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# Frailty in perioperative patients in three South African academic hospitals

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**Background**. Frailty is a state characterised by diminished physiological reserve that leaves an individual vulnerable to external stressors and delays recovery. Frailty assessments are proving to be more valuable in predicting poor perioperative outcomes than other well-known perioperative risk assessment tools. Very few studies using validated frailty assessment tools have been done to assess the prevalence of frailty in South Africa (SA), and none have assessed the intraoperative implications of frailty in a surgical population.

**Objectives.** To determine the demographics and frailty levels of patients presenting for surgery at three academic hospitals in Johannesburg, compare intraoperative complications between the frail and non-frail patients, and compare the association between frailty scores and American Society of Anesthesiologists Physical Status (ASA-PS) scores.

**Methods.** We prospectively enrolled 299 patients aged 18 - 90 years undergoing various types of elective surgery between mid-November 2016 and mid-March 2017 in three SA academic hospitals. Frailty was assessed using the nine-point Clinical Frailty Scale (CFS) and defined as a score of  $\geq$ 5. The CFS and demographic and clinical data were documented by the anaesthetists assigned to the respective elective lists. The primary outcome measure was intraoperative complications (hypotension, desaturation, and need for vasopressors and blood transfusion). We also compared associations between the patients' comorbidities and frailty and those between the CFS and ASA-PS scores.

**Results.** Of a total of 299 patients included in the study (mean age (standard deviation) 50.6 (15.8) years), 156 (52%) were women and 67 (22%) were classified as frail. Compared with patients who were not classified as frail, the frail group had significantly higher incidences of hypotension (odds ratio (OR) 1.87, 95% confidence interval (CI) 1.083 - 3.259; p=0.02) and desaturation (OR 3.79, 95% CI 1.367 - 10.54; p=0.01), and were more likely to need vasopressors (OR 2.81, 95% CI 1.607 - 4.912; p=0.00) and blood transfusion (OR 3.26, 95% CI 1.138 - 9.368; p=0.02). On multivariable logistic regression analysis, adjusting for factors related to frailty such as age, gender and comorbidities, desaturation was significantly associated with frailty (adjusted OR (aOR) 4.21, 95% CI 1.31 - 13.53; p=0.01), and the frail were more likely to require blood transfusion (aOR 5.36, 95% CI 1.50 - 19.16; p=0.01) and were older and had more comorbidities. Higher ASA-PS scores were also strongly associated with frailty.

**Conclusions.** The prevalence of frailty was high among surgical patients. Consistent with other studies, frailty was associated with older age and multiple comorbidities. The association between frailty and intraoperative complications found in this study may indicate and help inform areas of further research.

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The ageing population is growing at a rapid rate worldwide, resulting in an increased number of ageing individuals requiring medical services and presenting for surgical procedures.<sup>[11]</sup> In 2010, twothirds of the world's population aged >60 years lived in developing countries,<sup>[2]</sup> and the World Health Organization (WHO) reports that the number of people aged >60 will effectively double by 2050.<sup>[3]</sup> In the USA, >40% of surgical procedures are performed on elderly patients.<sup>[4]</sup> The increasing number of operations being performed on elderly patients has made it necessary to identify and manage geriatric conditions that place these individuals at increased surgical and anaesthetic risk.<sup>[4,5]</sup>

Frailty is the depletion of physiological and functional reserve across many organ systems, leading to a diminished capacity to withstand environmental stressors such as surgery.<sup>[6]</sup> An independent predictor of mortality, morbidity, complications and increased length of stay perioperatively,<sup>[7]</sup> it is likely to become one of the most problematic public health issues and will place a heavy strain on already overburdened healthcare systems.<sup>[8]</sup> Although often associated with advanced chronological age, frailty can also affect the non-elderly. Rockwood *et al.*<sup>[9]</sup> found the prevalence of frailty to be 2.0% in individuals aged >30 years,<sup>[9]</sup> which is similar to the findings of Kehler *et al.*<sup>[10]</sup> that the prevalence of frailty in Canadians between the ages of 18 and 34 was 1.8 - 5.3%, depending on the frailty assessment used.

A number of assessment methods have been used as indicators of overall health and physiological status. The American Society of Anaesthesiologists Physical Status (ASA-PS) scoring system is a scale from 1 through 6 in increasing order of risk, with an 'E' to denote emergency surgery, that is assigned to a patient by the anaesthesiologist before surgery.<sup>[11]</sup> Originally designed to describe the overall health status of a patient, it has since been used as a risk stratification tool and found to be predictive of postoperative morbidity and mortality.<sup>[12]</sup> The Revised Cardiac Risk Index is a simplified version of an earlier index.<sup>[13]</sup> It has six predictive factors with one point assigned per factor and with an increasing number of points corresponding to increased rates of perioperative cardiac risk.<sup>[14]</sup> Frailty assessments have been shown to be better predictors of postoperative complications, mortality and morbidity in the aged than other preoperative assessments.<sup>[15-17]</sup>

There are many assessment models that have been used to assess frailty. The two most commonly used models are the frailty phenotype<sup>[18]</sup> and the frailty index or deficit accumulation.<sup>[19]</sup> Both models have mainly been used for research purposes, and there have been concerns about their usefulness in a clinical setting. Screening tools such as the Clinical Frailty Scale (CFS) from Dalhousie University in Canada were subsequently developed for practical clinical use.<sup>[20]</sup> The CFS is a 9-point scale that uses pictographs to depict degrees of functional dependency. The scale ranges from 1 (very fit) to 9 (terminally ill), and a score of  $\geq$ 5 suggests frailty. Most studies using any one of the validated frailty measuring tools have been conducted in developed countries on predominantly white patients, with few published studies in the developing world and fewer still in Africa.<sup>[21]</sup> One study utilising data from the WHO study on global ageing and adult health found the prevalence of frailty to be 38% in South Africa (SA) and 37.9% in Ghana in adults aged  $\geq$ 50 years.<sup>[22]</sup> Another SA study compared frailty between HIV-positive and HIV-negative adults.<sup>[23]</sup> The mean ages of the HIV-positive and HIV-negative groups were 41.1 years and 42 years, respectively. There was a 19.4% prevalence of frailty in the HIV-positive adults compared with 13.3% in those who were HIV-negative.

Combining risk assessment tools such as the ASA-PS and frailty assessments to improve postoperative risk prediction could increase the probability of correct identification of patients who are likely to suffer adverse intraoperative and postoperative adverse events, and has been suggested by some authors.<sup>[5,24]</sup> However, combinations could make these tools more complex to use and limit their applicability to a wider population, and are likely to require extensive validation studies before they can be put into practice. Although much has been written about the postoperative adverse outcomes and mortality related to frailty, little is known about how frailty impacts on anaesthetic complications intraoperatively, or on the nature of utilisation of health services such as the intensive care unit (ICU) by patients assessed as frail. Correct risk assessment will identify the most vulnerable and may help facilitate discussions about realistic treatment goals.

# Objectives

To determine the prevalence of frailty and assess the intraoperative implications of frailty in a surgical population in three academic hospitals in Johannesburg, SA. We also assessed the association between frailty and comorbid illnesses and the association between the ASA-PS score and the CFS. Finally, we determined whether participants assessed as frail were more likely to go to a high-care unit or an ICU facility postoperatively.

# Methods

Approval for the study was obtained from the Human Research Ethics Committee (Medical) of the University of the Witwatersrand (ref. no. M160756). Participants aged 18 - 90 years presenting for elective surgery at Chris Hani Baragwanath Academic Hospital (CHBAH), Charlotte Maxeke Johannesburg Academic Hospital (CMJAH) and Helen Joseph Hospital (HJH) between mid-November 2016 and mid-March 2017 were included. The study period was effectively 4 months. Combined, these hospitals operate on an average of 2 500 elective cases monthly. This was a prospective, observational and cross-sectional study that employed convenience sampling. The surgical specialties included were general, gynaecological, orthopaedic, maxillofacial, ear, nose and throat, urological, neurosurgical and vascular. We excluded patients known to be pregnant and those with a diagnosis of mental impairment or dementia without any access to family members for information on frailty.

Frailty was defined using the CFS,<sup>[20]</sup> which is a validated 9-point assessment from the Canadian Study on Health and Aging (Table 1). Patients were defined as frail if they scored  $\geq$ 5 on the CFS. Copies of the data collection sheet and the CFS were placed in the operating theatre receiving areas at the three sites. Patients presenting for any of the included elective surgeries were identified during the preoperative assessments by the anaesthetists assigned to various elective lists. The anaesthetists included medical officers and registrars at different levels of training. After obtaining written informed consent from the patient or family member, demographic data and clinical data, which included comorbidities and planned procedure, were documented. An ASA-PS score and CFS was assigned on the data collecting sheet. Intraoperative complications and postoperative destination were documented.

All data were entered into a password-protected Microsoft Excel 2016 spreadsheet (Microsoft, USA). Patient-identifiable data were removed. The data were analysed using Stata version 13 (StataCorp, USA). Descriptive statistics were tabulated using means and percentages. The  $\chi^2$  test and Fisher's exact test were used to describe the association between categorical variables. Logistic regression analysis was used to assess factors associated with frailty. Univariate logistic regression analysis was performed for each of the variables. Multivariable logistic regression analysis was performed to adjust for the possible confounding variables. A *p*-value <0.05 was considered statistically significant for all analyses.

# Results

A total of 312 data collection forms were filled in and returned by the attending anaesthetists. We collected data on these 312 patients. Thirteen were excluded in the final analysis (7 patients did not undergo the planned surgery, and 6 forms were not filled in and/or consent was not documented), leaving a total of 299 patients. There were 156 women (52%) and 143 men (48%). The mean (standard deviation) age was 50.6 (15.8) years (range 21 - 90).

The CFSs assigned ranged from 1 (fit) to 8 (very severely frail). Of the patients, 67 (22%) were classified as frail, with 39 (58%) of these being female and 28 (42%) male. There were more frail women than men, as shown in Table 2. This was not statistically significant (p=0.263).

At CHBAH 27 out of 106 patients (25.4%) were frail, at CMJAH 28 out of 112 (25%) were frail, and at HJH 12 out of 81 (15%) were frail. Of all the patients, 222 (74%) had an ASA-PS score of 1 or 2 whereas 75 (26%) had an ASA-PS score of 3 or 4. Higher ASA-PS scores were strongly associated with frailty relative to an ASA-PS score of 1, with an ASA-PS score of 3 being the most highly associated with frailty (odds ratio (OR) 19.01, 95% confidence interval (CI) 6.870 - 52.614; p<0.001).

Between the ages of 18 and 34 years, 4% of patients were assessed as frail, in the 35 - 49 age group 13%, in the 50 - 64 age group 37%, and in the over-65 age group 45%. Age was significantly associated with frailty (OR 1.06, 95% CI 1.034 - 1.084; p<0.001. Frailty was also associated with an increased number of comorbidities. Older patients and those with more comorbidities were also more likely to experience intraoperative complications such as hypotension and need for vasopressors (Table 3).

Table 1. Clinical Frailty Scale <sup>[20]</sup>				
1. Very fit	Robust, active, energetic and motivated. Commonly exercise. Fittest for their age			
2. Well	No active disease symptoms, but less fit than category 1			
3. Managing well	Medical problems are well controlled, but not regularly active			
4. Vulnerable	Not dependent on others for daily help. Symptoms limit activities			
5. Mildly frail	More evident slowing. Need help in higher-order instrumental activities of daily living			
6. Moderately frail	Need help with all outside activities. Need help with instrumental and non-instrumental activities of daily living			
7. Severely frail	Completely dependent for personal care. Not at high risk of dying			
8. Very severely frail	Completely dependent. Approaching the end of life			
9. Terminally ill	Approaching the end of life, or life expectancy <6 months in those who are not otherwise evidently frail			

	n (%)	Age (years), mean (SD)	Gender, <i>n</i> (%)
Overall	299	50.6 (15.8)	Women 156 (52.2)
			Men 143 (47.8)
Frail (CFS ≥5)	67 (22.4)	62.29 (15.45)	Women 39/67 (58.2)
			Men 28/67 (41.8)
Non-frail	232 (77.6)	47.30 (14.32)	Women 117/232 (50.4)
			Men 115/232 (49.6)

Before adjusting for confounding variables, frailty was significantly associated with an increased incidence of hypotension and desaturation, need for blood transfusion and need for vasopressors. We then adjusted for confounding variables (age, gender, comorbidities) that are related to frailty and intraoperative complications. After adjustment, desaturation was significantly associated with frailty (adjusted odds ratio (aOR) 4.12, 95% CI 1.202 - 14.139; p=0.024) as well as need for blood transfusion (aOR 5.36, 95% CI 1.50 - 19.16; p=0.01), as shown in Table 3. Age, gender and comorbidities were not significantly related to frailty in either univariate or multivariable logistic regression analysis when assessing desaturation and need for blood transfusion.

Patients assessed as frail also had an increased likelihood of going to high care postoperatively (OR 3.01, 95% CI 1.073 - 8.447; p=0.036).

# Discussion

The prevalence of frailty is said to be higher in developing countries than in developed countries, where it ranges from 5.4% to 44% in community-dwelling elderly<sup>[2]</sup> and is reported to be between 4.1% and 50.3% in patients of all ages presenting for surgical procedures, depending on the assessment tool used and the population sampled.<sup>[25]</sup> The present study found that 22% of patients undergoing elective surgery in three large academic hospitals in Johannesburg were frail. This percentage is lower than our estimate of 30% and is also lower than the reported rates of frailty among the hospitalised elderly.<sup>[4,15]</sup> Our study, however, was one of the few that included age groups  $\geq$ 18 years, and on further analysis it was found that 45% of patients aged >65 were frail, which is on the higher end of prevalence rates generally reported for surgical patients. The distribution of frailty according to type of surgery is shown in Table 4.

Other studies have used the CFS to predict mortality and adverse outcomes. Bagshaw *et al.*<sup>[26]</sup> used the CFS in critically ill adults aged  $\geq$ 50 years and showed a prevalence of frailty of 32.8%. They also showed that in-hospital mortality was higher in the frail (32% v. 16% in the non-frail) and that major adverse events were more common (39% v. 29%). Similar to our study, the frail were older and had more comorbidities, which is also consistent with many other studies.<sup>[18,27,28]</sup>

Hewitt *et al.*<sup>[29]</sup> showed that 28% of patients aged  $\geq$ 65 in three acute surgical admission units were frail according to the CFS. The frail individuals spent longer in hospital and were more likely to die within 30 and 90 days after surgery.

The present study also demonstrates the ability of the CFS to predict outcomes even when used by clinicians without prior training on how to use the scale. In a comparable study, the CFS was used by junior doctors without any prior training in patients aged >65 years admitted to an acute general medical unit.<sup>[30]</sup> In this study, frailty was associated with functional decline (p=0.011) and mortality within 3 months (p=0.012).

There are no current studies with which we can directly compare our study, as most studies have focused on postoperative outcomes.<sup>[31]</sup> We could not find any studies that aimed to associate frailty with intraoperative complications, although a number of studies have shown a relationship between intraoperative complications and postoperative outcomes. The need for intraoperative blood transfusion was shown to be associated with increased morbidity and mortality in anaemic surgical patients.<sup>[32]</sup> Intraoperative hypotension was demonstrated to be associated with myocardial injury, acute kidney injury and death.<sup>[33]</sup>

The present study identifies the intraoperative environment as one associated with a high risk of adverse outcomes in the frail. These increased adverse outcomes in the frail should ultimately emphasise the need for intraoperative vigilance and highlight the significance of frailty for all perioperative physicians, including the anaesthetist and not just the surgeon or intensivist.

Univariate logistic regression analysis showed that frailty was associated with an increased risk of intraoperative complications such as hypotension, desaturation, need for vasopressors and blood transfusion. Frailty was only significantly associated with desaturation and need for blood transfusion after multivariable logistic regression analysis, which is possibly a function of a small sample size, with fewer frail patients in the sample than anticipated. Patients with higher levels of frailty on the CFS were more likely to receive an ASA-PS score of  $\geq$ 3, which is in keeping with the findings of Robinson *et al.*<sup>[34]</sup> There was an association between frailty and

Table 3. Effect of fra	ulty on intraop	on intraoperative complications			l. l		
	Unr	Univariate logistic model to assess			Multivariable logistic regression controlling for other		
Factors		effect of frailty	the walking		of the second se	complications	
Association between f	frailty and hypot	95% CI	<i>p</i> -value	aUK	95% CI	<i>p</i> -value	
Frailty	francy and hypot	linei iiita					
No	1						
Ves	2.08	0.99 - 1.36	0.051	_	_	_	
Association between f	2.00 frailty and hypot	0.99 - 4.30	0.031	-	-	-	
Frailty	francy and hypot						
No	1						
Yes	1.87	1.08 - 3.25	0.025	1.87	0.81 - 4.34	0.14	
Age category (years	s)	1.00 5.25	0.020	1.07	0.01 1.01	0.11	
18 - 34	1						
35 - 49	3.04	1.23 - 7.49	0.01	0.80	0.24 - 2.68	0.72	
50 - 64	4 38	1.23 7.19	0.001	1.76	0.54 - 5.70	0.34	
>65	8.0	3.06 - 20.87	0.000	1.65	0.44 - 6.16	0.45	
Sex	010	20107	01000	1100	0111 0110	0110	
Female	1						
Male	0.81	0.509 - 1.300	0.394	-	-	-	
Comorbidities, <i>n</i>							
0	1						
1	1.94	1.08 - 3.50	0.02	0.77	0.31 - 1.90	0.57	
2	2.67	1.33 - 5.34	0.005	0.74	0.25 - 2.16	0.58	
3	1.84	0.78 - 4.35	0.16	0.64	0.17 - 2.38	0.51	
4	1.10	0.20 - 6.06	0.90	-	-	-	
Association between f	frailty and desatu	uration					
Frailty							
No	1						
Yes	3.79	1.36 - 10.54	0.01	4.21	1.31 - 13.53	0.01	
Association between f	frailty and need	for blood transfusion					
Frailty	1						
No	1						
Yes	3.26	1.13 - 9.36	0.02	5.36	1.50 - 19.16	0.01	
Association between f	frailty and arrhy	thmia					
Frailty	, ,						
No	1						
Yes	2.34	0.38 - 14.35	0.35	-	-	-	
Association between f	frailty and need	for vasopressors					
Frailty		•					
No	1						
Yes	2.81	1.60 - 4.91	0.00	1.57	0.83 - 2.98	0.16	
Age category (years	s)						
18 - 34	1						
35 - 49	1.76	0.70 - 4.46	0.22	1.67	0.64 - 4.31	0.28	
50 - 64	4.18	1.69 - 10.33	0.002	3.24	1.24 - 8.47	0.01	
≥65	8	3.06 - 20.87	0.00	6.23	2.17 - 17.82	0.001	
Sex							
Female	1						
Male	0.79	0.48 - 1.28	0.34	1.03	0.60 - 1.79	0.89	
Comorbidities, n							
0	1						
1	1.63	0.87 - 3.84	0.12	1.05	0.52 - 2.09	0.88	
2	3.33	1.63 - 3.77	0.00	1.74	0.77 - 3.91	0.18	
3	3.02	1.27 - 7.14	0.01	1.74	0.66 - 4.58	0.26	
4	1.38	0.25 - 7.61	0.71	0.41	0.06 - 2.58	0.34	

# Table 3. Effect of frailty on intraoperative complications

OR = odds ratio; CI = confidence interval: aOR = adjusted odds ratio.

Type of surgery	Frail, <i>n</i> (%)	Non-frail, <i>n</i> (%)	Total, <i>n</i> (%)	
Orthopaedic	44 (65.7)	56 (24.1)	100 (33.4)	
General surgery	9 (13.4)	73 (31.5)	82 (27.4)	
Urology	5 (7.5)	33 (14.2)	38 (12.7)	
Gynaecology	0	30 (12.9)	30 (10.0)	
Neurosurgery	3 (4.5)	9 (3.9)	12 (4.0)	
Ear, nose and throat	2 (3.0)	10 (4.3)	12 (4.0)	
Vascular	4 (6.0)	6 (2.6)	10 (3.3)	
Plastics	0	8 (3.4)	8 (2.7)	
Maxillofacial	0	6 (2.6)	6 (2.0)	
Not specified	0	1 (0.4)	1 (0.3)	
Total	67 (100)	232 (100)	299 (100)	

postoperative destination, with the frail being more likely to go to a high-care unit, but there was no significant association between frailty and ICU admission owing to sample size. The clinical significance of this is unclear.

Although the prevalence of frailty is highest among the elderly, there is a noteworthy prevalence among younger age groups. This prevalence could increase with the increase in communicable and noncommunicable diseases such as HIV, tuberculosis and traumatic injury.

#### Study strengths and limitations

A strength of the study is that it is the only study of its kind to be conducted in SA looking specifically at the effect of frailty on intraoperative complications. The study also included patients from a few of the largest hospitals in sub-Saharan Africa. The patients were largely from a low socioeconomic group and were representative of the patient population that is seen at most SA hospitals, as only a small proportion of the SA population can afford private hospital care.

Limitations include the fact that assessment of frailty in a perioperative setting is confounded by the effects of the presenting surgical condition. To minimise the effects of acute surgical conditions, only patients presenting for elective procedures were included.

Data were documented by different anaesthesia providers at different levels of anaesthetic training, and inter-observer reliability was not assessed. It was, however, important to demonstrate the utility of the CFS as a screening tool in a real-life clinical setting where it was intended for use by non-geriatricians. The CFS has shown good inter-observer reliability in a previous study.<sup>[35]</sup>

This was a single-centre study with a relatively small sample size. Future larger studies may focus on the association between frailty and postoperative complications, morbidity, length of stay and mortality in an SA surgical population.

#### Conclusions

The prevalence of frailty was high among surgical patients. Consistent with other studies, frailty was associated with older age and multiple comorbidities. The association between frailty and intraoperative complications found in this study may indicate and help inform areas of further research.

#### Declaration. None.

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