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Shrestha, D, Wisely, N, Bampouras, T, Subar, D, Shelton, C and Gaffney, C Exploring the association between socioeconomic status and cardiopulmonary exercise testing measures: a cohort study based on routinely collected data. PLoS ONE. ISSN 1932-6203 (Accepted)

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1 Original Article

Exploring the association between socioeconomic status and cardiopulmonary exercise testing measures: a
 cohort study based on routinely collected data.

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- 22 Keywords: Social determinant of health, CPX, health inequities, gas exchange threshold, anaerobic threshold,
- 23 peak VO₂
- 24 Short title: Inequalities in cardiopulmonary exercise testing measures

25 Abstract

26 **Background:**

27 Cardiopulmonary exercise testing (CPET) provides objective measures of cardiorespiratory fitness and can 28 support surgical risk stratification. As socioeconomic status is a factor known to influence patient health and

29 outcomes, we analysed how CPET-derived measures vary across levels of socioeconomic status in patients 30 being considered for elective surgery.

31 Methods:

32 A database of patients who underwent CPET between 2011 and 2024 was analysed. Measures including oxygen 33 consumption ($\dot{V}O_2$) at gas exchange threshold (GET), peak $\dot{V}O_2$, and ventilatory equivalent for carbon dioxide 34 (VE/VCO₂) were compared across socioeconomic deprivation quintiles. Multivariable linear and logistic 35 regression models assessed the effects of age, sex, body mass index (BMI), Revised Cardiac Risk Index (RCRI), 36 and deprivation quintiles on CPET measures. Hierarchical regression models incorporating the Indices of 37 Deprivation (IoD) domains and Access to Healthy Assets and Hazards (AHAH) scores determined whether wider 38 social determinants of health explained the variance in CPET measures.

39 **Results:**

40 A total of 3344 patients (2476 male) were included, referred prior to procedures in vascular (2006), colorectal

41 (650), upper GI (267), urology (205), and other (216) surgical specialties. Lower socioeconomic status was

42 associated with younger age (p<0.001), higher BMI (p=0.022), higher smoking prevalence (p<0.001), and RCRI

43 ≥3 (p=0.013). CPET measures were lower in the most deprived quintile (Q1) compared to the least (Q5): mean

44 GET was 11.0 vs. 11.5 ml·kg⁻¹·min⁻¹ and peak VO₂ was 14.8 vs. 16.3 ml·kg⁻¹·min⁻¹ (p<0.05). Deprivation remained

45 an independent predictor of lower GET and peak VO2, even after adjustment. Several IoD and AHAH domains

46 explained small but significant variance in CPET measures.

47 **Conclusion:**

48 Patients from more deprived areas exhibit risk factors for poor health and lower cardiorespiratory fitness as

49 measured by CPET. These findings add to our understanding of socioeconomic disparities in physiological

50 reserve among surgical patients and may support the need for more holistic approaches to peri-operative

51 care.

52 Introduction

A cardiopulmonary exercise test (CPET) provides objective measures of cardiorespiratory and metabolic fitness and is commonly used to aid the preoperative assessment of patients undergoing surgery. CPETs are increasingly being performed in the United Kingdom (UK), with around 30,000 tests performed annually (1). The results are used to inform the prediction of a patient's risk of postoperative complications, tailor patient optimisation and prehabilitation, and plan postoperative care (2, 3). CPET data can also aid patient-centred shared decision making, contributing to a more informed consent.

59 The COVID-19 pandemic shed light on the growing health disparities in the UK (4) and catalysed initiatives to 60 reduce health inequalities, building on those set out in the 2019 National Health Service (NHS) Long Term Plan 61 (5). The NHS in England emphasised the need to restore services inclusively, analyse health inequalities using 62 robust datasets, and accelerate preventative programmes targeting those at greatest risk of poor outcomes 63 (6). The Health and Care Act 2022 mandates that integrated care systems consider health inequalities in all 64 decision making (7). Regarding perioperative pathways, the 2023/24 NHS standard contract set out that 65 providers must "implement a system of early screening, risk assessment and health optimisation for all adult 66 service users waiting for inpatient surgery" (8). Therefore, understanding how CPET can be applied in the 67 context of significant health inequalities is an important consideration.

Socioeconomic deprivation is associated with poorer surgical outcomes, including higher rates of postoperative complications and mortality (9-11). The reasons for these disparities are likely multifactorial, and may stem from disparities in access and utilisation of healthcare services (12, 13), lifestyle factors (14) and co-morbidities (15). Patients from deprived backgrounds also report inferior healthcare experiences (16). Identifying whether cardiorespiratory fitness differs by socioeconomic status could help uncover one potential physiological mechanism contributing to these inequalities.

74 Although CPET measures are known to predict adverse surgical outcomes (2), the relationship between 75 socioeconomic deprivation and cardiorespiratory fitness in preoperative patients is yet to be explored. 76 However, studies in healthy young adults do report lower levels of cardiorespiratory fitness in those from 77 poorer socioeconomic backgrounds (17, 18) and a meta-analysis including 9435 people from four population-78 level studies demonstrated a positive association between higher educational attainment and 79 cardiorespiratory function (19). These studies suggest that socioeconomic status is associated with differences 80 in cardiorespiratory fitness in the general population, but it remains unclear whether similar patterns are 81 observed in preoperative CPET measures among surgical patients.

Within the study setting of Greater Manchester, forty percent of patients who live within the areas served by the Greater Manchester Integrated Care Board live in the twenty percent most socioeconomically-deprived areas in England (20). The region faces significant deprivation-related social challenges (21) and disparities in health outcomes such as life expectancy between small geographical areas (22).

By examining the relationship between socioeconomic factors and CPET measures, this study aims to inform equitable preoperative care strategies and contribute to efforts to reduce health inequalities. We hypothesised that patients from more socioeconomically deprived areas would have lower Gas Exchange Threshold (GET) and peak oxygen consumption (peak $\dot{V}O_2$) values, and higher ventilatory equivalent for carbon dioxide (VE/ $\dot{V}CO_2$), even after adjusting for clinical and demographic factors.

91 Methods

92 *Ethics*

Ethical approval was granted by Lancaster University (FHM-2024-4326-IRAS-2) and the NHS Health Research
 Authority (24/HRA/1302). Data were collected prospectively as part of routine care, between 02/09/2011 and

19/04/2024. The requirement for individual patient consent was waived in accordance NHS data governance
policies. Following ethical approval, the database was accessed for research purposes on 17/06/2024 by a
member of the direct care team and an anonymised database was passed on to the researchers for analysis.

98 Inclusion/Exclusion Criteria

99 The patient cohort included individuals being considered for surgery, who were referred from a range of NHS 100 sites in Greater Manchester. The major specialties from which they were referred included vascular, colorectal, 101 urology, and upper gastrointestinal surgery, and all specialties were included in the study. Patients who were

102 unable to cycle or did not have a postcode were excluded from the analysis.

103 Study Design

104 This study is a retrospective analysis of a prospectively collected database consisting of consecutive adult 105 patients undergoing CPET at Wythenshawe Hospital, Manchester, UK.

106 Variables recorded consisted of patient characteristics including age, sex, body mass index (BMI), smoking

107 status, revised coronary risk index (RCRI), static spirometry and CPET measures. The CPET measures that were

included in the study were GET, peak $\dot{V}O_2$, and $VE/\dot{V}CO_2$ at GET. Of the spirometry measures, the forced

109 expiratory volume in first second (FEV₁) to forced vital capacity (FVC) ratio (FEV₁/FVC) was used for the analysis.

Although commonly referred to as the "anaerobic threshold" in the clinical and perioperative literature (3),
 'GET' is used in this study to reflect the physiological basis of the measurement, which corresponds to the first

410 A CET is used in this study to renect the physiological basis of the measurement, which corresponds to the inst

- ventilatory threshold. A GET<11 ml·kg⁻¹·min⁻¹ was used as an exploratory marker of reduced cardiorespiratory
 fitness, consistent with values cited in the perioperative literature as associated with poor outcomes of intra-
- abdominal surgery (23, 24). While optimal thresholds in GET for risk prediction may vary between surgical
- 115 specialties (2), this value serves as a pragmatic reference point for comparison in preoperative assessment.

116 In the absence of a co-morbidity score or list, the RCRI was used as a surrogate for cardiovascular co-morbidities

and FEV₁/FVC for respiratory co-morbidities. An RCRI ≥3 is thought to be associated with a 15% [95% CI 11.1-

118 20.0%) risk of major cardiac event, defined as death, myocardial infarction, or cardiac arrest at 30 days after

119 noncardiac surgery (25) and this risk was compared by deprivation quintiles.

Analysis was performed on the whole dataset including all specialties and a sub-group analysis was carried outon vascular patients only (supplement 1).

122 CPET methodology

123 Each CPET was undertaken by a Perioperative Exercise Testing and Training society (POETTS) accredited 124 consultant anaesthetist and conducted in accordance with their consensus guidelines (3). Tests were 125 performed using an electromagnetically braked cycle ergometer (Corival CPET Cycle Ergometer, Lode B.V. 126 Zernikepark 16 9747 AN Groningen, The Netherlands). The exercise protocol included a patient specific ramp 127 of either 12, 15 or 20 Watts per minute aiming for around 10 minutes of exercise until symptoms limited. A 128 rapid gas analyser (Ultima™ CardiO2[®] Gas Exchange Analysis System, MGC Diagnostics Corporation, 350 Oak 129 Grove Parkway, Saint Paul, MN 55127) was used alongside continuous 12-lead ECG, pulse oxygen saturation 130 and non-invasive blood pressure monitor, whilst the patient was at rest, undertaking unloaded cycling and 131 pedalling against a predetermined and increasing resistance. Cardiorespiratory data was managed using Breeze 132 Suite software (MCG Diagnostics, Saint Paul, MN, US). The GET was determined using a combination of the V-133 slope, ventilatory equivalents and ventilation curve analysis. Peak oxygen consumption was calculated by 134 recording the highest value for oxygen consumption during the test (26).

135 Index of Multiple Deprivation (IMD) quintiles

- 136
 Patients' English Index of Multiple Deprivation (IMD) 2019 deciles were derived from individual postcodes, to
- stratify the cohort by their neighbourhood deprivation as a surrogate for their socioeconomic status. The IMD
- incorporates scores from seven indices of deprivation (IoD) domains: 'income', 'employment', 'health and disability', 'education, skills and training', 'crime', 'barriers to housing and services', and 'living environment'
- 140 (27). These domains included a total of 37 indicators, each measuring the proportion of the population
- 141 experiencing specific types of deprivation. In England, 32,844 Lower-layer Super Output Areas (LSOA), each
- 142 containing an average of 1,500 residents, are ranked according to their IMD score. For this analysis, patients'
- 143 IMD deciles were re-categorised into quintiles, where Q1 represents the 20% most deprived areas, and Q5 the
- 144 20% least deprived areas.

145 Access to Healthy Assets and Hazards (AHAH) scores

146 In addition to the IoDs, the Access to Healthy Assets and Hazards (AHAH) database was also used to explore 147 wider social factors which could contribute towards the variance in CPET measures. The AHAH index is an open 148 access index of accessibility to, both positive and negative, environmental health-related amenities and 149 exposures across LSOAs in Great Britain (28, 29). The AHAH domains consists of retail environment (access to 150 fast food outlets, pubs, tobacconists, gambling outlets), health services (access to GPs, hospitals, pharmacies, 151 dentists, leisure services), physical environment (blue space, green space - passive), and air quality (NO₂, PM10, 152 SO₂). Version 3, released in 2022 was used for this analysis. In the case of IoD and AHAH domain scores, the 153 higher the score, the greater the deprivation.

154 Statistical analysis

- Normality was confirmed if the ratio of skewness and kurtosis values to their respective errors was within ± 2. Baseline patient characteristics, CPET and spirometry measures were compared across the IMD quintiles (Q1-Q5) using descriptive statistics. For continuous variables, significant differences across the quintiles were assessed using one-way analysis of variance (ANOVA), followed by Tukey's post-hoc test to determine interquintile differences and the corrected values are reported. For categorical variables, comparisons were made using log-linear regression.
- 161 Multivariable linear regression models were conducted to assess the association between each CPET outcome 162 variable - peak $\dot{V}O_2$, GET and VE/ $\dot{V}CO_2$ – and predictor variables including deprivation quintile, age, sex, BMI, 163 smoking status, RCRI, and FEV₁/FVC. Regression coefficients with 95% confidence intervals were reported to 164 quantify the effect size of each predictor on the outcomes. A binomial multivariable logistic regression model 165 was used to identify significant predictors of having GET<11 ml·kg-1·min-1 and, specifically, whether 166 deprivation was a significant predictor of GET<11 ml·kg-1·min-1 when accounting for the other predictor 167 variables. Odds ratios with 95% confidence intervals were reported to estimate the strength of associations.
- Hierarchical multiple linear regression analyses were used to understand the association between IoD domain scores and AHAH domain scores, and three CPET outcome variables (peak VO₂, GET and VE/VCO₂). A baseline model, including age and sex, was generated to which each additional deprivation domain was added to create new models. Age and sex were included as baseline covariates in the first block of the hierarchical regression model as they are non-modifiable demographic factors known to influence CPET measures (30, 31). This approach allowed us to assess the additional variance in CPET outcomes explained by area-level deprivation domains (IoD and AHAH) beyond underlying physiological variation.
- 175 R^2 values were reported, representing the overall model's explanation of variance in each dependent variable. 176 The change in R-squared (ΔR^2) and F-statistic (F (ΔR^2)) was reporting, representing the model's ability to explain 177 additional variance in the outcome variable with the addition of individual deprivation domains, whilst 178 accounting for age and sex (included in the baseline model).

- For all regression models, variables were checked for multicollinearity (variance inflation factor <5, tolerance >0.5) and Cook's Distance values were within acceptable ranges (-2.5 to 2.5), confirming that no data transformation was required. Statistical significance was set at p < 0.05 for all analyses. Data analysis was conducted using Jamovi (version 2.4.8, The Jamovi Project, Sydney, Australia).
- 183

184 Results

A total of 3344 patients underwent CPET between September 2011 and April 2024. Most patients were assessed for vascular surgery (2006, 60%), followed by colorectal 650 (19.4%), urology 205 (6.1%), upper GI 267 (8%), and other specialties accounted for 216 (6.5%).

Patient characteristics are reported in Table 1. The percentage of patients in each quintile, Q1 (most deprived)
Q5 (least deprived), was 28.7%, 18.7%, 15%, 18.8% and 18.8% respectively. 2476 (74%) patients were male
and the proportion of males to females varied significantly across the deprivation groups. The mean patient
age was 72.1 (9.6) years. There was a significant difference in age (p<0.001) across the five deprivation groups;
the mean age in the most deprived quintile (Q1) was 69.5 compared to 74.1 in the least deprived quintile (Q5).
BMI was also higher in the most deprived quintile (Q1) (p=0.022) with mean BMI 28.3 (6.4) vs. 27.5 kg.m² (5.0)
in Q1 and Q5, respectively.

There was a significant difference in smokers across the quintiles (p<0.001), with a greater proportion of smokers in more deprived groups (Q1, 32.1% vs. Q5, 13.4%; p<0.001) and, similarly, there was a significantly greater proportion of patients who have never smoked in the least deprived quintile (Q1, 14.8% vs. Q5, 30.2%; p<0.001). There was a difference in the proportion of patients with RCRI \ge 3 (main effect: p<0.001; Q1, 13.2% vs. Q5, 8.9%; p=0.013). The mean FEV₁/FVC was significantly lower 65.9% (13.2) in Q1 compared to 70.0% (12.7) in Q5 (p=0.002). The main differences in the proportion of patients with FEV₁/FVC < 70% was seen in Q4 vs Q1 (46.9% vs. 54.0%, p=0.006) and Q4 vs Q2 (46.9% vs. 54.4%, p=0.008).

202

203 Table 1. Comparison of patient characteristics by IMD quintile.

	Missing	Q1	Q2	Q3	Q4	Q5	p-value **
People in Greater	uata						
Manchester		40.2	21.4	13 5	13.6	11 3	
Integrated Care		40.2	21.4	15.5	15.0	11.5	
Board (2023) (%)							
Total n (%)		960 (28.7)	626 (18.7)	500 (15)	629 (18.8)	629 (18.8)	
Age	1	69.5,10.0	71.1, 9.9	72.9, 9.5	74.1, 8.5	74.1, 9.0	< 0.001
Comparison to Q1*			0.011	<0.001	<0.001	<0.001	
Comparison to Q2*				0.014	<0.001	<0.001	
Comparison to Q3*					0.177	0.179	
Comparison to Q4*						1.000	
BMI kg/m²	15	28.3, 6.4	27.8, 5.3	27.6, 5.3	27.7, 5.4	27.5, 5.0	0.022
Comparison to Q1*			0.386	0.113	0.210	0.022	
Comparison to Q2*				00.958	0.998	0.794	
Comparison to Q3*					0.995	0.996	
Comparison to Q4*						0.928	
Sex – Male n (%)	0	684 (71.3)	460 (73.5)	367 (73.4)	482 (76.6)	483 (76.7)	<0.001
Comparison to Q1			0.333	0.385	0.018	0.015	
Comparison to Q2				0.975	0.198	0.176	
Comparison to Q3					0.212	0.190	
Comparison to Q4						0.947	
Smoker n (%)	1	308 (32.1)	148 (26.3)	111 (22.2)	90 (14.3)	84 (13.4)	<0.001
Comparison to Q1			<0.001	<0.001	<0.001	<0.001	
Comparison to Q2				0.568	<0.001	<0.001	
Comparison to Q3					<0.001	<0.001	
Comparison to Q4						0.616	
Never smoked n (%)	1	142 (14.8)	107 (17.1)	119 (23.8)	158 (25.1)	190 (30.2)	<0.001
Comparison to Q1			0.219	<0.001	<0.001	<0.001	
Comparison to Q2				0.005	<0.001	0.001	
Comparison to Q3					0.598	0.017	
Comparison to Q4						0.046	
RCRI ≥ 3 n (%)	1	127 (13.2)	57 (9.1)	40 (8)	69 (11)	56 (8.9)	<0.001
Comparison to Q1			0.013	0.003	0.181	0.009	
Comparison to Q2				0.512	0.272	0.900	
Comparison to Q3					0.095	0.589	
Comparison to Q4						0.221	
FEV ₁ /FVC %	27	65.9, 13.2	66.4, 13.2	70.2, 12.3	70.5, 12.3	70.0, 12.7	<0.001
Comparison to Q1*			0.954	0.165	0.004	0.002	
Comparison to Q2*				0.594	0.075	0.042	
Comparison to Q3*					0.868	0.766	
Comparison to Q4*						0.999	
FEV ₁ /FVC < 70%	27	513 (54.0)	339 (54.4)	244 (49.2)	294 (46.9)	310 (49.9)	<0.001
n (%)			0.075	0.000		0.415	
Comparison to Q1			0.872	0.083	0.006	0.113	
Comparison to Q2				0.083	0.008	0.113	
Comparison to Q3					0.443	0.809	
Comparison to Q4						0.284	

204 205 Patient characteristics (n = 3344), displayed for each IMD quintile (Q1 – most deprived to Q5 – least deprived). Data is presented as mean, SD, unless otherwise stated. Percentages are representative of the proportion of

206 patients in each IMD quintile with the variable, apart from the first row which is a percentage of the total

207 population. Significant p-values(p<0.05) are in bold. *Tukey-adjusted post-hoc p-values. **Main effect of 208 ANOVA.

209 In comparison of CPET measures by IMD quintile (Table 2), GET varied significantly across the five quintiles

210 (p=0.001), with a difference of 0.5 ml·kg⁻¹·min⁻¹ between Q1 and Q5 (p=0.04). 9.2% more patients had a GET<11

211 ml·kg-1·min-1 in Q1 than in Q5 (p=0.002). Peak $\dot{V}O_2$ decreased with deprivation, with a difference in means of

212 1.5 ml·kg⁻¹·min⁻¹ between Q1 and Q5. VE/ $\dot{V}CO_2$ varied significantly between Q1 and Q5 with a difference in

213 means of 1.3, Q1-Q5 (35.9 \pm 7.3 vs. 34.6 \pm 6.6, p=0.003). There was no significant difference in right or left

214 hand grip strengths in the 1201 patients who had this measured.

215 Table 2. Comparison of CPET variables by IMD quintile

	Missing data	Q1	Q2	Q3	Q4	Q5	p-value **
Total n (%)		960 (28.7)	626 (18.7)	500 (15)	629 (18.8)	629 (18.8)	
GET (ml·kg ⁻¹ ·min ⁻¹)	748	11.0, 2.2	11.3, 2.2	11.5, 2.6	11.3, 2.4	11.5, 2.5	0.001
Comparison to Q1*			0.123	0.003	0.087	0.004	
Comparison to Q2*				0.705	1.000	0.847	
Comparison to Q3*					0.759	0.998	
Comparison to Q4*						0.890	
GET<11 ml·kg	748	346 (50%)	206 (42.2)	171(42.4)	232 (45.8)	207 (40.8)	<0.001
¹•min⁻¹							
n (%)							
Comparison to Q1			0.008	0.015	0.140	0.002	
Comparison to Q2				0.948	0.260	0.658	
Comparison to Q3					0.316	0.626	
Comparison to Q4						0.113	
peak VO₂ (ml∙kg	39	14.8, 3.9	15.4, 3.8	16.1, 4.1	16.0, 4.1	16.3, 4.19	<0.001
¹∙min⁻¹)							
Comparison to Q1*			0.031	<0.001	<0.001	<0.001	
Comparison to Q2*				0.061	0.072	0.002	
Comparison to Q3*					0.999	0.903	
Comparison to Q4*						0.767	
VE/VCO ₂	37	35.9, 7.3	35.1, 6.6	35.0, 6.6	35.1,6.3	34.6, 6.6	0.006
Comparison to Q1*			0.180	0.152	0.187	0.003	
Comparison to Q2*				1.000	1.000	0.712	
Comparison to Q3*					0.999	0.866	
Comparison to Q4*						0.699	
Right Hand Grip (kg)	2143	29.8,10.0	29.7,10.4	29.8,10.4	30.1, 9.9	29.8, 10.1	0.996
Left Hand Grip (kg)	2143	27.7, 9.6	27.7, 9.9	27.5, 9.7	28.6, 9.7	27.6, 9.58	0.750

216 CPET variables comparison displayed for each IMD quintile (Q1 – most deprived to Q5 – least deprived). Data

217 is presented as mean, SD, unless otherwise stated. Percentages are representative of the proportion of patients

in each IMD quintile with the variable, apart from the first row which is a percentage of the total population.

219 Significant p-values(p<0.05) are in bold. *Tukey-adjusted post-hoc p-values. **Main effect of ANOVA.

The logistic regression model, for GET<11 ml·kg⁻¹·min⁻¹ (Table 3), was statistically significant, $\chi^2(13) = 435$, p <0.001. The model explained 20.9% (Nagelkerke R²) of the variance in GET<11 ml·kg⁻¹·min⁻¹, and correctly classified 72.8% of cases. Age, sex, BMI, RCRI and FEV₁/FVC were significant predictors of having a GET<11 ml·kg⁻¹·min⁻¹. Female sex increased the likelihood of having a GET<11 ml·kg⁻¹·min⁻¹ significantly (OR 3.97 [CI 3.20, 4.92], p<0.001). Adjusting for patient characteristics, compared to Q1, patients in Q2, 3, and 5 were less likely to have a GET<11 ml·kg⁻¹·min⁻¹ (OR 0.73, p=0.016; 0.74, p=0.034; 0.72, p=0.013, respectively). Smoking was not a significant factor in predicting a GET<11 ml·kg⁻¹·min⁻¹ (p >0.05).

- The regression model, for GET (Table 3), was statistically significant F(13) = 51.6, p<0.001. The model explained 20.4% (adjusted R² = 0.204) of the variance in GET. Age, female sex, BMI, and RCRI were significant negative predictors of GET, whilst FEV₁/FVC was a significant positive predictor of GET. Compared to Q1, Q2-Q5 had a significant positive association with GET, (Q5 vs Q1, β = 0.34, p=0.007). Smoking was not a significant independent predictor for GET.
- The regression model, for peak \dot{VO}_2 (Table 3), was statistically significant F(13) = 99.6, p<0.001. The model explained 28.2% (adjusted R² = 0.282) of the variance in peak \dot{VO}_2 . Age, female sex, BMI, smoking and RCRI had a significant negative association with peak \dot{VO}_2 . Deprivation was a significant negative predictor of peak \dot{VO}_2 , with a β value of 1.16 [Cl 0.80, 1.52] for Q5 vs Q1 (p<0.001), suggesting that individuals in the least deprived quintile had a peak \dot{VO}_2 that was 1.16 ml·kg⁻¹-min⁻¹ greater than the most deprived quintile.
- The regression model, for VE/VCO₂ (Table 3), was statistically significant F(13) = 72.0, p<0.001. The model explained 22% (adjusted R^2 = 0.220) of the variance in VE/VCO₂. Age, female sex, smoking, and RCRI were significant predictors of higher (adverse) VE/VCO₂. Conversely, BMI, FEV₁/FVC and all levels of deprivation were associated with lower (more favourable) VE/VCO₂. However, patients in the least deprived quintile (Q5) had a
- 241 VE/ $\dot{V}CO_2$ that was 1.43 lower than the most deprived quintile, Q1 (β =-1.43 [Cl -2.06, -0.80], p=0.001).

242 Table 3. Results of regression models

	GET<11 ml∙kg⁻¹	∙min⁻¹	GET ml·kg ⁻¹ ·min ⁻¹		peak VO2 ml·kg ⁻¹ ·min ⁻¹		VE/VCO2	
Predictor	OR [CI]	p-value	Estimate (β)	p-value	Estimate (β)	p-value	Estimate (β)	p-value
Age	1.04 [1.03, 1.05]	<0.001	-0.04 [-0.05, -0.03]	<0.001	-0.10 [-0.11, -0.09]	<0.001	0.17 [0.15, 0.19	<0.001
Sex (F-M)	3.97 [3.20, 4.92]	<0.001	-1.59 [-1.79, -1.39]	<0.001	-2.97 [-3.24, -2.69]	<0.001	0.89 [0.41, 1.36]	<0.001
BMI	1.10 [1.08, 1.12]	<0.001	-0.11 [-0.13, -0.10]	<0.001	-0.20 [-0.22, -0.17]	<0.001	-0.24 [-0.28, -0.20]	<0.001
Current Smoker	0.98 [0.78, 1.23]	0.853	-0.06 [-0.28, 0.15]	0.572	-0.61 [-0.91, -0.30]	<0.001	1.61 [1.08, 2.14]	<0.001
(Yes-No)								
RCRI (2-1)	1.85 [1.53, 2.23]	<0.001	-0.75 [-0.93, -0.56]	<0.001	-1.19 [-1.46, -0.92]	<0.001	1.04 [0.58, 1.51]	<0.001
RCRI (3-1)	3.78 [2.68, 5.34]	<0.001	-1.47 [-1.79, -1.14]	<0.001	-2.45 [-2.90, -2.00]	<0.001	2.43 [1.65, 3.22]	<0.001
RCRI (4-1)	8.78 [3.81, 20.21]	<0.001	-2.40 [-3.04, -1.76]	<0.001	-4.06 [-4.90, -3.22]	<0.001	4.13 [2.60, 5.66]	<0.001
RCRI (5-1)	3.37 [0.83, 13.75]	0.090	-1.93 [-3.24, -0.62]	0.004	-2.42 [-4.55, -0.29]	0.026	1.86 [-1.86, 5.57]	0.327
FEV ₁ /FVC	0.98 [0.97, 0.99]	<0.001	0.03 [0.02, 0.03]	<0.001	0.06 [0.05, 0.07]	<0.001	-0.11 [-0.12, -0.09]	<0.001
IMD quintile (2-1)	0.73 [0.56, 0.94]	0.016	0.27 [0.03, 0.52]	0.029	0.40 [0.05, 0.75]	0.027	-0.78 [-1.39, -0.17]	0.012
IMD quintile (3-1)	0.74 [0.56, 0.98]	0.034	0.46 [0.19, 0.72]	<0.001	1.05 [0.67, 1.43]	<0.001	-1.07 [-1.74, -0.41]	0.002
IMD quintile (4-1)	0.85 [0.65, 1.09]	0.203	0.30 [0.05, 0.55]	0.018	1.07 [0.71, 1.43]	<0.001	-0.95 [-1.58, -0.33]	0.003
IMD quintile (5-1)	0.72 [0.56, 0.93]	0.013	0.34 [0.09, 0.59]	0.007	1.16 [0.80, 1.52]	<0.001	-1.43 [-2.06, -0.80]	<0.001

243 Multivariable logistic and linear regression models with independent variables - GET<11 ml·kg-1·min-1, AT, peak VO2, and VE/VCO₂. Q1 (most deprived) is the

244 reference value for deprivation. Significant p-values (p<0.05) are in bold.

- In the hierarchical regression model, to explain the variance in GET (Table 4), the baseline model, including age and sex were significant predictors (F = 106.2, p < 0.001), and explained 7.6% ($R^2 = 0.076$) of the variance in GET. Of the IoD domains, four of seven indicators, including the 'health deprivation and disability score', 'employment score', 'education, skills and training score', and 'crime score' each contributed a significant but small (1-1.6%) amount in explaining the variance in GET (p<0.001). Of the five AHAH domain scores, air quality, passive green space, and retail domain scores explained between 0.1%-0.3% of further variance in GET after adjusting for age and sex.
- In the hierarchical regression model, to explain the variance in peak \dot{VO}_2 (Table 5), the baseline model, including age and sex were significant predictors (F = 206, p < 0.001), and explained 11.1% (R² = 0.111) of the variance in peak \dot{VO}_2 . Of the IoD domains, five of seven predictors, including 'health deprivation and disability score', 'employment score', 'income score', 'education, skills and training score', and 'living environment score' were significant predictors of peak \dot{VO}_2 . These predictors explained between 0.04-2.9% of the additional variance from the baseline model. All AHAH domain scores were predictive of peak \dot{VO}_2 . The air quality score explained the greatest additional variance of 1.3%.
- In the hierarchical regression model, to explain the variance in VE/ $\dot{V}CO_2$ (Table 6), the baseline model, including age and sex were significant predictors were significant predictors (F = 135, p < 0.001) and explained 7.6% (R² = 0.076) of the variance in VE/ $\dot{V}CO_2$. 'Income', 'employment, education, skills and training', 'health deprivation and disability', and 'crime' scores were all significant predictors of VE/ $\dot{V}CO_2$, accounting for 0.1-1.6% of variance in VE/ $\dot{V}CO_2$. Of the AHAH domains, green space availability and retail domain scores were statistically significant predictors and explained 0.1-0.3% further variance in VE/ $\dot{V}CO_2$.

Predictors	R ²	F (R ²)	p-value (R ²)	ΔR ²	F (ΔR ²)	p-value (∆R ²)		
Indices of deprivation (2019) domain scores								
Age, Sex (baseline model)	0.076	106.2	<0.001	-	-	-		
+ Health deprivation and	0.092	87.7	<0.001	0.016	46.9	<0.001		
disability score								
+ Employment score	0.088	83.4	<0.001	0.012	34.9	<0.001		
+ Income deprivation score	0.087	82.7	<0.001	0.012	32.9	<0.001		
+ Education, skills, and	0.087	82.3	<0.001	0.011	31.8	<0.001		
training score								
+ Crime score	0.086	81.3	<0.001	0.010	29.3	<0.001		
+ Barriers to housing and	0.077	71.5	<0.001	7.05x x10 ⁻⁴	1.98	0.160		
services score								
+ Living environment	0.076	70.9	<0.001	1.07x10 ⁻⁴	0.301	0.583		
deprivation score								
Acces	s to Healtl	ny Assets	& Hazards (AHA	H) domain scor	res			
+ Air quality domain score	0.085	80.7	<0.001	0.001	27.4	<0.001		
+ NVDI value indicating	0.080	75.4	<0.001	0.005	12.70	<0.001		
Passive Green Space								
+ Retail domain score	0.079	73.8	<0.001	0.003	8.48	0.004		
+ Health domain score	0.076	71.4	<0.001	5.71x10 ⁻⁴	1.60	0.206		
+Distance to nearest	0.076	70.8	<0.001	3.61x10 ⁻⁶	0.010	0.920		
leisure facility								

265 **Table 4. Association between GET and IoD and AHAH domains**.

266 Predictor variables are in descending order of association with AT. Significant p-values (p<0.05) are in bold.

Predictors	R ²	F (R ²)	p-value (R ²)	ΔR ²	F (ΔR ²)	p-value (∆R ²)		
Indices of deprivation (2019) domain scores								
Age, Sex (baseline model)	0.111	206	<0.001	-	-	-		
+ Health deprivation and	0.140	179	<0.001	0.029	111	<0.001		
disability score								
+ Employment score	0.138	175	<0.001	0.027	101	<0.001		
+ Income deprivation score	0.138	176	<0.001	0.027	104	<0.001		
+ Education, skills, and	0.136	173	<0.001	0.025	96	<0.001		
training score								
+ Living environment	0.115	143	<0.001	0.004	15.3	<0.001		
deprivation score								
+ Crime score	0.111	138	<0.001	2.09 x10 ⁻⁴	0.774	0.379		
+ Barriers to housing and	0.111	138	<0.001	8.74 x10 ⁻⁵	0.324	0.569		
services score								
Acces	s to Healt	ny Assets	& Hazards (AHA	H) domain sco	res			
+ Air quality domain score	0.124	156	<0.001	0.013	49.0	<0.001		
+ Retail domain score	0.119	148	<0.001	0.008	28.5	<0.001		
+ NVDI value indicating	0.119	148	<0.001	0.008	28.2	<0.001		
Passive Green Space								
+ Health domain score	0.116	144	<0.001	0.004	16.4	<0.001		
+Distance to nearest	0.112	139	<0.001	0.001	4.05	0.044		
leisure facility								

Table 5. Association between peak $\dot{V}O_2$ and IoD and AHAH domains.

269 Predictor variables are in descending order of association with AT. Significant p-values(p<0.05) are in bold.

Table 6. Association between VE/ $\dot{V}CO_2$ and IoD and AHAH domains.

Predictors	R ²	F (R ²)	p-value (R ²)	ΔR ²	F (ΔR ²)	p-value (∆R ²)		
Indices of deprivation (2019) domain scores								
Age, Sex (baseline model)	0.076	135	<0.001	-	-	-		
+ Income deprivation score	0.092	111	<0.001	0.016	57.3	<0.001		
+ Employment score	0.091	110	<0.001	0.015	53.5	<0.001		
+ Education, skills, and	0.090	109	<0.001	0.014	51.2	<0.001		
training score								
+ Health deprivation and	0.088	107	<0.001	0.013	45.4	<0.001		
disability score								
+ Crime score	0.077	92.0	<0.001	0.001	4.97	0.026		
+ Barriers to housing and	0.076	90.6	<0.001	3.00 x10 ⁻⁴	1.07	0.300		
services score								
+ Living environment	0.076	90.6	<0.001	2.71 x10 ⁻⁴	0.970	0.325		
deprivation score								
Acces	s to Healtl	ny Assets	& Hazards (AHA	H) domain scor	res			
+ NVDI value indicating	0.079	94.2	<0.001	0.003	11.2	<0.001		
Passive Green Space								
+ Retail domain score	0.077	91.6	<0.001	0.001	3.91	0.048		
+ Health domain score	0.077	91.4	<0.001	9.65x10 ⁻⁴	3.45	0.063		
+ Air quality domain score	0.076	91	<0.001	5.98 x10 ⁻⁴	2.14	0.144		
+Distance to nearest	0.076	90.4	<0.001	1.58x10 ⁻⁴	0.566	0.452		
leisure facility								

271 Predictor variables are in descending order of association with AT. Significant p-values(p<0.05) are in bold.

272 Discussion

273 This study is the first in the UK to explore the relationship between social determinants of health and 274 cardiorespiratory function, in adult patients who have undergone CPET as part of an elective surgical pathway. 275 We report four key findings: 1) patients from more deprived areas had greater cardiovascular and respiratory 276 risk factors, while they also had a lower age and higher BMI; 2) Deprivation was an independent risk factor for 277 lower cardiorespiratory function; 3) a gradient of adverse risks with increasing deprivation was apparent in 278 terms of BMI, smoking status, FEV₁/FVC, and three CPET measures; 4) Several indicators for the wider 279 determinants of health were small but significant factors in explaining the variance in CPET measures. We 280 discuss these findings in turn and consider their implications for understanding socioeconomic disparities in 281 cardiorespiratory fitness and how the findings may inform equitable perioperative care strategies.

1. Age, BMI, and comorbidity differences by socioeconomic deprivation

283 In our cohort, patients from more deprived quintiles were younger but had a higher BMI, greater prevalence 284 of smoking, and more respiratory compromise, as indicated by lower FEV1/FVC ratios. They were also more 285 likely to have a RCRI score \geq 3 and at greater risk of multimorbidity, which is consistent with the Scottish 286 database study of over 1.7 million of the general population by Barnett et al. (15). In the emergency surgical 287 setting, Poulton et al. demonstrated similar trends in deprivation-related age and co-morbidities in the 288 National Emergency Laparotomy Audit data (32). This clustering of cardiovascular and respiratory risk factors 289 among younger, more deprived patients may suggest that physiological decline occurs earlier in this group and 290 may partly explain the variation in CPET performance across socioeconomic deprivation quintiles.

291 2. Association between CPET measures and socioeconomic deprivation

292 In our cohort, patients from deprived areas were more likely to have a lower GET and a GET<11 ml kg⁻¹ min⁻¹. 293 The GET represents the VO₂ at which carbon dioxide production begins to rise disproportionately to oxygen 294 uptake. While GET is considered less dependent on maximal effort than peak VO₂ and is widely used as a 295 reproducible marker of functional capacity, its identification can vary depending on protocol, patient 296 characteristics (such as age, sex and comorbidities), and early test termination (31, 33). In our study, 712 297 patients who had a peak $\dot{V}O_2$ recorded, did not have a GET documented, likely reflecting these factors. This 298 finding is consistent with larger clinical cohort studies such as Marzolini et al (31), where an identifiable GET 299 was only observed in 69% of females and 88% of males, despite structured protocols. In our data, missing GET 300 was more common in the most deprived quintiles, Q1 (26.25%) and Q2 (21.25%) versus Q4 (18.76%) and Q5 301 (18.44%). This pattern may reflect differences in exercise tolerance or premature fatigue in more deprived 302 patients, potentially linked to underlying comorbidities or reduced physiological reserve (34). It also highlights 303 the importance of a nuanced interpretation of CPET results within the broader clinical context, as incomplete 304 tests may disproportionately affect those already at higher risk.

Patients in the most deprived groups also had a significantly lower peak $\dot{V}O_2$, i.e. the highest $\dot{V}O_2$ value achieved during the CPET. Quintile 1 had a mean peak $\dot{V}O_2$ of 14.8 ml·kg⁻¹·min⁻¹, which is clinically significant, as a peak $\dot{V}O_2 < 15 \text{ ml·kg}^{-1} \cdot \text{min}^{-1}$ is a significant predictor of early mortality in patients following AAA repair (35). A lower peak $\dot{V}O_2$ in the more deprived groups could be, partly, reflective of greater risk of co-morbidities.

A VE/VCO₂>34 has been shown to be predictive of adverse postoperative outcomes (2, 36). The mean VE/VCO₂,
 in our study, ranged between 34.6-35.9, with increasing values from the most to least deprived quintiles.

311 **3.** The social gradient in risk factors and CPET measures

The 'social gradient' in health, as demonstrated across our findings, is a well-established phenomenon (37) described across a range of health risks and outcomes in the general population, including smoking (14),

314 multimorbidity (15) and premature mortality (38). Health inequalities impact across the socioeconomic

315 spectrum, and is not limited to those in the most deprived socioeconomic groups (37). Greater psychosocial 316 stressors (39) and reduced access to health services (40, 41), further compound the effects of deprivation. 317 Evidence also suggests that the cumulative effect of adverse risks is greater in patients from more deprived 318 be deprived access to health services (40, 41), further compound the effects of deprivation.

backgrounds and that they are disproportionately affected (42, 43).

319 Another possible explanation for the observed social gradient in health involves the biosocial perspective (44-320 46), which considers the interplay between exogenous socio-environmental factors and endogenous biological 321 processes. It offers a holistic view of the individual, their susceptibility to disease, ability to adapt to adverse 322 risks from a physiological, psychological and social perspective, and goes beyond attributing differences to 323 "traditional" patient modifiable risk factors. Although our study did not measure biological markers, emerging 324 evidence suggests that socioeconomic status, for instance, is associated with differences in gut microbiome 325 independent of healthy diet, BMI and health deficits (47), potentially affecting inflammation and metabolic 326 function (48). Similarly, variation in epigenetic markers, linked with accelerated ageing (49), inflammation and 327 responses to stress (50, 51) have been demonstrated in patients from different socioeconomic backgrounds. 328 These biological shifts, amongst others, could contribute to the deprivation-related variance in 329 cardiorespiratory fitness through impaired metabolic efficiency (52, 53). While speculative, this hypothesis 330 highlights the need for future research that integrates social and biological determinants of health and are 331 important considerations as we strive to practice personalised medicine (54).

332 4. The wider determinants of health and CPET measures

333 From a wider public health perspective, we considered the variance in CPET measures in terms of a range of 334 social determinants of health. Factors such as health deprivation, employment, income, education, living 335 environment, air quality, green space availability, and access to adverse retail environments were all small but 336 significant contributors to the variance in cardiorespiratory fitness and are critical areas for policymakers to 337 consider in efforts to improve population health. An understanding of these broader determinants, and the 338 ways they intersect to impact individuals, can guide the multidisciplinary team in designing prehabilitation 339 programs. Providing support that is responsive to these factors requires a coordinated approach between 340 primary care, secondary care, and local authorities, incorporating strategies like social prescribing, 341 partnerships with community organisations, and initiatives to overcome individual logistical barriers. Future 342 research should focus on understanding patients' needs to inform service design that is equitably accessible 343 and sensitive to the intersectionality of deprivation-related factors.

344 Informing equitable perioperative care strategies

345 Although this study focuses on patients being considered for planned surgery, the deprivation-related 346 differences in cardiorespiratory function we observed are reflective of those in non-surgical populations (19). 347 Cardiorespiratory fitness is modifiable and, in the general UK population, is an independent risk factor for all-348 cause mortality (55, 56). In our study, patients from more deprived backgrounds were younger and exhibited 349 poorer cardiopulmonary fitness along with a higher prevalence of risk factors for adverse health. These findings 350 highlight the importance of the NHS' 'making every contact count' approach (57) which promotes early 351 opportunistic interventions to improve health before patients reach the point of surgical referral or 352 preoperative assessment. However, in the context of perioperative care, given our findings, and that patients 353 from deprived backgrounds also face greater challenges in managing their health (40, 58), an earlier and more 354 comprehensive assessment could be beneficial.

The differences in cardiorespiratory fitness by socioeconomic status observed in our study have important implications for prehabilitation. In the surgical patient population, multimodal prehabilitation is potentially an effective intervention in modifying cardiorespiratory fitness. This has been demonstrated in randomised controlled trials through improvements in functional capacity, measured by increases in 6-minute walking test (6MWT) or CPET measures (59, 60), corresponding to improvements in postoperative outcomes for both cancer and benign surgical patients (61, 62). Importantly, CPET measures can guide personalised goals and exercise interventions in prehabilitation (63), particularly when using CPET protocols that reliably enable identification of key metabolic thresholds such as the GET (33, 64). This represents a departure from a 'one size fits all' approach to exercise. Embedding this personalisation into routine perioperative care could help maximise benefit, particularly for patients with lower baseline fitness and higher preoperative risk, as seen in more deprived groups.

CPET, if used equitably, could provide vital information for patient optimisation. In addition to its role in patient postoperative risk stratification, CPET, along with spirometry, can be used to identify undiagnosed or unoptimised cardiorespiratory conditions, which could prompt further investigations (65). For example, ECG changes in response to exercise, or delayed heart rate recovery after exercise correspond with RCRI≥3 and increased adverse cardiac events (66).

371 Health inequalities are evident throughout the perioperative pathway, with patients from deprived 372 backgrounds experiencing longer waiting times for elective surgery (67). Moving away from a 'waiting list' 373 model to a 'preparation list' provides a unique opportunity to bridge this gap (68, 69). By using this time 374 proactively through patient engagement initiatives, such as 'surgical schools,' healthcare professionals can 375 enhance health literacy and provide culturally nuanced, holistic and inclusive support. For patients from 376 deprived backgrounds, early identification of risk through comprehensive preoperative assessment, and 377 tailored support during this period could help alleviate the adverse effects of deprivation. This is particularly 378 important, as patients from socioeconomically disadvantaged backgrounds are often those who stand to 379 benefit the most from prehabilitation but may be less likely to participate in such programs (70).

380 Additional findings

381 In addition to the observed differences by socioeconomic status, our analysis also identified important 382 disparities in CPET measures by sex, which may have further implications for personalised risk stratification 383 and optimisation strategies. The differences in CPET measured by sex have been previously reported by 384 Thomas et al. in 703 patients, who reported significantly lower GET and peak VO₂ in females, after adjusting 385 for weight (71). Similar findings of differences in peak VO_2 were reported in the post hoc analysis of the METS 386 study, which additionally identified optimal sex-specific peak VO₂ thresholds for postoperative complication 387 prediction (30). Our analysis adds to these findings; in our cohort, females were almost four times more likely to have a GET<11 ml·kg⁻¹·min⁻¹, after adjusting for age, BMI, smoking, RCRI, FEV₁/FVC, and deprivation. 388 389 Similarly, adjusting for these, females were likely to have significantly lower GET by 1.59 ml kg⁻¹ min⁻¹, peak VO₂ 390 by 2.97 ml kg⁻¹ min⁻¹ and greater VE/VCO₂ by 0.89. Other significant findings by Thomas et al. included that, on 391 average, of the participants included in 17 studies related to perioperative CPET, 68.5% were male. In our 392 cohort, 74% were male. This may be reflective of the epidemiology of the surgical specialties involved (e.g., 393 vascular surgery), but more research is required to clarify whether this is the case. For example, although 394 peripheral artery disease and abdominal aortic aneurysms are more prevalent in men (72), this would not 395 account for our 80% male vascular cohort. Sex-related inequalities in the surgical pathway have been described 396 in several areas (73-75) but the mechanisms remain under researched. Tailoring interventions to address these 397 physiological differences could optimise outcomes for both sexes, especially in patients from higher-risk 398 groups, though further research is needed to support this.

In our cohort, there were greater proportions of patients in the least deprived quintiles compared to the deprivation distribution within the Greater Manchester Integrated Care Board. Given that research suggests that deprived patients have a higher incidence of medical conditions and utilise greater NHS resources (76), we would expect the deprivation distribution in our cohort to reflect that of the general population, or be skewed towards more deprived patients. Possible explanations for this discrepancy could include a greater 404 tendency for emergency rather than elective presentations amongst more deprived groups, higher levels of 405 self-advocacy in more affluent patients, or clinician bias in referrals for CPET. Further research into the patient 406 pathway would be required to ensure equitable access at each stage. However, it should be noted that our 407 cohort consisted of 60% vascular patients. For those being referred following abdominal aortic aneurysm 408 screening, the referral pathway is clearly defined. The results are not dissimilar where a subgroup analysis was 409 performed for vascular patients alone (supplement 1).

410 Strengths

411 Despite the known challenges in identifying metabolic thresholds in clinical CPET settings, particularly in 412 heterogeneous or deconditioned populations (77) this study has several methodological strengths. First, it 413 draws on a large cohort spanning over a decade of routine CPETs. Second, all tests were conducted using cycle 414 ergometry with ramp-incremental protocols, which are more conducive to threshold detection than treadmill-415 based step protocols. Third, the testing protocol was standardised with interpretation by POETTS-accredited 416 consultant anaesthetists. Notably, approximately 60% of tests were performed by a single consultant 417 anaesthetist, reducing inter-operator variability, with the remaining tests conducted by five other trained 418 consultant anaesthetists.

419 Limitations

This study is not without limitations. First, it should be acknowledged that, even when standardised rampincremental protocols are applied and tests are conducted by trained personnel, many CPET protocols remain suboptimal for the accurate determination of metabolic thresholds such as the GET. Recent evidence shows that a significant proportion of clinical CPETs fail to detect GET reliably, which may impact both the interpretation and the comparability of fitness-related variables (33).

Second, the Index of Multiple Deprivation (IMD), as an area-level socioeconomic measure, does not capture individual-level deprivation, which could introduce some misclassification of socioeconomic status at the individual level. Second, the RCRI and FEV₁/FVC ratio were used as surrogates for comorbidities due to the unavailability of detailed comorbidity data. Additionally, there is heterogeneity within the study cohort, as it includes patients from multiple specialties and some who may not have proceeded to surgery. Not all patients who underwent CPET were subsequently placed on a surgical waiting list, which may limit comparability with studies that included only patients who ultimately underwent surgery.

Furthermore, the database did not capture each patient's specific diagnosis. While surgical specialty and broad categories of intended procedures were available, these do not distinguish between underlying pathology or indicate the severity or the functional impact of the condition. Future prospective studies should aim to collect more granular clinical data to adjust for clinical factors that may influence CPET performance.

We also acknowledge that our study is based on data from a single healthcare trust and is representative of the specific socioeconomic and healthcare challenges locally, limiting the generalisability to other populations. A national CPET database, or future studies in multi-centre cohorts, would allow for external validation of our findings. Nevertheless, a strength of the study is that it offers insights that can be contextualised to the region, that might otherwise be obscured in national-level datasets.

441 Conclusion

In summary, our study highlights significant socioeconomic disparities in cardiorespiratory fitness among preoperative patients, emphasising the need for equity-focused approaches in perioperative care. By incorporating both medical and social determinants into prehabilitation and risk stratification, healthcare providers can better support patients, ultimately contributing to improved surgical outcomes and addressing broader health inequalities.

447 **Data statement**:

The data underlying this study were collected as part of routine clinical care within the NHS. Patients were not specifically consented for their data to be made publicly available, and the study's ethical approvals do not permit open access sharing. Anonymised data may be made available upon reasonable request, subject to review and approval by the Anaesthetic Department, Wythenshawe Hospital, care of Dr Richard Lowe (richard.lowe@mft.nhs.uk).

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454 References

Reeves T, Bates S, Sharp T, Richardson K, Bali S, Plumb J, et al. Cardiopulmonary exercise testing
 (CPET) in the United Kingdom-a national survey of the structure, conduct, interpretation and funding.
 Perioperative Medicine (London, England). 2018;7:2.

458 2. Moran J, Wilson F, Guinan E, McCormick P, Hussey J, Moriarty J. Role of cardiopulmonary exercise
459 testing as a risk-assessment method in patients undergoing intra-abdominal surgery: a systematic
460 review. British Journal of Anaesthesia. 2016;116(2):177-91.

461 3. Levett DZH, Jack S, Swart M, Carlisle J, Wilson J, Snowden C, et al. Perioperative cardiopulmonary
462 exercise testing (CPET): consensus clinical guidelines on indications, organization, conduct, and
463 physiological interpretation. British Journal of Anaesthesia. 2018;120(3):484-500.

464 4. Marmot M, Allen J, Goldblatt P, Herd E, Morrison J. Build Back Fairer: The COVID-19 Marmot 465 Review. The Pandemic, Socioeconomic and Health Inequalities in England. London: Institute of Health 466 Equity; 2020.

467 5. NHS England. The NHS Long Term Plan 2019. Available from: 468 https://www.longtermplan.nhs.uk/about/

469 6. NHS England. 2021/22 Priorities and operational planning guidance: Implementation guidance
470 NHS England 2021. Available from: https://www.england.nhs.uk/publication/2021-22-priorities-and471 operational-planning-guidance/

4727.UKGovernment.HealthandCareAct2022.Availablefrom:473https://www.legislation.gov.uk/ukpga/2022/31/contents/enacted

- 474 8. NHS England. Earlier screening, risk assessment and health optimisation in perioperative 475 and integrated care boards pathways: guide for providers 2023. Available from: 476 https://www.england.nhs.uk/long-read/earlier-screening-risk-assessment-and-health-optimisation-in-477 perioperative-pathways/
- Poulton TE, Salih T, Martin P, Rojas-Garcia A, Raine R, Moonesinghe SR. Systematic review of the
 influence of socioeconomic deprivation on mortality after colorectal surgery. British Journal of Surgery.
 2018;105:959-70.
- 481 10. Stamatiou D, Naumann DN, Foss H, Singhal R, Karandikar S. Effects of ethnicity and
 482 socioeconomic status on surgical outcomes from inflammatory bowel disease. International journal of
 483 colorectal disease. 2022;37(6):1367-74.
- 484 11. Wan YI, McGuckin D, Fowler AJ, Prowle JR, Pearse RM, Moonesinghe SR. Socioeconomic
 485 deprivation and long-term outcomes after elective surgery: analysis of prospective data from two
 486 observational studies. British journal of anaesthesia. 2021;126(3):642-51.
- Propper C. Socio-economic-inequality in distribution of healthcare in the UK. Institute for Fiscal
 Studies. 2022. Available from: https://ifs.org.uk/inequality/socio-economic-inequality-in-thedistribution-of-healthcare-in-the-uk/
- 490 13. Smith SG, McGregor LM, Raine R, Wardle J, Von Wagner C, Robb KA. Inequalities in cancer
 491 screening participation: examining differences in perceived benefits and barriers. Psycho-Oncology.
 492 2016;25(10):1168-74.
- 49314.Kotz D, West R. Explaining the social gradient in smoking cessation: it's not in the trying, but in the494succeeding. Tobacco Control. 2009;18(1):43-6.
- 495 15. Barnett K, Mercer SW, Norbury M, Watt G, Wyke S, Guthrie B. Epidemiology of multimorbidity and
 496 implications for health care, research, and medical education: a cross-sectional study. The Lancet.
 497 2012;380(9836):37-43.
- 498 16. Wilson RJT, Yates DRA, Walkington JP, Davies SJ. Ventilatory inefficiency adversely affects
 499 outcomes and longer-term survival after planned colorectal cancer surgery. British Journal of
 500 Anaesthesia. 2019;123(2):238-45.
- 501 17. Finger JD, Krug S, Gosswald A, Hartel S, Bos K. Cardiorespiratory fitness among adults in
 502 Germany: results of the German Health Interview and Examination Survey for Adults (DEGS1).
 503 Bundesgesundheitsblatt Gesundheitsforschung Gesundheitsschutz. 2013;56(5-6):772-8.

504 18. Shishehbor MH, Gordon-Larsen P, Kiefe CI, Litaker D. Association of neighborhood
505 socioeconomic status with physical fitness in healthy young adults: The Coronary Artery Risk
506 Development in Young Adults (CARDIA) study. American Heart Journal. 2008;155(4):699-705.

507 19. Ombrellaro KJ, Perumal N, Zeiher J, Hoebel J, Ittermann T, Ewert R, et al. Socioeconomic
508 Correlates and Determinants of Cardiorespiratory Fitness in the General Adult Population: a Systematic
509 Review and Meta-Analysis. Sports Medicine - Open. 2018;4(1).

510 20. NHS England. Change to integrated care board (ICB) core services target allocation (waterfalls)
 511 NHS England 2023. Available from: <u>https://www.england.nhs.uk/publication/change-to-icb-core-</u>
 512 services-target-allocation-waterfalls/. Accessed:15/09/2024

- 513 21. Marmot M, Allen J, Broyce T, Goldblatt P, Morrison J. Building Back Fairer in Greater Manchester:
 514 Health Equity and Dignified Lives London: Institute of Health Equity. 2021. Available from:
 515 https://www.instituteofhealthequity.org/resources-reports/build-back-fairer-in-greater-manchester516 health-equity-and-dignified-lives
- 517 22. Purdam K. The devolution of health funding in Greater Manchester in the UK: A travel map of life 518 expectancy. Environment and Planning A. 2017;49(7):1453-7.
- 51923.Older P, Hall A, Hader R. Cardiopulmonary Exercise Testing as a Screening Test for Perioperative520Management of Major Surgery in the Elderly. Chest. 1999;116(2):355-61.
- 521 24. Lai CW, Minto G, Challand CP, Hosie KB, Sneyd JR, Creanor S, et al. Patients' inability to perform 522 a preoperative cardiopulmonary exercise test or demonstrate an anaerobic threshold is associated with 523 inferior outcomes after major colorectal surgery. British Journal of Anaesthesia. 2013;111(4):607-11.
- 524 25. Duceppe E, Parlow J, Macdonald P, Lyons K, McMullen M, Srinathan S, et al. Canadian 525 Cardiovascular Society Guidelines on Perioperative Cardiac Risk Assessment and Management for 526 Patients Who Undergo Noncardiac Surgery. Canadian Journal of Cardiology. 2017;33(1):17-32.
- 527 26. Chambers DJ, Wisely NA. Cardiopulmonary exercise testing-a beginner's guide to the nine-panel
 528 plot. British Journal of Anaesthesia Education. 2019;19(5):158-64.
- 529 27. McLennan D, Noble S, Noble M, Emma Plunkett, Wright G, Gutacker N. The English Indices of
- 530 Deprivation 2019 Technical Report Ministry of Housing, Communities & Local Government. 2019. 531 Available from: https://www.gov.uk/government/publications/english-indices-of-deprivation-2019-532 technical-report
- 533 28. Green MA, Daras K, Davies A, Barr B, Singleton A. Developing an openly accessible multi-534 dimensional small area index of 'Access to Healthy Assets and Hazards' for Great Britain, 2016. Health 535 and Place. 2018;54:11-19.
- 536 29. Daras K, Green MA, Davies A, Barr B, Singleton A. Open data on health-related neighbourhood
 537 features in Great Britain. Scientific Data. 2019; 6:107.
- 30. Alfitian J, Riedel B, Ismail H, Ho KM, Xie S, Zimmer P, et al. Sex-related differences in functional
 capacity and its implications in risk stratification before major non-cardiac surgery: a post hoc analysis
 of the international METS study. eClinicalMedicine. 2023;64:102223.
- 54131.Marzolini S, Oh P, Peterman JE, Wallace P, Yadollahi A, Rivera-Theurel F, et al. Sex Differences and542Correlates of the Utility of the Cardiopulmonary Exercise Test for Prescribing Exercise at Entry to Cardiac543Rehabilitation. Canadian Journal of Cardiology. 2025;41(3):481-90.
- 544 32. Poulton TE, Moonesinghe R, Raine R, Martin P, National Emergency Laparotomy Audit project 545 team. Socioeconomic deprivation and mortality after emergency laparotomy: an observational 546 epidemiological study. British Journal of Anaesthesia. 2020;124(1):73-83.
- 547 33. Keltz RR, Hartley T, Huitema AA, McKelvie RS, Suskin NG, Keir DA. Do Clinical Exercise Tests
 548 Permit Exercise Threshold Identification in Patients Referred to Cardiac Rehabilitation? Canadian Journal
 549 of Cardiology. 2023;39(11):1701-11.
- 550 34. Shishehbor MH, Litaker D, Pothier CE, Lauer MS. Association of Socioeconomic Status With 551 Functional Capacity, Heart Rate Recovery, and All-Cause Mortality. Journal of the American Medical 552 Association 2006;295(7):784-92
- 552 Associaton. 2006;295(7):784-92.

35. Hartley RA, Pichel AC, Grant SW, Hickey GL, Lancaster PS, Wisely NA, et al. Preoperative
cardiopulmonary exercise testing and risk of early mortality following abdominal aortic aneurysm repair.
British Journal of Surgery. 2012;99(11):1539-46.

556 36. Wilson RJT, Davies S, Yates D, Redman J, Stone M. Impaired functional capacity is associated with 557 all-cause mortality after major elective intra-abdominal surgery. British Journal of Anaesthesia. 558 2010;105(3):297-303.

55937.Marmot MG, Smith GD, Stansfeld S, Patel C, North F, Head J, et al. Health inequalities among560British civil servants: The Whitehall II study. The Lancet. 1991;337(8754):1387.

561 38. Stringhini S, Carmeli C, Jokela M, Avendaño M, Muennig P, Guida F, et al. Socioeconomic status 562 and the 25 × 25 risk factors as determinants of premature mortality: a multicohort study and meta-563 analysis of 1.7 million men and women. The Lancet. 2017;389(10075):1229-37.

39. Matthews KA, Gallo LC, Taylor SE. Are psychosocial factors mediators of socioeconomic status
and health connections? A progress report and blueprint for the future. Annals of the New York Acadeny
of Sciences. 2010;1186:146-73.

56740.Moss C, Munford LA, Sutton M. Associations between inflexible job conditions, health and568healthcare utilisation in England: retrospective cross-sectional study. BMJ Open. 2022;12(12):e062942.

41. Asaria M, Ali S, Doran T, Ferguson B, Fleetcroft R, Goddard M, et al. How a universal health system
reduces inequalities: lessons from England. Journal of Epidemiology and Community Health.
2016;70(7):637-43.

42. Foster HME, Celis-Morales CA, Nicholl BI, Petermann-Rocha F, Pell JP, Gill JMR, et al. The effect of socioeconomic deprivation on the association between an extended measurement of unhealthy lifestyle factors and health outcomes: a prospective analysis of the UK Biobank cohort. The Lancet Public Health. 2018;3(12):e576-e85.

57643.Pampel FC, Rogers RG. Socioeconomic status smoking and health a test of competing theories577of cumulative advantage. Journal of Health and Social Behavior. 2004;45(3):306-21.

57844.Harris KM, McDade TW. The Biosocial Approach to Human Development, Behavior, and Health579Across the Life Course. The Russell Sage Foundation Journal of the Social Sciences. 2018;4(4):2-26.

580 45. Link BG, Phelan J. Social Conditions As Fundamental Causes of Disease. Journal of Health and
581 Social Behavior. 1995;35:80-94.

Glass TA, McAtee MJ. Behavioral science at the crossroads in public health: extending horizons,
envisioning the future. Social Science & Medicine. 2006;62(7):1650-71.

58447.Bowyer R, Jackson M, Le Roy C, Ni Lochlainn M, Spector T, Dowd J, et al. Socioeconomic Status585and the Gut Microbiome: A Twins UK Cohort Study. Microorganisms. 2019;7(1):17.

58648.de Vos WM, Tilg H, Van Hul M, Cani PD. Gut microbiome and health: mechanistic insights. Gut.5872022;71(5):1020-32.

588 49. Fiorito G, Polidoro S, Dugué P-A, Kivimaki M, Ponzi E, Matullo G, et al. Social adversity and 589 epigenetic aging: a multi-cohort study on socioeconomic differences in peripheral blood DNA 590 methylation. Scientific Reports. 2017;7:16266.

591 50. Needham BL, Smith JA, Zhao W, Wang X, Mukherjee B, Kardia SLR, et al. Life course 592 socioeconomic status and DNA methylation in genes related to stress reactivity and inflammation: The 593 multi-ethnic study of atherosclerosis. Epigenetics. 2015;10(10):958-69.

594 51. McGuinness D, McGlynn LM, Johnson PC, MacIntyre A, Batty GD, Burns H, et al. Socio-economic 595 status is associated with epigenetic differences in the pSoBid cohort. International Journal of 596 Epidemiology. 2012;41(1):151-60.

597 52. Estaki M, Pither J, Baumeister P, Little JP, Gill SK, Ghosh S, et al. Cardiorespiratory fitness as a 598 predictor of intestinal microbial diversity and distinct metagenomic functions. Microbiome. 2016;4:42.

59. Follitt RA, Kaufman JS, Rose KM, Diez-Roux AV, Zeng D, Heiss G. Cumulative life course and adult
socioeconomic status and markers of inflammation in adulthood. Journal of Epidemiology & Community
Health. 2008;62(6):484-91.

54. Vicente AM, Ballensiefen W, Jonsson JI. How personalised medicine will transform healthcare by 2030: the ICPerMed vision. Journal of Translational Medicine. 2020;18:180. 55. Blair SN, Kampert JB, Kohl HW, Barlow CE, Macera CA, Ralph S. Paffenbarger, et al. Influences of Cardiorespiratory Fitness and other precursors on cardiovascular disease and all cause mortality in men and women. Journal of the American Medical Association. 1996;276(3):205-10.

Kodama S, Saito K, Tanaka S, Maki M, Yachi Y, Asumi M et al. Cardiorespiratory Fitness as a
Quantitative Predictor of All-Cause Mortality and Cardiovascular Events in Healthy Men and Women.
Journal of the American Medical Association. 2009;301(19):2024-35.

61057.NHS England. Making Every Contact Count (MECC): Consensus statement NHS England 2016.611Availablefrom: https://www.england.nhs.uk/publication/making-every-contact-count-mecc-612consensus-statement/. Accessed: 05/09/2024.

- 58. Woodward A, Nimmons D, Davies N, Walters K, Stevenson FA, Protheroe J, et al. A qualitative
 exploration of the barriers and facilitators to self-managing multiple long-term conditions amongst
 people experiencing socioeconomic deprivation. Health Expectations. 2024;27(2):e14046.
- 59. Minnella EM, Bousquet-Dion G, Awasthi R, Scheede-Bergdahl C, Carli F. Multimodal
 prehabilitation improves functional capacity before and after colorectal surgery for cancer: a five-year
 research experience. Acta Oncologica. 2017;56(2):295-300.

60. Molenaar CJL, Minnella EM, Coca-Martinez M, Ten Cate DWG, Regis M, Awasthi R, et al. Effect of Multimodal Prehabilitation on Reducing Postoperative Complications and Enhancing Functional Capacity Following Colorectal Cancer Surgery: The PREHAB Randomized Clinical Trial. Journal of the American Medical Association Surgery. 2023;158(6):572-81.

- 623 61. Lambert JE, Hayes LD, Keegan TJ, Subar DA, Gaffney CJ. The Impact of Prehabilitation on Patient 624 Outcomes in Hepatobiliary, Colorectal, and Upper Gastrointestinal Cancer Surgery: A PRISMA-625 Accordant Meta-analysis. Annals of Surgery. 2021;274(1):70-7.
- 626 62. Liang MK, Bernardi K, Holihan JL, Cherla DV, Escamilla R, Lew DF, et al. Modifying Risks in Ventral
 627 Hernia Patients With Prehabilitation: A Randomized Controlled Trial. Annals of surgery. 2018;268(4):674628 80.
- 629 63. Van Rooijen S, Carli F, Dalton S, Thomas G, Bojesen R, Le Guen M, et al. Multimodal 630 prehabilitation in colorectal cancer patients to improve functional capacity and reduce postoperative 631 complications: the first international randomized controlled trial for multimodal prehabilitation. BMC 632 Cancer. 2019;19(1):98.
- 633 64. Inglis EC, Iannetta D, Rasica L, Mackie MZ, Keir DA, Macinnis MJ, et al. Heavy-, Severe-, and
 634 Extreme-, but Not Moderate-Intensity Exercise Increase Vo2max and Thresholds after 6 wk of Training.
 635 Medicine & Science in Sports & Exercise. 2024;56(7):1307-16.
- 636 65. Richardson K, Levett DZH, Jack S, Grocott MPW. Fit for surgery? Perspectives on preoperative 637 exercise testing and training. British Journal of Anaesthesia. 2017;119:i34-i43.
- 638 66. Abbott TEF, Pearse RM, Cuthbertson BH, Wijeysundera DN, Ackland GL. Cardiac vagal 639 dysfunction and myocardial injury after non-cardiac surgery: a planned secondary analysis of the 640 measurement of Exercise Tolerance before surgery study. British Journal of Anaesthesia. 641 2019;122(2):188-97.
- 642 67. Robertson R, Blythe N, Jefferies D. Tackling health inequalities on NHS waiting lists: learning from 643 local case studies. The King's Fund. 2023. Available from: 644 https://assets.kingsfund.org.uk/f/256914/x/98aabc1536/tackling_health_inequalities_in_waiting_lists_ 645 2023.pdf.
- 646 68. Levy N, Selwyn DA, Lobo DN. Turning 'waiting lists' for elective surgery into 'preparation lists'.
 647 British Journal of Anaesthesia. 2021;126(1):1-5.
- 648 69. Centre for Perioperative Care (CPOC). Tackling the elective surgery backlog perioperative care 649 solutions to the waiting list. 2021 Available from: https://www.cpoc.org.uk/cpoc-releases-new-poll-how-650 perioperative-care-can-help-reduce-waiting-list.
- 70. Lee D, Wang A, Augustin B, Buajitti E, Tahasildar B, Carli F, et al. Socioeconomic status influences
 participation in cancer prehabilitation and preparation for surgical recovery: A pooled retrospective
 analysis using a validated area-level socioeconomic status metric. European Journal of Surgical
 Oncology. 2023;49(2):512-20.

Thomas G, West MA, Browning M, Minto G, Swart M, Richardson K, et al. Why women are not
small men: sex-related differences in perioperative cardiopulmonary exercise testing. Perioperative
Medicine. 2020;9:18.

McGinigle KL, Browder SE, Strassle PD, Shalhub S, Harris LM, Minc SD. Sex-related disparities in
intervention rates and type of intervention in patients with aortic and peripheral arterial diseases in the
National Inpatient Sample Database. Journal of Vascular Surgery. 2021;73(6):2081-9 e7.

73. Howard R, Ehlers A, Delaney L, Solano Q, Shen M, Englesbe M, et al. Sex disparities in the
treatment and outcomes of ventral and incisional hernia repair. Surgical endoscopy. 2023;37(4):3061-8.
74. Jin G, Liu C, Fei X, Xu M. A systematic review and meta-analysis on sex disparities in the outcomes

664 of fenestrated branched endovascular aortic aneurysm repair. Journal of Vascular Surgery. 665 2023;77(6):1822-32 e3.

Merdji H, Long MT, Ostermann M, Herridge M, Myatra SN, De Rosa S, et al. Sex and gender
differences in intensive care medicine. Intensive Care Medicine. 2023;49(10):1155-67.

668 76. Cookson R, Propper C, Asaria M, Raine R. Socio-Economic Inequalities in Health Care in England.
669 Fiscal Studies. 2016;37(3-4):371-403.

670 77. Keir DA. Invited Commentary: Problems Using the Cardiopulmonary Exercise Test in Cardiac

671 Rehabilitation: It Is Time to Retire Outdated Protocols and Replace Them With Better Ones. Canadian

672 Journal of Cardiology. 2025;41(3):491-3.

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674 Supplementary Information

675 S1 – Vascular patient analysis