

Modified frailty index predicts complications and death after non-bariatric gastrectomies

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Background: The modified frailty index (mFI) has been shown to predict mortality and morbidity after major operations. The aim of the present study was to assess the mFI as a preoperative predictor of short-term postoperative complications and 30-day mortality in patients undergoing gastrectomy for non-bariatric diseases.

Methods: The American College of Surgeons National Surgical Quality Improvement Program (ACS NSQIP) database was queried for patients who underwent total or partial gastrectomy from 2005 to 2011. A mFI was calculated based on 11 variables as previously described. The population divided into the following four categories based on the mFI score: the non-frail (mFI 0), the low frail (mFI 1), the intermediate frail (mFI 2) and frail (mFI \geq 3). Thirty-day mortality and postoperative complications were evaluated.

Results: Overall, 5,711 patients underwent a gastrectomy for non-bariatric diseases. Higher mFI score was associated with higher rates of mortality (from 1.2% in the non-frail group to 10.7% in frail group, P<0.001), overall morbidity (26.7% vs. 51.1%, P<0.001), postoperative Clavien IV complication (6% vs. 24.6%, P<0.001), serious complications (19.3% vs. 42.6%, P<0.001), sepsis-related complications (8.4% vs. 16.4%, P<0.001), cardiopulmonary complications (5% vs. 20.7%, P<0.001) and failure to rescue (5.7% vs. 21.8%, P<0.001).

Conclusions: Higher mFI score in patients undergoing non-bariatric gastrectomy, is associated with a stepwise greater risk of postoperative morbidity and mortality. MFI Score can be easily calculated preoperatively, from the patient's history, and it can be used as an exceptionally useful criterion for treatment planning.

Keywords: Gastrectomy; outcomes; modified frailty index (mFI); failure to rescue; NSQIP

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Introduction

Total or subtotal gastrectomy is the only curative treatment for certain malignancies of the stomach. It is still considered to be a major surgical procedure with substantial perioperative morbidity and mortality. Patients undergoing surgery for gastric malignancy are usually older with many comorbidities and hence with higher odds of postoperative complications (1). The rate of postoperative serious complications after gastrectomy is around 24%, and the overall mortality has been estimated between 4-5%(2,3). However, frail patients might have higher rates of

Page 2 of 10

Translational Gastroenterology and Hepatology, 2021

postoperative complications compared with the non-frail patients of the same age. The outcomes of gastrectomy in frail patients have not been adequately studied, and outcome predictors for gastrectomy have not been published, other than isolated risk factors (2).

Several authors have reported comorbidities as relative contraindications for gastrectomy without calculating the cumulative effect of several coexisting comorbidities in the postoperative complication rate of any patient. Conventional risk assessment scores such as the American Society of Anesthesiologist (ASA) score or the Postoperative Score to Predict Postoperative Mortality (POSPOM) are of limited clinical use; either because they include a small number of preoperative factors or because they are too complex (4). Obviously, there is an imperative need for studying multiple preoperative clinical risk factors and developing a simple, objective and reliable risk assessment score which would estimate the patient's cumulative health deficit. Recent studies have explored the role of frailty in estimating the postoperative risk after different types of operations. A practical way of frailty measurement is the modified frailty index (mFI), which was first described by Velanovich et al. (5). Although several studies exist on mFI score as a surgical risk calculator, there are no studies on mFI score for patients undergoing gastrectomy for nonbariatric diseases.

In this study, we evaluated the utility of the mFI score as a surgical risk estimator for postoperative complication by using the NSQIP database. We also evaluated the association of each individual frailty components with postoperative serious complications and mortality. We hypothesized that mFI score would be able to identify patients at high risk for short-term (30 days) postoperative complications, and it would be a useful tool for preoperative surgical risk stratification. Our primary hypothesis was that higher frailty levels would be associated with higher rates of post-gastrectomy complications and mortality.

Methods

This is a retrospective cohort analysis of prospectively collected data from the ACS-NSQIP database. The ACS-NSQIP is a national quality improvement program with participation of more than 600 hospitals in 49 of the 50 states. Data is collected on over 150 variables, including preoperative risk factors, intraoperative variables, and postoperative mortality and morbidity (6).

The NSQIP participant use files were queried from 2005-

2011 for patients who underwent partial or total gastrectomy, either open or laparoscopic for either malignant or benign disease of the stomach. The patients were identified based on primary Current Procedural Terminology (CPT) codes: 43620 (Gastrectomy, total; with esophagoenterostomy), 43621 (Gastrectomy, total; with Roux-en-Y reconstruction), 43622 (Gastrectomy, total; with formation of intestinal pouch, any type), 43631(Gastrectomy, partial, distal; with gastroduodenostomy), 43632 (Gastrectomy, partial, distal; with gastrojejunostomy), 43633 (Gastrectomy, partial, distal; with Roux-en-Y reconstruction) and 43634 (Gastrectomy, partial, distal; with formation of intestinal pouch). Our exclusion criteria were cases with no available data on mFI score variables, cases with prior operation within 30 days, cases with open wound or wound infection preoperatively, and cases with concurrent other major operations like liver resections, pancreatectomies, splenectomies, colectomies or small bowel resections.

All outcomes were monitored in the first 30 postoperative days after gastrectomy. MFI score was calculated based on methodology recently described by Velanovich et al. (5). Every patient was allotted one point for each of the following preoperative comorbidity if abnormal: (I) patient functional status (dependent or not dependent); (II) diabetes mellitus (treated with insulin or oral medications); (III) hypertension (requiring treatment); (IV) congestive heart failure (CHF); (V) myocardial infarction (MI); (VI) either prior cardiac surgery and/or percutaneous coronary angioplasty and/or history of angina; (VII) chronic obstructive pulmonary disease (COPD) and/or history of pneumonia; (VIII) rest pain/gangrene secondary to peripheral vascular disease (PVD) and/or PVD with treatment (angioplasty or revascularization or amputation); (IX) impaired sensorium within 48 hours prior to the surgical procedure (not in the context of current neurologic disease like dementia); (X) history of transient ischemic attack (TIA) and/or cerebrovascular accident (CVA) without neurologic deficit and (XI) CVA with neurologic deficit. The total mFI was calculated for every patient. The possible mFI score range from zero to eleven, but in our study population the mFI score range from zero to seven (Table 1). For ease of interpretation and easier future clinical use, the total population was divided into four groups to determine the relationship of frailty to mortality, morbidity and failure to rescue. This was a similar method described by Augustin et al. (7). The four groups of mFI are the following: no frailty group (with mFI score zero), low frailty group (with mFI score 1), intermediate frailty group (with mFI score 2)

Table 1 Mortality,	Clavien IV	complications and	d failure to rescue	distribution	of patients	undergoing	gastrectomy	based or	n modified	frailty i	ndex
score											

Modified frailty index	Total, n (%)	Mortality, n (%)	Clavien IV complications ¹ , n (%)	Failure to rescue ² , n (%)
Overall	5,711	221 (3.9)	635 (11.1)	189 (13.1)
0	2,101 (36.8)	25 (1.2)	126 (6.0)	23 (5.7)
1	1,922 (33.7)	67 (3.5)	191 (9.9)	53 (11.5)
2	1,073 (18.8)	63 (5.9)	167 (15.6)	56 (17.6)
3	412 (7.2)	37 (9.0)	96 (23.3)	30 (17.9)
4	143 (2.5)	19 (13.3)	37 (25.9)	18 (28.6)
5	45 (0.8)	5 (11.1)	11 (24.4)	4 (17.4)
6	11 (0.2)	3 (27.3)	5 (45.5)	3 (50.0)
7	4 (0.1)	2 (50.0)	2 (50.0)	2 (100.0)

¹, *Clavien IV complications* is any one of the following complications in the NSQIP database: cardiac arrest requiring CPR; myocardial infarction; pulmonary embolism; postoperative acute renal failure requiring dialysis; septic shock; failure to wean from ventilator >48 hours and postoperative need for unplanned reintubation. ², *failure to rescue* is defined as death after serious complication.

and frail group (with mFI score \geq 3) (*Table 2*).

Preoperative patient demographic, clinical and laboratory characteristics were analyzed in an attempt to investigate for possible confounding factors. The analyzed preoperative variables were age, gender, race, body mass index (BMI), ASA score \geq 3, preoperative body weight loss >10% in the last six months, preoperative chronic steroid use, postoperative diagnosis (malignancy *vs.* benign disease), preoperative chemotherapy, preoperative radiotherapy, type of gastrectomy, preoperative platelets \leq 50×10³/mm³ and preoperative hematocrit <30% (*Table 2*). BMI and age were analyzed as continuous variables, while all the other variables were analyzed as categorical (*Table 2*).

Postoperative outcomes assessed were: 30-day mortality, overall morbidity, Clavien IV complications (CLIVC), cardiopulmonary complications (included postoperative pulmonary embolism and/or cardiac arrest and/or MI and/ or failure to wean), sepsis-related complications (included postoperative sepsis and/or septic shock), organ space surgical site infection (SSI), serious postoperative complications and failure to rescue after a serious complication (Table 3). CLIVC are life-threatening complications that lead to single organ or multi-organ dysfunction, requiring intensive care unit (ICU) management (8). In our study we defined CLIVC as any one of the following complications in the NSQIP database: cardiac arrest requiring CPR; MI; pulmonary embolism; postoperative acute renal failure requiring dialysis; septic shock; failure to wean from ventilator >48 hours, and postoperative need for unplanned reintubation. Serious complications is defined as (9): pneumonia, unplanned intubation, pulmonary embolism, failure to wean from ventilation, acute renal failure, cerebrovascular accident/ stroke with neurological deficit, MI, cardiac arrest requiring CPR, organ space surgical site infection, deep incisional surgical site infection, wound disruption, sepsis, septic shock, bleeding which needed transfusion and progressive renal insufficiency. Failure to rescue was defined as death after serious complication (9,10). Organ space SSI rates were used as a surrogate for postoperative anastomotic leak in order to evaluate whether the postoperative mortality and complications were associated with the technical failure or with patient's lower physiologic reserves.

Statistical analysis was performed using the program SPSS Statistics Version 24. Data on categorical variables were expressed as frequencies and proportions (%) and were compared between groups of frailty using the Chi-Square test. Data on continuous variables were summarized with descriptive statistics such as means and standard deviations (SDs), and group comparisons of these variables were performed using the ANOVA test. Univariable logistic regression models were used to examine the associations of frailty and other individual predictor variables with the CLIVC and mortality one at a time. Furthermore, multivariable logistic regression was performed to understand the relationship of frailty with the odds of patients having a CLIVC or mortality, after controlling the statistically significant factors in the univariate analysis. Age and BMI transformed to categorical variables in univariate

Page 4 of 10

Translational Gastroenterology and Hepatology, 2021

Table 2 Preoperative demographic characteristics and comorbidities

Variable	Total population, n=5,711 (100%)	No frailty mFI =0, n=2,101 (36.8%)	Low frailty mFI =1, n=1,922 (33.7%)	Intermediate frailty mFI =2, n=1,073 (18.8%)	Frail mFl ≥3, n=615 (10.8%)	P value
Age, mean (± SD) (missing: 75/1.3 %)	62.3 (14.5)	54.6 (13.8)	64.7 (13.4)	68.3 (11.8)	71.3 (10.7)	<0.001
Age ≥65 (%)	2,668 (46.7)	496 (23.7)	1,034 (54.5%)	678 (65.0)	460 (76.5)	<0.001
Gender female, n (%) (missing: 7/0.1 %)	2,929 (51.3)	1,159 (55.3)	1,026 (53.4)	509 (47.5)	235 (38.2)	<0.001
Race Caucasian, n (%) (missing: 516/9%)	3,814 (66.8)	1,395 (66.4)	1,278 (66.5)	728 (67.8)	413 (67.2)	0.85
BMI, mean (± SD) (missing: 121/2.1%)	27.54 (7.98)	26.3 (7.8)	27.83 (7.96)	29 (8.37)	28.3 (7.27)	<0.001
ASA ≥ 3, n (%) (missing: 8/0.1%)	3,848 (67.5)	1,013 (48.3)	1,333 (69.5)	912 (85.2)	590 (95.9)	<0.001
Weight loss, n (%) (missing: 0/0%)	794 (13.9)	331 (15.8)	281 (14.6)	116 (10.8)	66 (10.7)	<0.001
Steroid use, n (%) (missing: 0/0%)	163 (2.9)	35 (1.7)	46 (2.4)	42 (3.9)	40 (6.5)	<0.001
Patients with malignancy, n (%) (missing: 0/0%)	3,317 (58.1)	1,161 (55.3)	1,151 (59.9)	645 (60.1)	360 (58.5)	0.01
Chemotherapy, n (%) (missing: 0/0%)	237 (4.1)	103 (4.9)	77 (4.0)	39 (3.6)	18 (2.9)	0.106
Radiotherapy, n (%) (missing: 8/0.1%)	104 (1.8)	48 (2.3)	27 (1.4)	20 (1.9)	9 (1.5)	0.185
Type of operation						<0.001
Distal/partial gastrectomy	4,175 (73.1)	1,457 (69.3)	1,411 (73.4)	821 (76.5)	486 (79)	
Total gastrectomy	1,326 (23.2)	557 (26.5)	444 (23.1)	211 (19.7)	114 (18.5)	
Gastrectomy with pouch formation	210 (3.7)	87 (4.1)	67 (3.5)	41 (3.8)	15 (2.4)	
Platelets ≤50, n (%) (missing: 205/3.6%)	30 (0.5)	14 (0.7)	9 (0.5)	3 (0.3)	4 (0.7)	0.5
Hematocrit <30, n (%) (missing: 199/3.5 %)	850 (15.4)	254 (12.6)	293 (15.8)	162 (15.6)	141 (23.5)	<0.001

BMI, body mass index; ASA score, American Society of Anesthesiologists' (ASA) classification of Physical Health Score. *Weight loss*: patients with a greater than 10% decrease in body weight in the six-month interval immediately preceding surgery as manifested by serial weights in the chart, as reported by the patient, or as evidenced by change in clothing size or severe cachexia. Patients who have intentionally lost weight as part of a weight reduction program do not qualify. *Steroid use*: patient who required regular administration of oral or parenteral corticosteroid medications (e.g., Prednisone, Decadron) in the 30 days prior to surgery for a chronic medical condition (e.g., COPD, asthma, rheumatologic disease, rheumatoid arthritis, inflammatory bowel disease). Topical corticosteroids applied to the skin or corticosteroids administered by inhalation or rectally are not included. Patients who only receive short course steroids (duration 10 days or less) in the 30 days prior to surgery are not included. *Chemotherapy*: if the patient had any chemotherapy treatment for cancer in the 30 days prior to surgery. Chemotherapy may include, but is not restricted to, oral and parenteral treatment with chemotherapeutic agents for malignancies such as colon, breast, lung, head and neck, and gastrointestinal solid tumors as well as lymphatic and hematopoietic malignancies such as lymphomas, leukemia, and multiple myeloma. Patient is not included if treatment consists solely of hormonal therapy. *Radiotherapy*: if the patient had any radiotherapy treatment for cancer in the 90 days prior to surgery. Count If the patient had radiation seeds implanted and the implantation was within 90 days prior to the operation.

and multivariable analysis for easier future clinical use. Patients were categorized in those with age 65 or older and patients with age <65. Respectively patients were grouped in those with BMI 30 or higher and patients with BMI <30. Wald tests were used to assess the significant contribution of each predictor variables, and unadjusted raw or adjusted odds ratios and their 95% confidence intervals (CIs) were reported as appropriate for both CLIVC and mortality. Two-tailed P values of less than 0.05 were deemed statistically significant. Finally, the impact of every individual frailty component was studied, and the odds of CLIVC and mortality calculated for each particular mFI

 Table 3 30 days mortality, overall morbidity, Clavien IV complications, cardiopulmonary complications, sepsis related complications, serious complications, failure to rescue and organ space surgical site infections (SSI)

Variable, n (%)	Total cohort	No frailty	Low frailty	Intermediate frailty	Frail	P value
N (%)	5,711 (100.0)	2,101 (36.8)	1,922 (33.7)	1,073 (18.8)	615 (10.8)	
Mortality	221 (3.9)	25 (1.2)	67 (3.5)	63 (5.9)	66 (10.7)	<0.001
Overall morbidity	1867 (32.7)	562 (26.7)	594 (30.9)	397 (37.0)	314 (51.1)	<0.001
Clavien IV complications	635 (11.1)	126 (6.0)	191 (9.9)	167 (15.6)	151 (24.6)	<0.001
Cardiopulmonary complications	518 (9.1)	106 (5.0)	149 (7.8)	136 (12.7)	127 (20.7)	<0.001
Sepsis related complications	598 (10.5)	176 (8.4)	191 (9.9)	130 (12.1)	101 (16.4)	<0.001
Serious complications	1,446 (25.3)	405 (19.3)	461 (24.0)	318 (29.6)	262 (42.6)	<0.001
Failure to rescue	189 (13.1)	23 (5.7)	53 (11.5)	56 (17.6)	57 (21.8)	<0.001
Organ space SSI	328 (5.7)	117 (5.6)	98 (5.1)	67 (6.2)	46 (7.5)	0.138
Patients with malignancy						
N (%)	3,317 (100.0)	1,161 (35.0)	1,151 (34.7)	645 (19.4%)	360 (10.9)	
Mortality	102 (3.1)	14 (1.2)	34 (3.0)	32 (5.0)	22 (6.1)	<0.001
Overall morbidity	1,019 (30.7)	313 (27.0)	337 (29.3)	209 (32.4)	160 (44.4)	<0.001
Clavien IV complications	300 (9.0)	69 (5.9)	92 (8.0)	76 (11.8)	63 (17.5)	<0.001
Cardiopulmonary complications	227 (6.8)	56 (4.8)	65 (5.6)	60 (9.3)	46 (12.8)	<0.001
Failure to rescue	96 (14.4)	9 (5.0)	23 (11.5)	26 (16.6)	38 (28.8)	<0.001
Organ space SSI	190 (5.7)	69 (5.9)	56 (4.9)	39 (6.0)	26 (7.2)	0.348
Patients with benign disease						
N (%)	2,394 (100.0)	940 (39.3)	771 (32.2)	428 (17.9)	255 (10.7)	
Mortality	119 (5.0)	11 (1.2)	33 (4.3)	31 (7.2)	44 (17.3)	<0.001
Overall morbidity	848 (35.4)	249 (26.5)	257 (33.3)	188 (43.9)	154 (60.4)	<0.001
Clavien IV complications	335 (14)	57 (6.1)	99 (12.8)	91 (21.3)	88 (34.5)	<0.001
Cardiopulmonary complications	291 (12.2)	50 (5.3)	84 (10.9)	76 (17.8)	81 (31.8)	<0.001
Failure to rescue	116 (4.6)	16 (1.6)	40 (4.8)	35 (8.1)	25 (9.9)	<0.001
Organ space SSI	138 (5.8)	48 (5.1)	42 (5.4)	28 (6.5)	20 (7.8)	0.334

component.

Results

Baseline characteristics of the population

The final population after applying the exclusion criteria was 5,711 patients (*Figure 1*). There were 2,101 (36.8%) patients with no frailty, 1,922 (33.7%) patient with low

frailty (mFI score 1), 1,073 (18.8%) with intermediate frailty (mFI score 2) and 615 (10.8%) frail patients (mFI score \geq 3) (*Table 2*). The mean age of the population was 62.3 (±14.5) and higher mFI groups had higher means of age compared to the non-frail group. The overall BMI was 27.5 (±7.98) with significantly higher BMI patients in the intermediate and frail groups. Of the total population, 2,929 (51.3%) patients were female, and 3,814 (66.8) patients were Caucasian. Higher frailty groups include increased rates of



Figure 1 Flow diagram of patients selected for analysis. mFI, modified frailty index.

male patients, ASA scores ≥ 3 and malignancy; and decreased percentages of body weight loss (>10% in the last six months before the operation), preoperative chemotherapy, and preoperative radiotherapy. Further information of the patient's demographic and preoperative tests are summarized in *Table 2*. For the multivariable analysis we grouped the procedures in distal/partial gastrectomies, total gastrectomies and gastrectomy with pouch formation.

Postoperative complications and estimates of covariate associations with mortality and serious morbidity

Clavien IV complication occurred in 635 (11.1%) patients during the first 30 postoperative days after the operation, and the 30-day mortality was 3.9%. *Table 3* and *Figure 2* provide a breakdown of morbidity and mortality according

Translational Gastroenterology and Hepatology, 2021

to the frail groups. A significant increase in overall complications, Clavien IV complications, cardiopulmonary complications, sepsis related complications, failure to rescue from serious complications, and mortality were seen with an increased mFI. However, no significant difference was found in the rate of organ space SSI between the mFI groups which indicates no surgical causes of postoperative complications and mortality. The prognostic value of the mFI score was confirmed also in the population with malignancy after performing a subgroup analysis (Table 3). Univariate regression analysis identified frailty as the strongest preoperative factor for having postoperative CLIVC (frail group: OR =5.1, CI =3.94-6.59) and mortality (frail group: OR =9.98, CI =6.24-15.97) (Table 4). On univariate analysis, CLIVC associated with age ≥ 65 , male gender, preoperative weight loss >10%, preoperative steroid use, hematocrit <30 and type of gastrectomy (total gastrectomy). Moreover, mortality was associated with age \geq 65, BMI \geq 30, preoperative weight loss >10%, preoperative chemotherapy, preoperative steroid use and hematocrit <30.

In multivariable regression analysis, after adjusting for confounders, low frail patients had more than 50% higher odds of having CLIVC and almost 100% higher odds of 30-days mortality in comparison to non-frail. Intermediate frail patients had 140% higher odds of CLIVC and 200% higher odds of 30-day mortality. Frail patients had more than 280% higher odds of CLIVC and 410% higher odds of 30-day mortality. In multivariable logistic regression model, the preoperative factors which were consistently statistically significant in CLIVC are the frailty status, age >65, male gender, weight loss >10%, steroid use, hematocrit <30 and total gastrectomy. The corresponding factors for mortality are frailty, age >65, weight loss >10%, preoperative chemotherapy, preoperative steroid use and hematocrit <30.

Because mFI is not weighted and there was a possibility that every component of frailty calculation has different impact on postoperative morbidity and mortality, we analyzed the individual contribution of each frailty component toward the risk of CLIVC and mortality (*Table 5*). It appears that functional status, hypertension, CHF, MI, CS or angina or PCA history, COPD or pneumonia, PVD or Rest Pain and Impaired Sensorium are the mFI factors that steadily influence the final mFI score predictive value on both mortality and CLIVC. Diabetes Mellitus and TIA or CVA without neurologic deficit seems to affect the mFI score only on the CLIVC prediction. On the other hand CVA with neurologic deficits seems to affect the mFI score only



Figure 2 Observed 30 days mortality, overall morbidity, Clavien IV complications, surgical site infections and sepsis related complications by frailty group.

		Univariat	e analysis		Multivariate analysis				
Risk factors	Clavien IV		Mortality	ty Clavien		V	Mortality	Mortality	
	uOR (95% CI)	P value	uOR (95% Cl)	P value	aOR (95% CI)	P value	aOR (95% CI)	P value	
Frailty groups									
No frailty	ref		ref		ref		ref		
Low frailty	1.73 (1.37–2.19)	<0.001	2.99 (1.89–4.77)	<0.001	1.51 (1.18–1.94)	0.001	1.94 (1.18–3.19)	0.009	
Intermediate frailty	2.89 (2.26–3.69)	<0.001	5.18 (3.24–8.28)	<0.001	2.4 (1.84–3.13)	<0.001	3.01 (1.79–5.05)	<0.001	
Frail	5.1 (3.94–6.59)	<0.001	9.98 (6.24–15.97)	<0.001	3.88 (2.91–5.16)	<0.001	5.12 (3.03–8.65)	<0.001	
Age ≥65	2.05 (1.72–2.43)	<0.001	4.615 (3.26–6.53)	<0.001	1.43 (1.19–1.75)	<0.001	3.08 (2.09–4.53)	<0.001	
Gender, male	1.4 (1.18–1.65)	<0.001	1.24 (0.95–1.63)	0.116	1.2 (1.01–1.44)	0.042	-	-	
BMI ≥30	1.07 (0.88–1.29)	0.49	0.69 (0.49–0.97)	0.031	_		0.87 (0.61–1.25)	0.451	
Weight loss >10%	1.26 (1.01–1.58)	0.04	1.76 (1.27–2.45)	0.001	1.32 (1.04–1.68)	0.022	2.03 (1.41–2.93)	<0.001	
Chemotherapy	0.77 (0.49–1.21)	0.26	1.99 (1.19–3.33)	0.008	_		2.38 (1.38–4.1)	0.002	
Steroid use	2.8 (1.95–4.03)	<0.001	2.84 (1.66–4.84)	<0.001	2.52 (1.72–3.71)	<0.001	2.30 (1.28–4.13)	0.005	
HCT <30	2.11 (1.74–2.57)	<0.001	2.29 (1.69–3.1)	<0.001	1.93 (1.57–2.37)	<0.001	1.92 (1.37–2.68)	<0.001	
Type of gastrectomy									
Distal gastrectomy	ref		ref		ref		-	-	
Total gastrectomy	1.26 (1.05–1.53)	0.015	0.694 (0.69–1.32)	0.796	1.42 (1.16–1.73)	<0.001	-	-	
Gastrectomy with pouch formation	1.21 (0.79–184)	0.39	0.84 (0.39–1.82)	0.664	1.37 (0.87–2.15)	0.171	-	-	

Table 4 Univariate and multivariate regression analysis of risk factors for postoperative Clavien IV complications and 30 days mortality

aOR, adjusted odds ratios; uOR, unadjusted odds ratios.

Page 7 of 10

Page 8 of 10

Translational Gastroenterology and Hepatology, 2021

Table 5 Unadjusted risk of Clavien I	V complications and mortality	y for each individual frailt	v components
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· · · · · ·	Mortality		Clavien IV complications		
Frailty components –	OR (95% CI)	P value	OR (95% Cl)	P value	
FS	7.19 (5.36–9.66)	<0.001	4.51 (3.63–5.59)	<0.001	
Diabetes mellitus	1.25 (0.9–1.74)	0.178	1.27 (1.03–1.55)	0.024	
Hypertension	2.49 (1.85–3.37)	<0.001	1.64 (1.38–1.94)	<0.001	
CHF	5.26 (2.63–10.53)	<0.001	3.88 (2.24–6.75)	<0.001	
MI	4.69 (2.07–10.63)	<0.001	2.944 (1.51–5.73)	0.001	
CS or Angina or PCA history	2.39 (1.73–3.32)	<0.001	2.09 (1.68–2.59)	<0.001	
COPD or pneumonia	3.05 (2.07–4.49)	<0.001	2.86 (2.19–3.73)	<0.001	
PVD or rest pain	2.53 (1.15–5.57)	0.021	2.13 (1.22–3.71)	0.008	
Impaired sensorium	8.77 (4.72–16.32)	<0.001	23.29 (12.8–42.3)	<0.001	
TIA or CVA without neurologic deficit	1.56 (0.89–2.73)	0.117	2.0 (1.44–2.78)	<0.001	
CVA with neurologic deficits	2.19 (1.7–4.12)	0.015	1.27 (0.78–2.08)	0.334	

FS: functional status-dependent vs. non-dependent: this variable focuses on the patient's abilities to perform activities of daily living (ADLs) in the 30 days prior to surgery. Activities of daily living are defined as 'the activities usually performed in the course of a normal day in a person's life'. ADLs include: bathing, feeding, dressing, toileting, and mobility. CHF, congestive heart failure; MI, myocardial infarction history; CS, history of cardiac surgery; PCA, percutaneous coronary angioplasty; PVD, peripheral vein disease; TIA, transient ischemic attack; CVA, cerebrovascular accident.

on the mortality prediction and not on CLIVC prediction.

Finally, the mFI score demonstrates good discrimination for both 30-day mortality (AUROC =0.703, 95% CI: 0.669– 0.738) and Clavien IV complications (AUROC =0.65, 95% CI: 0.627–0.673). Furthermore, the discriminative ability of the model with adjustment of covariates it seems to be even better for both 30-day mortality (AUROC =0.773, 95% CI: 0.742–0.805) and Clavien IV complications (AUROC =0.692, 95% CI: 0.67–0.714).

Discussion

Gastrectomy remains as the main curative treatment option for any patient with either malignant or benign gastric tumor. However, gastrectomy is associated with significant morbidity and mortality (2,3). These can be attributed to: the complexity of the particular operation and also to the fact that patient undergoing gastrectomy are usually older with many comorbidities (1,11). Models used in the past lacking predictive value or are too laborious to use (11). Moreover, the majority of recent studies for preoperative risk estimation in gastrectomy have focused on either single risk factors or single postoperative outcomes, both of which have limited clinical use (2,12,13). We used a simplified method of calculation mFI tailored to the ACS-NSQIP (5).

Our results provide compelling evidence that mFI is a powerful predicting model for 30-day mortality, CLVIC and postoperative failure to rescue after a serious complication. The present study revealed that frail patients undergoing gastrectomy have almost 290% higher odds of having a CLIVC complication and more that 400% higher odds of having postoperative mortality. Also, we found that frail patients undergoing gastrectomy have significantly higher rates of any complication, cardiopulmonary complications, sepsis-related complications and failure to rescue from a serious complication. However, frail patients undergoing gastrectomy were not found to have significantly higher rates of postoperative organ space SSI which would suggest anastomotic leaks. This supports the idea of reduced physiologic reserves, and not technical errors as the root cause. Our results are analogous to existing studies of mFI for other operations like hepatectomies, pancreatectomies, necrosectomies and gynecologic operations. The mFI score is a highly useful tool for daily clinical use (14-17).

This study has several limitations, both in data extraction and database design. First, the preoperative variables are predefined for specific comorbidities and laboratories, and much critical information is lacking. For example, the

information regarded the total glycated hemoglobin for diabetic patients is not available. Therefore, we cannot differentiate patients with poorly controlled diabetes mellitus from those with optimal control. Also, the postoperative outcomes are limited to the postoperative 30day period. Furthermore, NSQIP database has significant limitations on technical-related complications report, and these complications can incorrectly be attributed to preoperative patient's frail status (18). Finally, our study is subjected to all potential limitations of a retrospective study.

Future work can be focused on incorporating more factors in the model and adding details such as genetic information, laboratory values as well as social such as smoking, alcohol or illicit drug use of the patient. Furthermore, the mFI score can be applied to new mobile or computer applications, and play a fundamental role in the decision-making process, something already proposed by Koble *et al.* (17).

In conclusion, mFI incorporation in preoperative assessment of patient undergoing gastrectomy adds valuable information regarding the postoperative morbidity, mortality and failure to rescue after a serious complication.

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Footnote

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Conflicts of Interest: All authors have completed the ICMJE uniform disclosure form (available at http://dx.doi. org/10.21037/tgh.2020.01.07). The authors have no conflicts of interest to declare.

Ethical Statement: The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are

appropriately investigated and resolved. The NSQIP data base is a de-identified patient database and consequently our study was exempt from institutional review board (IRB) approval.

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Page 10 of 10

Translational Gastroenterology and Hepatology, 2021

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