SMD: skeletal muscle radiodensity. SMI: skeletal muscle index

Table 2. The relationship between MUST, clinical and pathological characteristics and **length of hospital stay** (>**7vs**<**7 days**) in patients undergoing surgery for colorectal cancer¹

| LOS | Variable | Univariate OR (95% CI) | P value | Multivariate) OR (95% CI) | P value |
|-----|---------------------------|---------------------------|---------|------------------------------|---------|
| | Age | 1.389(1.05,1.82) | 0.019 | 1.28(0.79,1.71) | 0.176 |
| | Sex | 1.23 (0.81,1.87) | 0.328 | - | - |
| | ASA | 1.35(1.03,1.76) | 0.026 | 1.22(0.87,1.72) | 0.234 |
| | TNM stage | 1.01(0.80,1.28) | 0.898 | - | - |
| | BMI | 0.79 (0.63,0.99) | 0.045 | 0.99(0.67,1.47) | 0.972 |
| | MUST | 1.88(1.31,2.69) | 0.001 | 2.17(1.45,3.26) | <0.001 |
| | Subcutaneous adiposity | 0.69(0.40,1.18) | 0.183 | - | - |
| | Visceral obesity | 0.66(0.41,1.07) | 0.094 | 0.62(0.29,1.32) | 0.217 |
| | Low SMI | 1.37(0.89,2.10) | 0.142 | - | - |
| | Low SMD | 1.09(0.70,1.67) | 0.696 | - | - |
| | mGPS | 1.38 (1.01,1.90) | 0.043 | 0.99(0.67,1.47) | 0.993 |
| | NLR(<3/3-5/>5) | 1.59(1.19,2.12) | 0.002 | 1.26(0.88 ,1.80) | 0.190 |
| | Complication | 9.10(5.26,15.75) | <0.001 | 11.04(5.96,20.44) | <0.001 |

¹ Binary logistic regression, OR: odd ratio, variables associated to a degree of p<0.1 at univariate analysis were entered into a backward conditional multivariate model. P<0.05 considered significant. mGPS: modified Glasgow Prognostic Score. NLR: Neutrophil-to-lymphocyte Ratio. SMD: skeletal muscle radiodensity. SMI: skeletal muscle index Table 3. The relationship between MUST, clinical and pathological characteristics and overall survival in patients undergoing surgery for colorectal cancer¹.

| OS | Variable | Univariate HR (95% CI) | P value | Multivariate HR (95% CI) | P value |
|----|---------------------------|---------------------------|---------|-----------------------------|---------|
| | Age | 1.78 (1.32,2.42) | <0.001 | 1.45(1.05,2.01) | 0.023 |
| | Sex | 1.63(1.00,2.64) | 0.048 | 1.61(0.91,2.85) | 0.109 |
| | ASA | 1.64(1.24,2.17) | <0.001 | 1.27(0.90,1.77) | 0.271 |
| | TNM stage | 1.88(1.40,2.53) | <0.001 | 2.19(1.56,3.08) | <0.001 |
| | BMI | 0.71(0.56,0.90) | 0.005 | 0.91(0.62,1.32) | 0.749 |
| | MUST | 1.69(1.30,2.21) | <0.001 | 1.45(1.06,1.99) | 0.020 |
| | Subcutaneous adiposity | 0.56(0.33,0.96) | 0.033 | 0.87(0.42,1.82) | 0.360 |
| | Visceral obesity | 0.81(0.49,1.33) | 0.411 | - | - |
| | Low SMI | 1.43(0.87,2.34) | 0.155 | - | - |
| | Low SMD | 1.88(1.13,3.14) | 0.015 | 1.08(0.56,2.08) | 0.848 |
| | mGPS | 1.52(1.13,2.06) | 0.005 | 1.25(0.90,1.73) | 0.303 |
| | NLR(<3/3-5/>5) | 1.38(1.03,1.85) | 0.026 | 1.39(1.01,1.90) | 0.037 |

¹Binary logistic regression, OR: odd ratio, variables associated to a degree of p<0.1 at univariate analysis were entered into a backward conditional multivariate model. P<0.05 considered significant. mGPS: modified Glasgow Prognostic Score. NLR: Neutrophil-to-lymphocyte Ratio. SMD: skeletal muscle radiodensity. SMI: skeletal muscle index Legends for figures :

Figure 1: MUST score (Elia., 2003) (9)

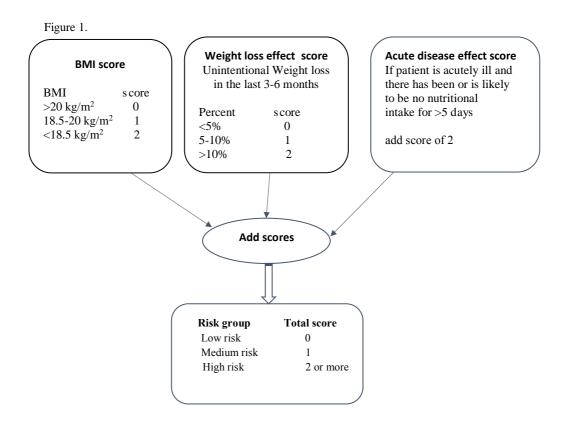


Figure 2: Kaplan–Meier curves showing the relationship between MUST score and overall survival in patients with colorectal cancer (Median follow-up: 38 month. MUST score 0 n = 288, MUST score 1 n = 31, MUST score $\ge 2 n = 44$.)

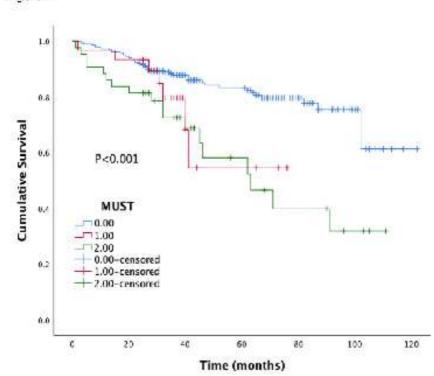


Figure 2