Global patient outcomes after elective surgery: Prospective cohort study in 27 low, middle and high income countries

Running title: Global patient outcomes after elective surgery

International Surgical Outcomes Study (ISOS) group*

*members of study group listed in supplementary file

Correspondence and requests for re-prints to:

Rupert M. Pearse
Adult Critical Care Unit
Royal London Hospital
London
E1 1BB
United Kingdom

e-mail: r.pearse@qmul.ac.uk Tel: +44 20 3594 0346

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Abstract

Introduction

As global initiatives increase patient access to surgical treatments, there remains a need to understand the adverse effects of surgery, and define appropriate levels of perioperative care at a global level.

Methods

Prospective international seven-day cohort study of outcomes following elective adult inpatient surgery. The primary outcome was in-hospital complications. Secondary outcomes were death following a complication (failure to rescue), and death in hospital. Process measures were admission to critical care immediately after surgery, or to treat a complication, and duration of hospital stay. A single definition of critical care was used for all countries.

Results

474 hospitals in 19 high, 7 middle and one low income country were included in the primary analysis. The dataset included 44814 patients with a median hospital stay of 4 (2-7) days. 7508 patients (16.8%) developed ≥1 postoperative complications, and 207 died (0.5%). The overall mortality amongst patients who developed complications was (2.8%). Mortality following complications ranged from 2.4% for pulmonary embolism to 43.9% for cardiac arrest. 4360 (9.7%) patients were admitted to a critical care unit as routine immediately after surgery, of whom 2198 (50.4%) developed a complication with 105 (2.4%) deaths. 1233 patients (16.4%) were admitted to a critical care unit to treat complications with 119 (9.7%) deaths. Despite lower baseline risk, outcomes were similar in low and middle, when compared to high income countries.

Conclusions

Poor patient outcomes are common after in-patient surgery. Global initiatives to increase access to surgical treatments should also address the need for safe perioperative care.

Introduction

Currently, 310 million patients undergo surgery worldwide each year, with more procedures taking place in high income countries.^{1, 2} Findings from epidemiological studies suggest that 4.8 billion people are unable to access safe surgical treatments,³ and that at least 143 million additional procedures are required each year, primarily in low and middle income countries.^{4, 5} However, as healthcare systems develop to improve access to surgical treatments, the number of patients who suffer postoperative complications will also increase.^{3, 4}

Postoperative complications increase treatment costs,⁶ and reduce both life expectancy and quality of life at a societal level.^{7, 8} Nonetheless, our global understanding of outcomes after surgery remains limited. Estimates from high income countries suggest postoperative complications occur in up to 20% of patients,^{9, 10} and short-term mortality may vary from 1 to 4%.¹¹⁻¹⁸ While effective perioperative care is considered essential to the safe provision of surgical treatments,⁸ the optimal level of such care has not been defined. Admission to a critical care unit is often considered necessary to prevent, or treat, life threatening complications. However, this standard of patient care is very expensive, and there is little or no evidence to confirm the critical care resource provision needed for a safe surgical service.

As we seek to ensure the global availability of surgical treatments to all patients, we need to understand how often patients develop complications after surgery, the severity of harm which results, and how hospital systems should be configured to safely respond. We performed the International Surgical Outcomes Study (ISOS) to evaluate the incidence and risk factors for complications and death after in-patient elective surgery at a global level, and to describe current standards of postoperative care.

Methods

Project organisation

ISOS was a seven-day international cohort study. Regulatory requirements differed between countries with some requiring research ethics approval and some requiring only data governance approval. In the UK, the study was approved by the Yorkshire & Humber Research Ethics Committee (Reference: 13/YH/0371). The inclusion criteria were all adult patients (age ≥18 years) undergoing elective surgery with a planned overnight stay in hospital. Each participating country selected a single data collection week between April and August 2014. Patients undergoing emergency surgery, day-case surgery or radiological procedures were excluded. Patient data included only that recorded as part of routine care. In some countries, patient consent was sought to allow the collection of supplementary data for pre-specified sub-studies. In each country, we approached individuals to act as national co-ordinators using contacts in national and international specialist societies in surgery and anaesthesia. Individual participating hospitals were then identified through a global online recruitment campaign led by the study management group, and through the direct approach of the national co-ordinators. Nominations for participation were then confirmed as appropriate through discussion with national co-ordinators. The study website provided all study documentation and guidance on study procedures (www.isos.org.uk/documents). ISOS was registered prospectively with an international trial registry (ISRCTN51817007).

Data collection

Data describing perioperative care facilities were collected for each hospital at the beginning of the study. Data describing consecutive patients were collected until hospital discharge on paper case record forms (Supplementary file). Complications were assessed according to predefined criteria and graded as mild, moderate or severe. ¹⁹ Data were censored at 30 days following surgery for patients who remained in hospital. Data were anonymised before entry onto a secure internet based electronic case record form designed specifically for ISOS, which incorporated automated checks for plausibility, consistency and completeness.

Outcome measures

The primary outcome measure was in-hospital postoperative complications. Secondary outcomes were death following a postoperative complication (failure to rescue), and in-hospital mortality. Process measures were admission directly to critical care after surgery, admission to critical care for treatment of a postoperative complication, and duration of hospital stay. A single prospective definition of critical care was used for all countries (a facility routinely capable of admitting patients who require invasive ventilation overnight).

Statistical analysis

We aimed to recruit as many hospitals and countries as possible, and asked investigators in those hospitals to enrol all eligible patients. No formal sample size calculation was performed. Only hospitals returning valid data describing 20 or more patients, and countries with ten or more participating hospitals were included in the primary analysis.

Association between surgical procedure category and patient outcomes

We assessed the association between surgical procedure category and complications or mortality both before and after adjustment for potential confounding factors. The unadjusted analysis was performed using a logistic regression model with the surgical procedure category included as a fixed factor. The adjusted analysis was performed using a three-level mixed-effects logistic regression model. Patients were entered at the first level, hospitals at the second and countries at the third level. This model accounted for correlation between patients in the same hospital or country. The following variables were included as fixed factors in the model: age, current smoker, American Society of Anesthesiologists physical status (ASA) score, severity of surgery, surgical procedure category and presence of ischaemic heart disease, heart failure, diabetes mellitus, chronic obstructive pulmonary disease/asthma, cirrhosis, stroke, and other co-morbid diseases. Factors were selected for biological plausibility, scientific rationale and a low rate of missing data. We used restricted cubic splines to account for a potential non-linear association between age and outcome. ²⁰ To assess the effect of predefined exclusions on our findings, we repeated our analyses for all patients in the database. For both the unadjusted and adjusted analyses, Hosmer-Lemeshow

goodness-of-fit statistics were used to test model calibration, and multi-collinearity was assessed using the variance inflation factor. The ability of the model to discriminate cases from non-cases was assessed using the area under the receiver operating characteristic curve (AUROC). Data are presented as mean (SD) and median (IQR) for continuous data, number (%) for binary data, or as odds ratios (OR) with 95% confidence intervals. Analyses were performed using Stata 14 (StataCorp, USA).

Results

Data describing 44814 patients were collected in 474 hospitals in the following countries and regions: Australia, Austria, Belgium, Brazil, Canada, China, Denmark, France, Germany, Greece, Hong Kong, Indonesia, Italy, Malaysia, Netherlands, New Zealand, Nigeria, Portugal, Romania, Russia, South Africa, Spain, Sweden, Switzerland, Uganda, United Kingdom, and the United States of America (Figure 1). Fewer than ten hospitals participated in India, Iraq and Mexico, and in accordance with the prospective statistical analysis plan, patients recruited in these countries were excluded from the primary analysis (Figure 2). Seven countries were classed middle income and one as low income, with 134 participating hospitals between them. Hospitals had a median of 550 (329-850) ward beds and 21 (10-38) critical care beds. The median critical care capacity (ratio of critical care beds to total hospital beds) was 4 (2-6) %. 310 hospitals (66%) were affiliated to a university. 77% of hospitals provided only government funded healthcare, 3% only privately funded healthcare, whilst 21% of hospitals were funded by both sources. Baseline patient data are presented in table 1.

Data validation

There was high concordance in a random 1% data sample selected for duplicate entry (95% for categorical variables, 92% for continuous variables), with very high concordance for clinical outcomes (99.7%). Investigators were granted immediate access to their uncleaned data once this was locked following entry, and were encouraged to review this for accuracy and completeness. All national co-ordinators confirmed the face validity of the baseline and crude outcome data for their countries. Only a small proportion of patients (451/44814 [1%]), were missing data for at least one of the factors included in the model. Due to the low proportion of missing data, we performed a complete case analysis, where patients with missing data were excluded from the analysis (Supplementary table 1). Hosmer-Lemeshow goodness-of-fit statistics indicated that the models were well calibrated, with a good match between observed and expected outcomes. The discrimination of the model was good with an AUROC of 0.80 (95% CI 0.80 - 0.81). Residuals showed that the assumptions for regression analyses were met. All variables had a variance inflation factor of less than five.

Clinical outcomes

A total of 7508 (16.8%) patients developed complications in hospital, and 207 died before hospital discharge (0.5%), indicating an overall mortality amongst patients who developed complications (failure to rescue) of (2.8%). 5254 (11.7%) patients developed a single postoperative complication whilst a further 2254 (5.0%) patients developed two or more complications. The breakdown of complications is presented in table 2. Infectious complications were the most frequent, in particular superficial surgical site infections. 2925 patients developed an unspecified complication ('other' category). There were significant variations in complications and mortality across surgical procedure categories and countries (Figure 3, Supplementary tables 2 and 3). Outcomes for patients according to planned admission to critical care immediately after surgery are presented in Table 3. 1233 patients (16.4%) were admitted to a critical care unit to treat complications of whom 119 (9.7%) died. 58 (28.0%) patients who died were not admitted to critical care at any stage during their admission, either immediately after surgery or for treatment of a complication. The clinical outcomes for all patients included in the database are presented in Supplementary table 4.

Process measures

The median stay in a post-anaesthetic care unit was 1 (0-2) hours. 4360 (9.7%) patients were admitted to a critical care unit as routine immediately after surgery. The median length of time spent in critical care for those with a planned admission directly after surgery was 1 (1-3) days. Of these patients, 2198 (50.4%) developed a complication with 105 (2.4%) deaths. 1233 (4.9%) patients were admitted to a critical care unit to treat complications of whom 119 (9.7%) died. The median length of time spent in critical care for patients admitted to treat a complication was 3 (1-6) days. The median overall hospital stay was 4 (2-7) days, increasing to 8 (5-14) days amongst those patients who developed complications.

Outcomes in low, middle and high income countries

Patient outcomes and process measures according to low and middle, or high income country status are presented in Table 4. One country in the low and middle income group, which returned a large patient sample, experienced much lower complication rates than other

participating nations. Patients in low and middle income countries tended to be younger with lower ASA scores. Crude complication rates were lower, but mortality rates overall, and for patients developing complications, were similar to those in high income countries, suggesting care for patients who develop complications may be less effective. There was a much lower rate of planned admission to critical care immediately after surgery in low and middle income countries.

Discussion

This international prospective cohort study has provided detailed outcome data on a population of more than 44,000 consecutive patients undergoing elective in-patient surgery in 27 low, middle and high income countries worldwide. The principal finding was that one in six patients experienced a complication before hospital discharge, and one in thirty-five patients who experienced a complication subsequently died without leaving hospital. The mortality amongst patients who developed complications (failure to rescue) of 2.8% indicates the continued need for a more effective treatment response for patients who develop postoperative complications. Despite lower baseline risk, crude patient outcomes were broadly similar in low and middle, compared to high income countries.

There are few large datasets of complication rates after surgery, and none we are aware of which provide data at an international level, although the findings of a recent study of almost 11,000 patients undergoing emergency abdominal surgery in 58 low, middle and high income countries indicate a high mortality following such procedures.²² Comparisons between country level datasets should be cautious because of international differences in patterns of surgical disease and genomics, as well as in healthcare systems. A variable degree of selection bias is also likely to result in important differences between reports which are few in number. Whilst overall complication rates in the current data were slightly lower than those previously reported in the USA,^{9, 23} this may simply be due to differences in patient risk factors and the surgical procedures included. In particular, ISOS only included patients undergoing elective surgery. Previous mortality estimates for unselected patient populations undergoing inpatient surgery vary between 1 and 4%.¹⁵⁻¹⁸ A recent study of postoperative mortality in Europe suggested an in-hospital mortality of 3% for elective in-patient surgery,¹¹ similar to the overall mortality rates in reports from the USA.^{9, 16, 23}

These data provide detailed insights into patterns of critical care admission after surgery. This is an expensive resource, and rates of admission in low and middle income countries appear to be much lower than high-income countries. The value of routine admission of high-risk patients to a critical care unit after surgery remains uncertain and allocation of this this

resource appears inconsistent. For example, admission to critical care after cardiac surgery is routine in most countries, whilst high-risk patients undergoing non-cardiac surgery may not be provided with this level of care despite a much higher mortality rate. 12-14 The findings of two recent healthcare registry studies in the UK suggest that provision of critical care may improve survival for surgical patients, although the effect may be subtle.^{24, 25} Meanwhile, a study of Medicare registry data in the US failed to identify any benefit of critical care admission.^{26, 27} Comparison of failure to rescue (rate of death after postoperative complications) between hospitals and healthcare systems may help us to understand the impact of postoperative critical care on patient outcomes. Whilst it seems unlikely that we could ever reduce the mortality from postoperative complications to zero, failure to rescue has provided a useful metric of the quality of postoperative care for surgical patients in high income countries. 9, 28-31 We could argue that, in a well-resourced system, very few patients should die after elective surgery without being admitted to a critical care unit. The current data confirm there is an important rate of failure to rescue at a global level, which is placed in context by the rates of use of critical care facilities. Global strategies to improve access to surgical treatments should take account of the increased demand for perioperative care services, in particular critical care, for those patients who develop complications.^{3,4} However, whilst the surgical population is very large, few countries have any reliable system to monitor the volume of activity and clinical outcomes. Understanding of the safety and effectiveness of surgical treatments is therefore limited and the need remains for robust audit and public reporting of outcomes after all surgery worldwide.8 Data driven improvement in quality of perioperative care may be possible even in resource limited environments.³²

The strengths of this study include the large number of consecutive patients enrolled worldwide. Importantly, critical care beds were classified according to a standard definition in all participating hospitals. We also distinguished between planned admission to critical care immediately after surgery as a part of routine postoperative management, and unplanned admission to critical care to treat a life threatening complication. By developing a very simple data set consisting primarily of categorical variables, we were able to minimise the amount of missing data. Patient-level variables were selected on the basis that they were objective,

routinely collected for clinical reasons, could be transcribed with a high level of accuracy, a low rate of missing data, and would be relevant to a risk adjustment model which included a wide variety of surgical procedures. The online data entry system was designed specifically for ISOS, and included a variety of internal error checks, whilst avoiding the redundant functionality of generic software designed for complex trials. The study also has a number of weaknesses. Despite the large sample size, we cannot consider this study as representative of current practice in all countries. ISOS was a pragmatic study and only a small proportion of hospitals took part in a small number of countries. Whilst we are pleased to have recruited hospitals in 30 countries, only 27 of these reached the predefined number of participating hospitals. We discussed participation with potential investigators in a number of countries who did not feel they had adequate resources to take part. This affected the participation of low, middle and high income countries. Many patients were enrolled in university hospitals whilst smaller, low volume centres are under-represented. This effect was greater in the low and middle income countries which took part. The risk adjustment methods used may not fully account for high mortality rates in hospitals specialising in more complex surgery. After risk adjustment, there were differences in postoperative outcomes between countries, but there are likely to be differences in casemix which are not fully represented in our baseline data.^{1, 2} We note that crude complication and mortality rates were lower in one high volume country, reducing the overall event rate. Given the pragmatic nature of this study, it was only possible to follow patients until hospital discharge. In countries where the availability of hospital beds is more limited, early hospital discharge of patients may have resulted in a lower measured complication rate. Although we planned to enrol every eligible patient undergoing surgery during the study period, we cannot be sure of the exact proportion of eligible patients included. Despite these limitations, assuming the volume of surgery during the cohort week is typical of the participating hospitals, these centres perform over 3 million in-patient surgical procedures each year, approximately 1% of the estimated volume of surgery taking place worldwide.1,2

Conclusions

The findings of this international cohort study indicate that a large number of patients develop complications after elective in-patient surgery. Global strategies to improve access to surgical treatments should take account of the increased demand placed on perioperative care services.

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Role of the funding source

This was an investigator initiated study funded by Nestle Health Sciences through an unrestricted research grant, by a National Institute for Health Research Professorship held by RP, and sponsored by Queen Mary University of London. ISOS investigators were entirely responsible for study design, conduct and data analysis. Members of the writing committee had full data access and were solely responsible for data interpretation, drafting and revision of the manuscript, and the decision to submit for publication. Nestle Health Sciences had no data access and no role in study design, conduct, analysis, or in drafting this report.

Contributors

RP conceived the study and designed this together with all members of the writing committee and steering committees. Patient recruitment and data collection were performed by the members of the ISOS study group (see supplementary file). TA and BK performed the data analysis with input from all members of the writing committee. The manuscript was drafted by RP and revised following critical review by all members of the writing committee and steering committees. The ISOS investigators would like to thank our patient representative, Naomi Pritchard, for her guidance and support throughout this project.

Data sharing

The authors are happy to consider data sharing requests from bona fide researchers. Enquiries should be addressed to the chief investigator at: admin@isos.org.uk.

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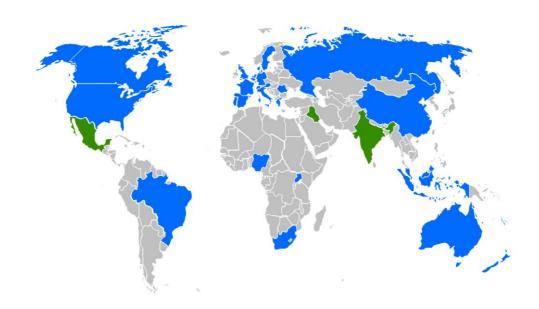
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Figure & table legends

Figure 1. Countries participating in the International Surgical Outcomes Study

Blue: countries included in primary analysis. Green: countries with fewer than ten participating hospitals included in secondary analysis.



Unsigned datasheet
(46 patients)

Unsigned datasheet
(46 patients)

Hospitals with fewer than
20 patients
(27 hospitals, 323
patients)

Countries with fewer than
10 hospitals
(3 countries, 8 hospitals,
462 patients)

474 hospitals available for analysis
(44,863 patients)

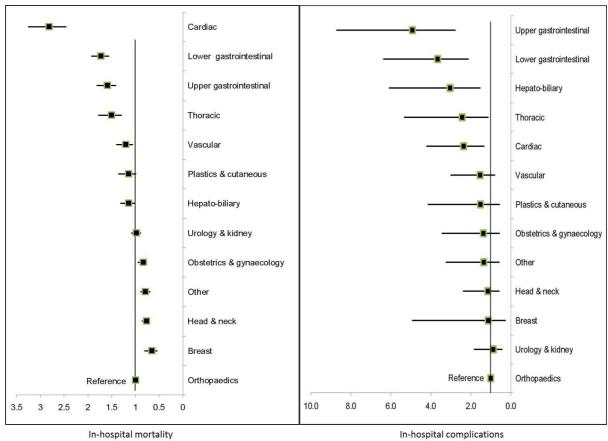
Missing outcome data
(49 patients)

Specialty hospitals
(3436 patients)

41,378 patients included in hospital level analysis

Figure 2. Patients, hospital and countries excluded from study.

Figure 3. Adjusted risk (odds ratio) of complications with 95% confidence intervals, and inhospital mortality in different surgical procedure categories.



In-hospital mortality

Table 1. Baseline patient characteristics.

All data presented as n (%). ASA, American Society of Anesthesiologists physical status score; COPD, chronic obstructive pulmonary disease.

	All patients	Patients with complications	Patients with no complications	Patients who died	Patients who survived	
	n = 44814	n = 7508	n = 37306	n = 207	n = 44607	
Age in years (mean SD)	55.3 (17.1)	61.8 (16.0)	54.1 (17.0)	69.1 (13.3)	55.3 (17.1)	
Age in years (median range)	57 (18-102)	64 (18-100)	55 (18-102)	73 (28-93)	57 (18-102)	
Male	20458 (45.7%)	3968 (19.4%)	16490 (80.6%)	121 (0.6%)	20337 (99.4%)	
Smoker	7913 (17.8%)	1305 (16.5%)	6608 (83.5%)	47 (0.6%)	7866 (99.4%)	
ASA score		•				
ı	11227 (25.1%)	848 (7.6%)	10379 (92.5%)	1 (0.1%)	11226 (99.9%)	
II	22265 (49.8%)	3005 (13.5%)	19260 (86.5%)	38 (0.2%)	22227 (99.8%)	
III	10193 (22.8%)	3090 (30.3%)	7103 (69.7%)	115 (1.1%)	10078 (98.9%)	
IV	1038 (2.3%)	554 (53.4%)	484 (46.6%)	53 (5.1%)	985 (94.9%)	
Severity of surgery		•	•			
Minor	8411 (18.8%)	672 (8.0%)	7739 (92.0%)	14 (0.2%)	8397 (99.8%)	
Intermediate	20203 (45.1%)	2494 (12.3%)	17709 (87.7%)	56 (0.3%)	20147 (99.7%)	
Major	16175 (36.1%)	4336 (26.8%)	11839 (73.2%)	137 (0.9%)	16038 (99.1%)	
Surgical procedure						

Orthopaedic	9459 (21.1%)	1556 (16.5%)	7893 (83.5%)	25 (0.3%)	9434 (99.7%)
Breast	1538 (3.4%)	128 (8.3%)	1410 (91.7%)	2 (0.1%)	1536 (99.9%)
Obstetrics & gynaecology	5674 (12.7%)	554 (9.8%)	5120 (90.2%)	6 (0.1%)	5668 (99.9%)
Urology & kidney	4871 (10.9%)	720 (14.8%)	4151 (85.2%)	10 (0.2%)	4861 (99.8%)
Upper gastro-intestinal	1986 (4.4%)	485 (24.4%)	1501 (75.6%)	29 (1.5%)	1957 (98.5%)
Lower gastro-intestinal	3073 (6.9%)	748 (24.3%)	2325 (75.7%)	32 (1.0%)	3041 (99.0%)
Hepato-biliary	2282 (5.1%)	366 (16.0%)	1916 (83%)	14 (0.6%)	2268 (99.4%)
Vascular	1599 (3.6%)	410 (25.6%)	1189 (74.4%)	15 (0.9%)	1584 (99.0%)
Head & neck	6510 (14.5%)	674 (10.4%)	5836 (89.6%)	12 (0.2%)	6498 (99.8%)
Plastics & cutaneous	1670 (3.7%)	244 (14.6%)	1426 (85.4%)	5 (0.3%)	1665 (99.7%)
Cardiac	1716 (3.8%)	979 (57.0%)	737 (43.0%)	40 (2.3%)	1676 (97.7%)
Thoracic	1157 (2.6%)	305 (26.4%)	852 (73.6%)	10 (0.9%)	1147 (99.1%)
Other	3270 (7.3%)	328 (10.0%)	2942 (90.0%)	7 (0.2%)	3263 (99.8%)
Co-morbid disease					
Ischaemic heart disease	4588 (10.3%)	1525 (33.2%)	3063 (66.8%)	67 (1.5%)	4521 (98.5%)
Heart failure	1882 (4.2%)	775 (41.2%)	1107 (58.8%)	49 (2.6%)	1833 (97.4%)
Diabetes mellitus	5171 (11.6%)	1319 (25.5%)	3852 (74.5%)	58 (1.1%)	5113 (98.9%)
Cirrhosis	342 (0.8%)	113 (33.0%)	229 (67.0%)	10 (2.9%)	332 (97.1%)
Metastatic cancer	1706 (3.8%)	508 (29.8%)	1198 (70.2%)	36 (2.1%)	1670 (97.9%)
Stroke	1492 (3.3%)	451 (30.2%)	1041 (69.8%)	38 (2.6%)	1454 (97.4%)

COPD / asthma	4094 (9.2%)	1012 (24.7%)	3082 (75.3%)	40 (1.0%)	4054 (99.0%)	
Other	18607 (41.6%)	4134 (22.2%)	14464 (77.8%)	134 (0.7%)	18473 (99.3%)	
Other measures	Other measures					
Laparoscopic surgery	7087 (15.8%)	905 (12.8%)	6182 (87.2%)	16 (0.2%)	7071 (99.8%)	
Cancer surgery	9006 (20.3%)	2005 (22.2%)	7001 (77.7%)	70 (0.8%)	8936 (99.2%)	

Table 2. Postoperative complications and mortality for 44814 patients undergoing elective surgery.

Data presented as n (%). ARDS, acute respiratory distress syndrome; N/A, category not applicable for this complication. Some patients may have developed more than one complication, and consequently in some cases the denominator is the number complications whilst in the left most column the denominator is the number of patients. The cell at the bottom of the far right column represents the number of deaths divided by the number of patients with at least one complication.

			Severity of complications			
	n=44814	Mild	Moderate	Severe	n = 207	
Infectious complications				<u> </u>	L	
Superficial surgical site	1320 (2.9)	681/1320 (51.6)	517/1320 (39.2)	122/1320 (9.2)	17/1320 (1.3)	
Deep surgical site	566 (1.3)	120/566 (21.2)	250/566 (44.2)	196/566 (34.6)	28/566 (4.9)	
Body cavity	340 (0.8)	97/340 (28.5)	136/340 (40.0)	107/340 (31.5)	24/340 (7.0)	
Pneumonia	708 (1.6)	240/708 (33.9)	325/708 (45.9)	143/708 (20.2)	55/708 (7.8)	
Urinary tract	681 (1.5)	294/681 (43.2)	333/681 (48.9)	54/681 (7.9)	13/681 (1.9)	
Bloodstream	417 (0.9)	140/417 (33.6)	162/417 (38.8)	115/417 (27.6)	48/417 (11.5)	
Total infectious complications	4032	1572/4032 (39.0)	1723/4032 (42.7)	737/4032 (18.3)	104/4032 (2.6)	
Cardiovascular complications		-	<u> </u>			
Myocardial infarction	139 (0.3)	45/139 (32.4)	43/139 (30.9)	51/139 (36.7)	26/139 (18.7)	
Arrhythmia	1222 (2.7)	468/1222 (38.3)	568/1222 (46.5)	186/1222 (15.2)	74/1222 (6.1)	
Pulmonary oedema	330 (0.7)	127/330 (38.4)	141/330 (42.8)	62/330 (18.8)	34/330 (10.3)	
Pulmonary embolism	78 (0.2)	17/78 (21.8)	33/78 (42.3)	28/78 (35.9)	5/78 (6.4)	

Stroke	111 (0.2)	31/111 (27.9)	28/111 (25.2)	52/111 (46.9)	18/111 (16.2)
Cardiac arrest	153 (0.3)	N/A	N/A	153/153 (100.0)	91/153 (59.5)
Total cardiovascular complications	2033	688/2033 (33.8)	813/2033 (40.0)	532/2033 (26.2)	141/2033 (6.9)
Other complications					
Gastro-intestinal bleed	201 (0.4)	95/201 (47.3)	66/201 (32.8)	40/201 (19.9)	24/201 (11.9)
Acute kidney injury	778 (1.7)	423/778 (54.4)	203/778 (26.1)	152/778 (19.5)	76/778 (9.8)
Post-operative bleed	1362 (3.0)	N/A	1147/1362 (84.2)	215/1362 (15.8)	55/1362 (4.0)
ARDS	142 (0.3)	46/142 (32.4)	41/142 (28.9)	55/142 (38.7)	34/142 (23.9)
Anastomotic leak	208 (0.5)	52/208 (25.0)	62/208 (29.8)	94/208 (45.2)	21/208 (10.1)
All others	2934 (6.5)	1342/2925 (45.9)	1200/2925 (41.0)	392/2925 (13.4)	83/2925 (2.8)
Total other complications	5625	1958/5625 (34.8)	2719/5625 (48.3)	948/5625 (16.9)	158/5625 (2.8)
Total number of complications	11690	4218/11690 (36.1)	5255/11690 (45.0)	2217/11690 (19.0)	207/7508 (2.8)

Table 3. Outcomes for patients according to planned admission to critical care immediately after surgery.

Data presented as n (%).

	All patients	Patients admitted to critical care immediately after surgery	Patients not admitted to critical care immediately after surgery
	n=44814	n=4360	n=39935
Mortality	207/44814 (0.5%)	105/4360 (2.4%)	99/39935 (0.2%)
Complication(s)	7508/44814 (16.8%)	2198/4360 (50.4%)	5270/39935 (13.2%)
Critical care admission to treat complication(s)	1233/7508 (16.4%)	857/2198 (39.0%)	365/5270 (6.9%)
Death following a complication (Failure to rescue)	207/7508 (2.8%)	105/2198 (4.8%)	99/5270 (1.9%)

Table 4. Hospital resources, process measures and patient outcomes in low, middle and high income countries. ASA, American Society of Anesthesiologists physical status score; Data presented as mean (SD), median (IQR), or n (%).

	Low and middle income countries (n=8)	High income Countries (n=19)
Number of hospitals	126	348
Number of patients	15806	29008
Hospital characteristics		
Total beds per hospital	825 (412 - 1318)	570 (361 - 835)
Critical care beds per hospital	25 (12 - 45)	20 (11 - 37)
Critical care capacity per hospital	2.8% (1.5% - 4.8%)	3.6% (2.4% - 5.9%)
Patient characteristics		
Age	50.8 (16.0)	57.8 (17.2)
ASA I & II	13766 (87.2%)	19726 (68.2%)
ASA III & IV	2029 (12.8%)	9202 (31.8%)
Co-morbid disease (any)	6488 (41.2)	19590 (67.6)
Metastatic cancer	297 (1.9)	1409 (4.9)
Process measures		
Post-anaesthetic care unit stay (hours)	1 (0 - 1)	1 (1 - 2)
Length of hospital stay (days)	5 (3 - 8)	3 (1 - 6)
Planned critical care admission	1051/15299 (6.9%)	3309/28996 (11.4%)
Critical care to treat complication(s)	317/15806 (2.0%)	916/28905 (3.2%)
Patient outcomes		
Complication(s)	1760/15806 (11.1%)	5748/29008 (19.8%)
Mortality	58/15806 (0.4%)	149/29008 (0.5%)
Mortality following complications	58/1760 (3.3%)	149/5748 (2.6%)

Global patient outcomes after elective surgery: Prospective cohort study in 27 low, middle and high income countries

Supplementary file

International Surgical Outcomes Study (ISOS) group*

*members of study group listed below

Writing committee:

Rupert M. Pearse, Professor of Intesnive Care Medicine, Queen Mary University of London, UK EC1M 6BQ Tahania Ahmad, Statistician, Queen Mary University of London, UK EC1M 6BQ Xiangming Fang, Professor of Anaesthesia, Zhejiang University, Hangzhou 310003, China Peter Holt, Senior Lecturer in Vascular Surgery, St Georges University of London, SW17 0RE Brennan C. Kahan, Senior statistician, Queen Mary University of London, UK E1 2AT Andrew Rhodes, Consultant in Intensive Care Medicine, St Georges University of London, SW17 0RE Duminda Wijeysundera, Assistant Professor, University of Toronto, Canada

Correspondence to:

Rupert M. Pearse Adult Critical Care Unit Royal London Hospital London E1 1BB

e-mail: r.pearse@qmul.ac.uk

United Kingdom

Tel: +44 20 3594 0346

Paper case record form (CRF) for International Surgical Outcomes Study

Patient name:	Date of birth: dd/mm/yyyy
International Surgical Outcomes	
For use with Outcomes	s definitions guide
Age years Gender M	F Current smoker ☐ Y ☐ N
ASA 🗌 I 🔛 II 🔲 III 🗎 IV	Black ethnicity (eGFR) 🔲 Y 🔲 N
Chronic Disease (tick all that apply):	
Coronary Artery Disease	☐ Heart Failure
☐ Diabetes Mellitus	Cirrhosis
☐ Metastatic cancer	☐ Stroke
COPD / Asthma	☐ Other
Most recent blood results (no more than 28 day	s before surgery):
	cocytes x10 ⁹ /L
	ntinineμmol/L
Souldin mino/E crea	иниеµшоу.с
Anaesthesia induction time & date:	H M M D D 0 M 2 0 1 4
Anaesthetic technique (tick all that apply)	
☐ General ☐ Spinal ☐ E	Epidural Sedation / Local
Surgical procedure category (single best answer):
☐ Orthopaedic	☐ Breast
Obstetrics & Gynaecology	Urology & Kidney
Upper gastro-intestinal	☐ Lower gastro-intestinal
☐ Hepato-biliary	☐ Vascular
☐ Head and neck	☐ Plastics / Cutaneous
☐ Cardiac	☐ Thoracic (lung & other)
☐ Thoracic (gut)	☐ Other
Severity of surgery	☐ Intermediate ☐ Major
Laparoscopic surgery	□ Y □ N
Cancer surgery	□ Y □ N
Surgical checklist used (eg WHO checklist)	□ Y □ N
Critical care immediately after surgery	□ Y □ N
Data entry staff use only	
ISOS patient Identifier:	TISOS •
,	

Patient name: Date of birth: dd/mm/yyyy

	Outcor	ne after surgery			
Infection					
Superficial surgical site	Mild 🗌	Moderate 🗌	Severe	None 🗌	
Deep surgical site	Mild 🗌	Moderate 🗌	Severe 🗌	None 🗌	
Body cavity	Mild 🗌	Moderate 🗌	Severe	None 🗌	
Pneumonia	Mild 🗌	Moderate 🗌	Severe	None 🗌	
Urinary tract	Mild 🗌	Moderate 🗌	Severe	None 🗌	
Bloodstream	Mild 🗌	Moderate 🗌	Severe 🗌	None 🗌	
Cardiovascular					
Myocardial infarction	Mild 🗌	Moderate 🗌	Severe 🗌	None 🗌	
Arrhythmia	Mild 🗌	Moderate 🗌	Severe 🗌	None 🗌	
Pulmonary oedema	Mild 🗌	Moderate 🗌	Severe 🗌	None 🗌	
Pulmonary embolism	Mild 🗌	Moderate 🗌	Severe 🗌	None 🗌	
Stroke	Mild 🗌	Moderate 🗌	Severe 🗌	None 🗌	
Cardiac arrest			Severe 🗌	None 🗌	
Other					
Gastro-intestinal bleed	Mild 🗌	Moderate 🗌	Severe 🗌	None 🗌	
Acute kidney injury	Mild 🗌	Moderate 🗌	Severe 🗌	None 🗌	
Post-operative bleed		Moderate 🗌	Severe 🗌	None 🗌	
ARDS	Mild 🗌	Moderate 🗌	Severe 🗌	None 🗌	
Anastomotic leak	Mild 🗌	Moderate 🗌	Severe 🗌	None 🗌	
Other	Mild 🗌	Moderate 🗌	Severe 🗌	None 🗌	
Treatment for post-operati	ive complicati	ions:			
Drug therapy, blood trans	sfusion or par	enteral nutrition	□ Y	■ N	
Surgical or radiological pr	ocedure			■ N	
Critical care admission			□ Y	■ N	
Hours in Post-Anaesthetic Care Unit after surgery					
Days in critical care after su		sarbery	h h		
Days in hospital after surgery					
Status at 30 days after surg	-	☐ Alive	<u> </u>	☐ Dead	
	<u></u>				
Data entry staff use only			TSC	OS 🥮	
ISOS patient Identifier:			Figure 1 and 1	Charles Study	

Supplementary table 1. Missing data in primary analysis cohort.

Data presented as n (%).

Variable	All patients (n = 44814)
Age	15 (<1)
Smoker	249 (1)
ASA score	91 (<1)
Severity of surgery	25 (<1)
Surgical procedure	9 (<1)
Ischaemic heart disease	107 (<1)
Heart failure	107 (<1)
Diabetes mellitus	102 (<1)
Cirrhosis	107 (<1)
Stroke	107 (<1)
COPD / asthma	107 (<1)
Other	107 (<1)
Patients missing at least one variable	451 (1)

Supplementary table 2. Complications after surgery.

Output of the three level hierarchical generalised mixed modelling (mixed effect logistic regression). Patients were entered in the first level, hospitals in second level and countries in the third level. Population average was used as a reference for the countries. Country was added as a random effect in the model. Odds ratios and 95% confidence intervals (CI) for each country were produced by using the random effects estimates of each country. ASA, American Society of Anesthesiologists physical status score; COPD, chronic obstructive pulmonary disease.

	Complications n (%)	Unadjusted OR (95% CI)	Adjusted OR (95% CI)	p-value
Age mean (SD)	61.8 (16.0)			
Age spline 1		1.01 (1.00 - 1.02)	1.00 (0.99 - 1.01)	0.63
Age spline 2		1.08 (1.04 - 1.12)	1.04 (0.99 - 1.08)	0.09
Age spline 3		0.77 (0.65 - 0.91)	0.88 (0.73 - 1.06)	0.18
Age spline 4		1.35 (0.99 - 1.83)	1.25 (0.89 - 1.76)	0.20
Current smoker	1305 (17.5)	0.98 (0.92 - 1.04)	1.02 (0.95 - 1.1)	0.57
ASA score				
1	848 (11.3)	Reference		
II	3005 (40.1)	1.91 (1.76 - 2.07)	1.38 (1.25 - 1.52)	<0.01
III	3090 (41.2)	5.32 (4.91 - 5.78)	2.39 (2.13 - 2.68)	<0.01
IV	554 (7.4)	14.01 (12.17 - 16.12)	4.30 (3.55 - 5.21)	<0.01
Severity of surgery				
Minor	672 (8.9)	Reference		
Intermediate	2494 (33.2)	1.62 (1.48 - 1.77)	1.76 (1.59 - 1.94)	<0.01
Major	4336 (57.8)	4.22 (3.87 - 4.60)	4.12 (3.71 - 4.56)	<0.01
Surgical procedure				
Orthopaedics	1556 (20.9)	Reference		
Breast	128 (1.7)	0.46 (0.38 - 0.55)	0.66 (0.54 - 0.81)	<0.01
Obstetrics & gynaecology	554 (7.4)	0.55 (0.49 - 0.60)	0.84 (0.75 - 0.95)	<0.01
Urology & kidney	720 (9.6)	0.87 (0.79 - 0.96)	0.98 (0.88 - 1.09)	0.67
Upper gastro-intestinal	485 (6.5)	1.63 (1.45 - 1.83)	1.59 (1.4 - 1.82)	<0.01
Lower gastro-intestinal	748 (10.0)	1.62 (1.47 - 1.79)	1.73 (1.55 - 1.93)	<0.01
Hepato-biliary	366 (4.9)	0.96 (0.85 - 1.09)	1.15 (1 - 1.32)	0.06
Vascular	410 (5.5)	1.74 (1.53 - 1.97)	1.21 (1.05 - 1.4)	0.01
Head & neck	674 (9.0)	0.58 (0.53 - 0.64)	0.77 (0.69 - 0.86)	<0.01
Plastics & cutaneous	244 (3.2)	0.86 (0.75 - 1.00)	1.15 (0.98 - 1.36)	0.09
Cardiac	979 (13.0)	6.70 (6.00 - 7.47)	2.82 (2.45 - 3.26)	<0.01
Thoracic	305 (4.1)	1.80 (1.57 - 2.08)	1.51 (1.28 - 1.78)	<0.01

Comorbidity Coronary artery disease 1525 (20.3) 2.85 (2.66 - 3.01) 1.03 (0.94 - 1.12) 0.59 Heart failure 775 (10.3) 3.76 (3.42 - 4.13) 1.24 (1.10 - 1.40) <0.01 Diabetes mellitus 1319 (17.6) 1.85 (1.73 - 1.98) 1.10 (1.02 - 1.20) 0.02 Cirrhosis 113 (1.5) 2.47 (1.97 - 3.10) 1.47 (1.12 - 1.91) 0.01 Stroke 451 (6.0) 2.22 (1.98 - 2.49) 1.13 (0.99 - 1.29) 0.07 COPD/Asthma 1012 (13.5) 1.73 (1.60 - 1.86) 1.12 (1.03 - 1.23) 0.01 Country Country AA 176 (19.9) 1.12 (0.95 - 1.32) 0.90 (0.66 - 1.23) - Country AA 176 (19.9) 1.12 (0.95 - 1.32) 0.90 (0.66 - 1.23) - Country A3 1.02 (19.3) 1.08 (0.87 - 1.33) 1.02 (0.72 - 1.43) - Country 310 (21.93) 1.08 (0.87 - 1.33) 1.02 (0.72 - 1.43) - A1 (29.3) 0.96 (0.	Other	328 (4.4)	0.56 (0.50 - 0.64)	0.79 (0.68 - 0.9)	<0.01
Heart failure 775 (10.3) 3.76 (3.42 - 4.13) 1.24 (1.10 - 1.40)	Co-morbidity				
Diabetes mellitus 1319 (17.6) 1.88 (1.73 - 1.98) 1.10 (1.02 - 1.20) 0.02 Cirrhosis 113 (1.5) 2.47 (1.97 - 3.10) 1.47 (1.12 - 1.91) 0.01 Stroke 451 (6.0) 2.22 (1.98 - 2.49) 1.13 (0.99 - 1.29) 0.07 COPD/Ashtma 1012 (13.5) 1.73 (1.60 - 1.86) 1.12 (1.03 - 1.23) 0.01 Other 4134 (55.2) 1.94 (1.85 - 2.04) 1.19 (1.12 - 1.27) <0.01	Coronary artery disease	1525 (20.3)	2.85 (2.66 - 3.01)	1.03 (0.94 - 1.12)	0.59
Cirrhosis 113 (1.5) 2.47 (1.97 - 3.10) 1.47 (1.12 - 1.91) 0.01 Stroke 451 (6.0) 2.22 (1.98 - 2.49) 1.13 (0.99 - 1.29) 0.07 COPD/Asthma 1012 (13.5) 1.73 (1.60 - 1.86) 1.12 (1.03 - 1.23) 0.01 Other 4134 (55.2) 1.94 (1.85 - 2.04) 1.19 (1.12 - 1.27) <0.01 Country	Heart failure	775 (10.3)	3.76 (3.42 - 4.13)	1.24 (1.10 - 1.40)	<0.01
Stroke 451 (6.0) 2.22 (198 - 2.49) 1.13 (0.99 - 1.29) 0.07 COPD/Asthma 1012 (13.5) 1.73 (1.60 - 1.86) 1.12 (1.03 - 1.23) 0.01 Other 4134 (55.2) 1.94 (1.85 - 2.04) 1.19 (1.12 - 1.27) <0.01 Country 2 2 1.94 (1.85 - 2.04) 1.19 (1.12 - 1.27) <0.01 AA 176 (19.9) 1.12 (0.95 - 1.32) 0.90 (0.66 - 1.23) - Z 102 (19.3) 1.08 (0.87 - 1.33) 1.02 (0.72 - 1.43) - Y 310 (31.4) 2.07 (1.81 - 2.36) 1.38 (1.01 - 1.90) - X 129 (13.3) 0.69 (0.58 - 0.83) 0.98 (0.72 - 1.34) - W 321 (21.5) 1.24 (1.09 - 1.40) 0.83 (0.62 - 1.10) - V 1060 (9.6) 0.48 (0.45 - 0.52) 0.69 (0.56 - 0.86) - U 123 (19.7) 1.11 (0.96 - 1.28) 1.08 (0.81 - 1.44) - T 174 (17.8) 0.98 (0.83 - 1.15) 1.12 (0.83 - 1.50) - S 529 (20.2) 1.14 (10.3 - 1.26) 1.18	Diabetes mellitus	1319 (17.6)	1.85 (1.73 - 1.98)	1.10 (1.02 - 1.20)	0.02
COPD/Asthma 1012 (13.5) 1.73 (1.60 - 1.86) 1.12 (1.03 - 1.23) 0.01 Other 4134 (55.2) 1.94 (1.85 - 2.04) 1.19 (1.12 - 1.27) <0.01 Country 2 1.12 (0.95 - 1.32) 0.90 (0.66 - 1.23) - Z 102 (19.3) 1.08 (0.87 - 1.33) 1.02 (0.72 - 1.43) - Y 310 (31.4) 2.07 (1.81 - 2.36) 1.38 (1.01 - 1.90) - X 129 (13.3) 0.69 (0.58 - 0.83) 0.98 (0.72 - 1.34) - W 321 (21.5) 1.24 (1.09 - 1.40) 0.83 (0.62 - 1.10) - V 1060 (9.6) 0.48 (0.45 - 0.52) 0.69 (0.56 - 0.86) - U 223 (19.7) 1.11 (0.96 - 1.28) 1.08 (0.81 - 1.44) - T 174 (17.8) 0.98 (0.83 - 1.15) 1.12 (0.83 - 1.50) - S 529 (20.2) 1.14 (1.03 - 1.26) 1.18 (0.89 - 1.57) - R 360 (20.6) 1.17 (1.04 - 1.32) 1.11 (0.88 - 1.42) - Q 192 (16.9) 0.92 (0.79 - 1.07) 0.90 (0.67 - 1.22) <th< td=""><td>Cirrhosis</td><td>113 (1.5)</td><td>2.47 (1.97 - 3.10)</td><td>1.47 (1.12 - 1.91)</td><td>0.01</td></th<>	Cirrhosis	113 (1.5)	2.47 (1.97 - 3.10)	1.47 (1.12 - 1.91)	0.01
Other 4134 (55.2) 1.94 (1.85 - 2.04) 1.19 (1.12 - 1.27) <0.01 Country 2	Stroke	451 (6.0)	2.22 (1.98 - 2.49)	1.13 (0.99 - 1.29)	0.07
Country AA 176 (19.9) 1.12 (0.95 - 1.32) 0.90 (0.66 - 1.23) - Z 102 (19.3) 1.08 (0.87 - 1.33) 1.02 (0.72 - 1.43) - Y 310 (31.4) 2.07 (1.81 - 2.36) 1.38 (1.01 - 1.90) - X 129 (13.3) 0.69 (0.58 - 0.83) 0.98 (0.72 - 1.34) - W 321 (21.5) 1.24 (1.09 - 1.40) 0.83 (0.62 - 1.10) - Y 1060 (9.6) 0.48 (0.45 - 0.52) 0.69 (0.56 - 0.86) - U 223 (19.7) 1.11 (0.96 - 1.28) 1.08 (0.81 - 1.44) - T 174 (17.8) 0.98 (0.83 - 1.15) 1.12 (0.83 - 1.50) - S 529 (20.2) 1.14 (1.03 - 1.26) 1.18 (0.89 - 1.57) - R 360 (20.6) 1.17 (1.04 - 1.32) 1.11 (0.88 - 1.42) - Q 192 (16.9) 0.92 (0.79 - 1.07) 0.90 (0.67 - 1.22) - P 70 (16.2) 0.88 (0.68 - 1.12) 1.09 (0.78 - 1.51) - O 141 (19.2) 1.07 (0.90 - 1.28) 1.02 (0.73 - 1.42) - N 79 (12.4) 0.64 (0.51 - 0.81) 1.10 (0.79 - 1.52) - M 298 (19.2) 1.07 (0.94 - 1.22) 1.01 (0.75 - 1.37) - L 265 (19.7) 1.11 (0.97 - 1.27) 1.11 (0.84 - 1.46) - K 37 (16.0) 0.86 (0.61 - 1.21) 1.10 (0.78 - 1.55) - J 168 (22.0) 1.27 (1.08 - 1.51) 1.10 (0.78 - 1.55) - I 304 (1.63) 0.88 (0.78 - 1.00) 0.94 (0.73 - 1.22) - H 125 (19.1) 1.07 (0.98 - 1.51) 1.10 (0.79 - 1.52) - H 125 (19.1) 1.07 (0.88 - 1.29) 0.94 (0.68 - 1.32) - G 39 (7.8) 0.38 (0.28 - 0.52) 0.85 (0.61 - 1.19) - E 266 (20.4) 1.16 (1.01 - 1.33) 1.07 (0.97 - 1.45) - E 163 (23.7) 1.40 (1.18 - 1.66) 1.13 (0.82 - 1.56) - D 180 (15.4) 0.82 (0.70 - 0.96) 0.92 (0.68 - 1.24) - C 588 (26.4) 1.62 (1.47 - 1.79) 1.03 (0.77 - 1.38) -	COPD/Asthma	1012 (13.5)	1.73 (1.60 - 1.86)	1.12 (1.03 - 1.23)	0.01
AA 176 (19.9) 1.12 (0.95 - 1.32) 0.90 (0.66 - 1.23) - Z 102 (19.3) 1.08 (0.87 - 1.33) 1.02 (0.72 - 1.43) - γ 310 (31.4) 2.07 (1.81 - 2.36) 1.38 (1.01 - 1.90) - χ 129 (13.3) 0.69 (0.58 - 0.83) 0.98 (0.72 - 1.34) - W 321 (21.5) 1.24 (1.09 - 1.40) 0.83 (0.62 - 1.10) - V 1060 (9.6) 0.48 (0.45 - 0.52) 0.69 (0.56 - 0.86) - U 223 (19.7) 1.11 (0.96 - 1.28) 1.08 (0.81 - 1.44) - T 174 (17.8) 0.98 (0.83 - 1.15) 1.12 (0.83 - 1.50) - S 529 (20.2) 1.14 (1.03 - 1.26) 1.18 (0.89 - 1.57) - R 360 (20.6) 1.17 (1.04 - 1.32) 1.11 (0.88 - 1.42) - Q 192 (16.9) 0.92 (0.79 - 1.07) 0.90 (0.67 - 1.22) - P 70 (16.2) 0.88 (0.68 - 1.12) 1.09 (0.78 - 1.51) - O 141 (19.2) 1.07 (0.94 - 1.28) 1.02 (0.73 - 1.42) - N 79 (12.4) 0.64 (0.51 - 0.81) 1.10 (0.75 - 1.37)	Other	4134 (55.2)	1.94 (1.85 - 2.04)	1.19 (1.12 - 1.27)	<0.01
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X 129 (13.3) 0.69 (0.58 - 0.83) 0.98 (0.72 - 1.34) - W 321 (21.5) 1.24 (1.09 - 1.40) 0.83 (0.62 - 1.10) - V 1060 (9.6) 0.48 (0.45 - 0.52) 0.69 (0.56 - 0.86) - U 223 (19.7) 1.11 (0.96 - 1.28) 1.08 (0.81 - 1.44) - T 174 (17.8) 0.98 (0.83 - 1.15) 1.12 (0.83 - 1.50) - S 529 (20.2) 1.14 (1.03 - 1.26) 1.18 (0.89 - 1.57) - R 360 (20.6) 1.17 (1.04 - 1.32) 1.11 (0.88 - 1.42) - Q 192 (16.9) 0.92 (0.79 - 1.07) 0.90 (0.67 - 1.22) - P 70 (16.2) 0.88 (0.68 - 1.12) 1.09 (0.78 - 1.51) - O 141 (19.2) 1.07 (0.90 - 1.28) 1.02 (0.73 - 1.42) - N 79 (12.4) 0.64 (0.51 - 0.81) 1.10 (0.79 - 1.52) - M 298 (19.2) 1.07 (0.94 - 1.22) 1.01 (0.75 - 1.37) - L 265 (19.7) 1.11 (0.97 - 1.27) 1.11 (0.84 - 1.46) - K 37 (16.0) 0.86 (0.61 - 1.21) 1.10 (0.78 - 1.55)	Z	102 (19.3)	1.08 (0.87 - 1.33)	1.02 (0.72 - 1.43)	-
W 321 (21.5) 1.24 (1.09-1.40) 0.83 (0.62-1.10) - V 1060 (9.6) 0.48 (0.45-0.52) 0.69 (0.56-0.86) - U 223 (19.7) 1.11 (0.96-1.28) 1.08 (0.81-1.44) - T 174 (17.8) 0.98 (0.83-1.15) 1.12 (0.83-1.50) - S 529 (20.2) 1.14 (1.03-1.26) 1.18 (0.89-1.57) - R 360 (20.6) 1.17 (1.04-1.32) 1.11 (0.88-1.42) - Q 192 (16.9) 0.92 (0.79-1.07) 0.90 (0.67-1.22) - P 70 (16.2) 0.88 (0.68-1.12) 1.09 (0.78-1.51) - O 141 (19.2) 1.07 (0.90-1.28) 1.02 (0.73-1.42) - N 79 (12.4) 0.64 (0.51-0.81) 1.10 (0.79-1.52) - M 298 (19.2) 1.07 (0.94-1.22) 1.01 (0.75-1.37) - L 265 (19.7) 1.11 (0.97-1.27) 1.11 (0.84-1.46) - K 37 (16.0) 0.86 (0.61-1.21) 1.10 (0.78-1.55) - J 168 (22.0) 1.27 (1.08-1.51) 1.10 (0.80-1.50) - I <t< td=""><td>Υ</td><td>310 (31.4)</td><td>2.07 (1.81 - 2.36)</td><td>1.38 (1.01 - 1.90)</td><td>-</td></t<>	Υ	310 (31.4)	2.07 (1.81 - 2.36)	1.38 (1.01 - 1.90)	-
V 1060 (9.6) 0.48 (0.45 - 0.52) 0.69 (0.56 - 0.86) - U 223 (19.7) 1.11 (0.96 - 1.28) 1.08 (0.81 - 1.44) - T 174 (17.8) 0.98 (0.83 - 1.15) 1.12 (0.83 - 1.50) - S 529 (20.2) 1.14 (1.03 - 1.26) 1.18 (0.89 - 1.57) - R 360 (20.6) 1.17 (1.04 - 1.32) 1.11 (0.88 - 1.42) - Q 192 (16.9) 0.92 (0.79 - 1.07) 0.90 (0.67 - 1.22) - P 70 (16.2) 0.88 (0.68 - 1.12) 1.09 (0.78 - 1.51) - O 141 (19.2) 1.07 (0.90 - 1.28) 1.02 (0.73 - 1.42) - N 79 (12.4) 0.64 (0.51 - 0.81) 1.10 (0.79 - 1.52) - M 298 (19.2) 1.07 (0.94 - 1.22) 1.01 (0.75 - 1.37) - L 265 (19.7) 1.11 (0.97 - 1.27) 1.11 (0.84 - 1.46) - K 37 (16.0) 0.86 (0.61 - 1.21) 1.10 (0.78 - 1.55) - J 168 (22.0) 1.27 (1.08 - 1.51) 1.10 (0.80 - 1.50) - I 304 (16.3) 0.88 (0.78 - 1.00) 0.94 (0.68 - 1.32)	X	129 (13.3)	0.69 (0.58 - 0.83)	0.98 (0.72 - 1.34)	-
U 223 (19.7) 1.11 (0.96 - 1.28) 1.08 (0.81 - 1.44) - T 174 (17.8) 0.98 (0.83 - 1.15) 1.12 (0.83 - 1.50) - S 529 (20.2) 1.14 (1.03 - 1.26) 1.18 (0.89 - 1.57) - R 360 (20.6) 1.17 (1.04 - 1.32) 1.11 (0.88 - 1.42) - Q 192 (16.9) 0.92 (0.79 - 1.07) 0.90 (0.67 - 1.22) - P 70 (16.2) 0.88 (0.68 - 1.12) 1.09 (0.78 - 1.51) - O 141 (19.2) 1.07 (0.90 - 1.28) 1.02 (0.73 - 1.42) - N 79 (12.4) 0.64 (0.51 - 0.81) 1.10 (0.79 - 1.52) - M 298 (19.2) 1.07 (0.94 - 1.22) 1.01 (0.75 - 1.37) - L 265 (19.7) 1.11 (0.97 - 1.27) 1.11 (0.84 - 1.46) - K 37 (16.0) 0.86 (0.61 - 1.21) 1.10 (0.78 - 1.55) - J 168 (22.0) 1.27 (1.08 - 1.51) 1.10 (0.80 - 1.50) - I 304 (16.3) 0.88 (0.78 - 1.00) 0.94 (0.68 - 1.32) - G 39 (7.8) 0.38 (0.28 - 0.52) 0.85 (0.61 - 1.19) <	W	321 (21.5)	1.24 (1.09 - 1.40)	0.83 (0.62 - 1.10)	-
T	V	1060 (9.6)	0.48 (0.45 - 0.52)	0.69 (0.56 - 0.86)	-
S 529 (20.2) 1.14 (1.03 - 1.26) 1.18 (0.89 - 1.57) - R 360 (20.6) 1.17 (1.04 - 1.32) 1.11 (0.88 - 1.42) - Q 192 (16.9) 0.92 (0.79 - 1.07) 0.90 (0.67 - 1.22) - P 70 (16.2) 0.88 (0.68 - 1.12) 1.09 (0.78 - 1.51) - O 141 (19.2) 1.07 (0.90 - 1.28) 1.02 (0.73 - 1.42) - N 79 (12.4) 0.64 (0.51 - 0.81) 1.10 (0.79 - 1.52) - M 298 (19.2) 1.07 (0.94 - 1.22) 1.01 (0.75 - 1.37) - L 265 (19.7) 1.11 (0.97 - 1.27) 1.11 (0.84 - 1.46) - K 37 (16.0) 0.86 (0.61 - 1.21) 1.10 (0.78 - 1.55) - J 168 (22.0) 1.27 (1.08 - 1.51) 1.10 (0.80 - 1.50) - I 304 (16.3) 0.88 (0.78 - 1.00) 0.94 (0.73 - 1.22) - H 125 (19.1) 1.07 (0.88 - 1.29) 0.94 (0.68 - 1.32) - G 39 (7.8) 0.38 (0.28 - 0.52) 0.85 (0.61 - 1.19) - F 266 (20.4) 1.16 (1.01 - 1.33) 1.07 (0.79 - 1.45) <t< td=""><td>U</td><td>223 (19.7)</td><td>1.11 (0.96 - 1.28)</td><td>1.08 (0.81 - 1.44)</td><td>-</td></t<>	U	223 (19.7)	1.11 (0.96 - 1.28)	1.08 (0.81 - 1.44)	-
R 360 (20.6) 1.17 (1.04 - 1.32) 1.11 (0.88 - 1.42) - Q 192 (16.9) 0.92 (0.79 - 1.07) 0.90 (0.67 - 1.22) - P 70 (16.2) 0.88 (0.68 - 1.12) 1.09 (0.78 - 1.51) - O 141 (19.2) 1.07 (0.90 - 1.28) 1.02 (0.73 - 1.42) - N 79 (12.4) 0.64 (0.51 - 0.81) 1.10 (0.79 - 1.52) - M 298 (19.2) 1.07 (0.94 - 1.22) 1.01 (0.75 - 1.37) - L 265 (19.7) 1.11 (0.97 - 1.27) 1.11 (0.84 - 1.46) - K 37 (16.0) 0.86 (0.61 - 1.21) 1.10 (0.78 - 1.55) - J 168 (22.0) 1.27 (1.08 - 1.51) 1.10 (0.80 - 1.50) - I 304 (16.3) 0.88 (0.78 - 1.00) 0.94 (0.73 - 1.22) - H 125 (19.1) 1.07 (0.88 - 1.29) 0.94 (0.68 - 1.32) - G 39 (7.8) 0.38 (0.28 - 0.52) 0.85 (0.61 - 1.19) - F 266 (20.4) 1.16 (1.01 - 1.33) 1.07 (0.79 - 1.45) - D 180 (15.4) 0.82 (0.70 - 0.96) 0.92 (0.68 - 1.24) - C 589 (26.4) 1.62 (1.47 - 1.79) 1.03 (0.77 - 1.38) - B 42 (23.0) 1.35 (0.96 - 1.88) 1.22 (0.86 - 1.72) -	Т	174 (17.8)	0.98 (0.83 - 1.15)	1.12 (0.83 - 1.50)	-
Q 192 (16.9) 0.92 (0.79 - 1.07) 0.90 (0.67 - 1.22) - P 70 (16.2) 0.88 (0.68 - 1.12) 1.09 (0.78 - 1.51) - O 141 (19.2) 1.07 (0.90 - 1.28) 1.02 (0.73 - 1.42) - P 79 (12.4) 0.64 (0.51 - 0.81) 1.10 (0.79 - 1.52) - D 128 (19.2) 1.07 (0.94 - 1.22) 1.01 (0.75 - 1.37) - D 128 (19.2) 1.07 (0.94 - 1.22) 1.01 (0.75 - 1.37) - D 129 (19.2) 1.01 (0.78 - 1.55) 1.11 (0.84 - 1.46) 1.12 (19.1) 1.12 (1.84 - 1.46) 1.13 (1.84 - 1.46) 1.14 (1.97 - 1.27) 1.14 (1.984 - 1.46) 1.15 (19.1) 1.10 (0.78 - 1.55) 1.10 (0.78 - 1.55) 1.10 (0.78 - 1.55) 1.10 (0.78 - 1.55) 1.10 (0.78 - 1.50)	S	529 (20.2)	1.14 (1.03 - 1.26)	1.18 (0.89 - 1.57)	-
P 70 (16.2) 0.88 (0.68 - 1.12) 1.09 (0.78 - 1.51) - O 141 (19.2) 1.07 (0.90 - 1.28) 1.02 (0.73 - 1.42) - N 79 (12.4) 0.64 (0.51 - 0.81) 1.10 (0.79 - 1.52) - M 298 (19.2) 1.07 (0.94 - 1.22) 1.01 (0.75 - 1.37) - L 265 (19.7) 1.11 (0.97 - 1.27) 1.11 (0.84 - 1.46) - K 37 (16.0) 0.86 (0.61 - 1.21) 1.10 (0.78 - 1.55) - J 168 (22.0) 1.27 (1.08 - 1.51) 1.10 (0.80 - 1.50) - I 304 (16.3) 0.88 (0.78 - 1.00) 0.94 (0.73 - 1.22) - H 125 (19.1) 1.07 (0.88 - 1.29) 0.94 (0.68 - 1.32) - G 39 (7.8) 0.38 (0.28 - 0.52) 0.85 (0.61 - 1.19) - F 266 (20.4) 1.16 (1.01 - 1.33) 1.07 (0.79 - 1.45) - E 163 (23.7) 1.40 (1.18 - 1.66) 1.13 (0.82 - 1.56) - D 180 (15.4) 0.82 (0.70 - 0.96) 0.92 (0.68 - 1.24) - C 589 (26.4) 1.62 (1.47 - 1.79) 1.03 (0.77 - 1.38) -	R	360 (20.6)	1.17 (1.04 - 1.32)	1.11 (0.88 - 1.42)	-
O 141 (19.2) 1.07 (0.90 - 1.28) 1.02 (0.73 - 1.42) - N 79 (12.4) 0.64 (0.51 - 0.81) 1.10 (0.79 - 1.52) - M 298 (19.2) 1.07 (0.94 - 1.22) 1.01 (0.75 - 1.37) - L 265 (19.7) 1.11 (0.97 - 1.27) 1.11 (0.84 - 1.46) - K 37 (16.0) 0.86 (0.61 - 1.21) 1.10 (0.78 - 1.55) - J 168 (22.0) 1.27 (1.08 - 1.51) 1.10 (0.80 - 1.50) - I 304 (16.3) 0.88 (0.78 - 1.00) 0.94 (0.73 - 1.22) - H 125 (19.1) 1.07 (0.88 - 1.29) 0.94 (0.68 - 1.32) - G 39 (7.8) 0.38 (0.28 - 0.52) 0.85 (0.61 - 1.19) - F 266 (20.4) 1.16 (1.01 - 1.33) 1.07 (0.79 - 1.45) - E 163 (23.7) 1.40 (1.18 - 1.66) 1.13 (0.82 - 1.56) - D 180 (15.4) 0.82 (0.70 - 0.96) 0.92 (0.68 - 1.24) - C 589 (26.4) 1.62 (1.47 - 1.79) 1.03 (0.77 - 1.38) - B 42 (23.0) 1.35 (0.96 - 1.88) 1.22 (0.86 - 1.72) <t< td=""><td>Q</td><td>192 (16.9)</td><td>0.92 (0.79 - 1.07)</td><td>0.90 (0.67 - 1.22)</td><td>-</td></t<>	Q	192 (16.9)	0.92 (0.79 - 1.07)	0.90 (0.67 - 1.22)	-
N 79 (12.4) 0.64 (0.51 - 0.81) 1.10 (0.79 - 1.52) - M 298 (19.2) 1.07 (0.94 - 1.22) 1.01 (0.75 - 1.37) - L 265 (19.7) 1.11 (0.97 - 1.27) 1.11 (0.84 - 1.46) - K 37 (16.0) 0.86 (0.61 - 1.21) 1.10 (0.78 - 1.55) - J 168 (22.0) 1.27 (1.08 - 1.51) 1.10 (0.80 - 1.50) - I 304 (16.3) 0.88 (0.78 - 1.00) 0.94 (0.73 - 1.22) - H 125 (19.1) 1.07 (0.88 - 1.29) 0.94 (0.68 - 1.32) - G 39 (7.8) 0.38 (0.28 - 0.52) 0.85 (0.61 - 1.19) - F 266 (20.4) 1.16 (1.01 - 1.33) 1.07 (0.79 - 1.45) - E 163 (23.7) 1.40 (1.18 - 1.66) 1.13 (0.82 - 1.56) - D 180 (15.4) 0.82 (0.70 - 0.96) 0.92 (0.68 - 1.24) - C 589 (26.4) 1.62 (1.47 - 1.79) 1.03 (0.77 - 1.38) - B 42 (23.0) 1.35 (0.96 - 1.88) 1.22 (0.86 - 1.72) -	Р	70 (16.2)	0.88 (0.68 - 1.12)	1.09 (0.78 - 1.51)	-
M 298 (19.2) 1.07 (0.94 - 1.22) 1.01 (0.75 - 1.37) - L 265 (19.7) 1.11 (0.97 - 1.27) 1.11 (0.84 - 1.46) - K 37 (16.0) 0.86 (0.61 - 1.21) 1.10 (0.78 - 1.55) - J 168 (22.0) 1.27 (1.08 - 1.51) 1.10 (0.80- 1.50) - I 304 (16.3) 0.88 (0.78 - 1.00) 0.94 (0.73 - 1.22) - H 125 (19.1) 1.07 (0.88 - 1.29) 0.94 (0.68 - 1.32) - G 39 (7.8) 0.38 (0.28 - 0.52) 0.85 (0.61 - 1.19) - F 266 (20.4) 1.16 (1.01 - 1.33) 1.07 (0.79 - 1.45) - E 163 (23.7) 1.40 (1.18 - 1.66) 1.13 (0.82 - 1.56) - D 180 (15.4) 0.82 (0.70 - 0.96) 0.92 (0.68 - 1.24) - C 589 (26.4) 1.62 (1.47 - 1.79) 1.03 (0.77 - 1.38) - B 42 (23.0) 1.35 (0.96 - 1.88) 1.22 (0.86 - 1.72) -	0	141 (19.2)	1.07 (0.90 - 1.28)	1.02 (0.73 - 1.42)	-
L 265 (19.7) 1.11 (0.97 - 1.27) 1.11 (0.84 - 1.46) - K 37 (16.0) 0.86 (0.61 - 1.21) 1.10 (0.78 - 1.55) - J 168 (22.0) 1.27 (1.08 - 1.51) 1.10 (0.80 - 1.50) - J 304 (16.3) 0.88 (0.78 - 1.00) 0.94 (0.73 - 1.22) - H 125 (19.1) 1.07 (0.88 - 1.29) 0.94 (0.68 - 1.32) - G 39 (7.8) 0.38 (0.28 - 0.52) 0.85 (0.61 - 1.19) - F 266 (20.4) 1.16 (1.01 - 1.33) 1.07 (0.79 - 1.45) - E 163 (23.7) 1.40 (1.18 - 1.66) 1.13 (0.82 - 1.56) - D 180 (15.4) 0.82 (0.70 - 0.96) 0.92 (0.68 - 1.24) - C 589 (26.4) 1.62 (1.47 - 1.79) 1.03 (0.77 - 1.38) - B 42 (23.0) 1.35 (0.96 - 1.88) 1.22 (0.86 - 1.72) -	N	79 (12.4)	0.64 (0.51 - 0.81)	1.10 (0.79 - 1.52)	-
K 37 (16.0) 0.86 (0.61 - 1.21) 1.10 (0.78 - 1.55) - J 168 (22.0) 1.27 (1.08 - 1.51) 1.10 (0.80 - 1.50) - I 304 (16.3) 0.88 (0.78 - 1.00) 0.94 (0.73 - 1.22) - H 125 (19.1) 1.07 (0.88 - 1.29) 0.94 (0.68 - 1.32) - G 39 (7.8) 0.38 (0.28 - 0.52) 0.85 (0.61 - 1.19) - F 266 (20.4) 1.16 (1.01 - 1.33) 1.07 (0.79 - 1.45) - E 163 (23.7) 1.40 (1.18 - 1.66) 1.13 (0.82 - 1.56) - D 180 (15.4) 0.82 (0.70 - 0.96) 0.92 (0.68 - 1.24) - C 589 (26.4) 1.62 (1.47 - 1.79) 1.03 (0.77 - 1.38) - B 42 (23.0) 1.35 (0.96 - 1.88) 1.22 (0.86 - 1.72) -	М	298 (19.2)	1.07 (0.94 - 1.22)	1.01 (0.75 - 1.37)	-
J 168 (22.0) 1.27 (1.08 - 1.51) 1.10 (0.80 - 1.50) - I 304 (16.3) 0.88 (0.78 - 1.00) 0.94 (0.73 - 1.22) - H 125 (19.1) 1.07 (0.88 - 1.29) 0.94 (0.68 - 1.32) - G 39 (7.8) 0.38 (0.28 - 0.52) 0.85 (0.61 - 1.19) - F 266 (20.4) 1.16 (1.01 - 1.33) 1.07 (0.79 - 1.45) - E 163 (23.7) 1.40 (1.18 - 1.66) 1.13 (0.82 - 1.56) - D 180 (15.4) 0.82 (0.70 - 0.96) 0.92 (0.68 - 1.24) - C 589 (26.4) 1.62 (1.47 - 1.79) 1.03 (0.77 - 1.38) - B 42 (23.0) 1.35 (0.96 - 1.88) 1.22 (0.86 - 1.72) -	L	265 (19.7)	1.11 (0.97 - 1.27)	1.11 (0.84 - 1.46)	-
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G 39 (7.8) 0.38 (0.28 - 0.52) 0.85 (0.61 - 1.19) - E 266 (20.4) 1.16 (1.01 - 1.33) 1.07 (0.79 - 1.45) - E 163 (23.7) 1.40 (1.18 - 1.66) 1.13 (0.82 - 1.56) - D 180 (15.4) 0.82 (0.70 - 0.96) 0.92 (0.68 - 1.24) - C 589 (26.4) 1.62 (1.47 - 1.79) 1.03 (0.77 - 1.38) - B 42 (23.0) 1.35 (0.96 - 1.88) 1.22 (0.86 - 1.72) -	I	304 (16.3)	0.88 (0.78 - 1.00)	0.94 (0.73 - 1.22)	-
E 266 (20.4) 1.16 (1.01 - 1.33) 1.07 (0.79 - 1.45) - E 163 (23.7) 1.40 (1.18 - 1.66) 1.13 (0.82 - 1.56) - D 180 (15.4) 0.82 (0.70 - 0.96) 0.92 (0.68 - 1.24) - C 589 (26.4) 1.62 (1.47 - 1.79) 1.03 (0.77 - 1.38) - B 42 (23.0) 1.35 (0.96 - 1.88) 1.22 (0.86 - 1.72) -	Н	125 (19.1)	1.07 (0.88 - 1.29)	0.94 (0.68 - 1.32)	-
E 163 (23.7) 1.40 (1.18 - 1.66) 1.13 (0.82 - 1.56) - D 180 (15.4) 0.82 (0.70 - 0.96) 0.92 (0.68 - 1.24) - C 589 (26.4) 1.62 (1.47 - 1.79) 1.03 (0.77 - 1.38) - B 42 (23.0) 1.35 (0.96 - 1.88) 1.22 (0.86 - 1.72) -	G	39 (7.8)	0.38 (0.28 - 0.52)	0.85 (0.61 - 1.19)	-
D 180 (15.4) 0.82 (0.70 - 0.96) 0.92 (0.68 - 1.24) - C 589 (26.4) 1.62 (1.47 - 1.79) 1.03 (0.77 - 1.38) - B 42 (23.0) 1.35 (0.96 - 1.88) 1.22 (0.86 - 1.72) -	F	266 (20.4)	1.16 (1.01 - 1.33)	1.07 (0.79 - 1.45)	-
C 589 (26.4) 1.62 (1.47 - 1.79) 1.03 (0.77 - 1.38) - 42 (23.0) 1.35 (0.96 - 1.88) 1.22 (0.86 - 1.72) -	E	163 (23.7)	1.40 (1.18 - 1.66)	1.13 (0.82 - 1.56)	-
B 42 (23.0) 1.35 (0.96 - 1.88) 1.22 (0.86 - 1.72) -	D	180 (15.4)	0.82 (0.70 - 0.96)	0.92 (0.68 - 1.24)	-
1166 (16.6) 0.0 (0.94, 0.06) 0.92 (0.72, 0.06)	С	589 (26.4)	1.62 (1.47 - 1.79)	1.03 (0.77 - 1.38)	-
A 1166 (16.6) 0.9 (0.84 - 0.96) 0.83 (0.72 - 0.96) -	В	42 (23.0)	1.35 (0.96 - 1.88)	1.22 (0.86 - 1.72)	-
	Α	1166 (16.6)	0.9 (0.84 - 0.96)	0.83 (0.72 - 0.96)	=

Supplementary table 3. Mortality after surgery.

Output of the three level hierarchical generalised mixed modelling (mixed effect logistic regression). Patients were entered in the first level, hospitals in second level and countries in the third level. Population average was used as a reference for countries. Country was added as a random effect in the model, Odds ratios and 95% confidence intervals (CI) for each country were produced by using the random effects estimates for each country. ASA, American Society of Anesthesiologists physical status score; COPD, chronic obstructive pulmonary disease.

	In-hospital mortality n (%)	Unadjusted OR Adjusted OR (95% CI) (95% CI)		p-value	
Age mean (SD)	69.1 (13.3)				
Age spline 1		1.15 (1.00 - 1.33)	1.12 (0.97 - 1.30)	0.12	
Age spline 2		0.83 (0.56 - 1.24)	0.79 (0.53 - 1.18)	0.25	
Age spline 3		1.57 (0.36 - 6.86)	1.94 (0.43 - 8.68)	0.39	
Age spline 4		0.91 (0.11 - 7.61)	0.75 (0.08 - 6.67)	0.80	
Current smoker	121 (58.5)	1.37 (0.99 - 1.90)	1.57 (1.09 - 2.25)	0.02	
ASA score					
1	1 (0.5)	Reference			
II	38 (18.4)	19.19 (2.63 - 139.8)	9.23 (1.24 - 68.50)	0.03	
III	115 (55.6)	128.10 (17.89 - 917.32)	32.79 (4.41 - 243.84)	<0.01	
IV	53 (25.6)	604.04 (83.45 - 4372.45)	111.98 (14.56 - 861.14)	<0.01	
Severity of surgery					
Minor	14 (6.8)	Reference			
Intermediate	56 (27.1)	1.67 (0.93 - 3.00)	1.44 (0.78 - 2.66)	0.24	
Major	137 (66.2)	5.12 (2.95 - 8.89)	3.01 (1.65 - 5.47)	<0.01	
Surgical procedure					
Orthopaedic	25 (12.1)	Reference			
Breast	2 (1.0)	0.49 (0.12 - 2.08)	1.14 (0.26 - 4.95)	0.86	
Obstetrics & gynaecology	6 (2.9)	0.40 (0.16 - 0.97)	1.38 (0.55 - 3.47)	0.50	
Urology & kidney	10 (4.8)	0.78 (0.37 - 1.62)	0.87 (0.41 - 1.84)	0.71	
Upper gastro-intestinal	29 (14.0)	5.59 (3.27 - 9.57)	4.92 (2.77 - 8.74)	<0.01	
Lower gastro-intestinal	32 (15.5)	3.97 (2.35 - 6.71)	3.66 (2.10 - 6.39)	<0.01	
Hepato-biliary	14 (6.8)	2.33 (1.21 - 4.49)	3.04 (1.52 - 6.10)	<0.01	
Vascular	15 (7.2)	3.57 (1.88 - 6.79)	1.54 (0.78 - 3.02)	0.21	
Head & neck	12 (5.8)	0.70 (0.35 - 1.39)	1.16 (0.57 - 2.39)	0.68	
Plastics & cutaneous	5 (2.4)	1.13 (0.43 - 2.96)	1.51 (0.55 - 4.16)	0.42	
Cardiac	40 (19.3)	9.01 (5.45 - 14.88)	2.36 (1.32 - 4.23)	<0.01	
Thoracic	10 (4.8)	3.29 (1.58 - 6.87)	2.43 (1.11 - 5.35)	0.03	

Other	7 (3.4)	0.81 (0.35 - 1.87)	1.36 (0.57 - 3.26)	0.49
Co-morbidities				
Coronary artery disease	67 (32.4)	4.23 (3.16 - 5.67)	0.99 (0.70 - 1.41)	0.97
Congestive heart failure	49 (23.7)	7.22 (5.22 - 9.98)	1.41 (0.96 - 2.07)	0.08
Diabetes mellitus	58 (28.0)	3.00 (2.21 - 4.07)	1.24 (0.89 - 1.73)	0.20
Cirrhosis	10 (4.8)	6.75 (3.55 - 12.86)	2.61 (1.27 - 5.33)	0.01
Stroke	38 (18.4)	6.66 (4.66 - 9.50)	2.58 (1.74 - 3.82)	<0.01
COPD/Asthma	40 (19.3)	2.39 (1.69 - 3.38)	1.14 (0.79 - 1.67)	0.48
Other	134 (64.7)	2.59 (1.94 - 3.44)	1.30 (0.95 - 1.78)	0.11
Country				
AA	2 (0.2)	0.46 (0.12 - 1.78)	0.57 (0.25 - 1.29)	-
Z	7 (1.3)	2.74 (1.30 - 5.77)	1.76 (0.81 - 3.85)	-
Υ	4 (0.4)	0.83 (0.32 - 2.18)	0.82 (0.37 - 1.83)	-
Х	17 (1.7)	3.63 (2.20 - 6.01)	3.95 (2.12 - 7.36)	-
W	6 (0.4)	0.83 (0.37 - 1.83)	0.57 (0.29 - 1.12)	-
V	12 (0.1)	0.22 (0.12 - 0.40)	0.64 (0.37 - 1.11)	-
U	7 (0.6)	1.27 (0.60 - 2.67)	1.29 (0.63 - 2.65)	-
Т	6 (0.6)	1.26 (0.57 - 2.79)	1.43 (0.68 - 3.02)	-
S	11 (0.4)	0.86 (0.47 - 1.57)	0.93 (0.5 - 1.75)	-
R	15 (0.9)	1.77 (1.04 - 3.00)	1.29 (0.74 - 2.26)	-
Q	3 (0.3)	0.54 (0.18 - 1.64)	0.75 (0.34 - 1.68)	-
P	1 (0.2)	0.47 (0.07 - 3.17)	1.10 (0.38 - 3.13)	-
0	5 (0.7)	1.40 (0.59 - 3.34)	1.19 (0.53 - 2.68)	-
N	2 (0.3)	0.64 (0.17 - 2.49)	1.48 (0.53 - 4.13)	-
М	4 (0.3)	0.53 (0.20 - 1.38)	0.76 (0.35 - 1.62)	-
L	2 (0.1)	0.30 (0.08 - 1.17)	0.58 (0.26 - 1.33)	-
К	2 (0.9)	1.78 (0.46 - 6.91)	1.67 (0.57 - 4.92)	-
J	5 (0.7)	1.35 (0.56 - 3.21)	1.12 (0.52 - 2.43)	-
ſ	20 (1.1)	2.22 (1.39 - 3.55)	1.65 (0.98 - 2.78)	-
Н	5 (0.8)	1.57 (0.66 - 3.76)	1.17 (0.52 - 2.61)	-
G	3 (0.6)	1.23 (0.40 - 3.72)	1.78 (0.67 - 4.73)	-
F	14 (1.1)	2.22 (1.29 - 3.83)	1.79 (0.97 - 3.29)	-
E	2 (0.3)	0.59 (0.15 - 2.29)	0.91 (0.37 - 2.24)	-
D	4 (0.3)	0.70 (0.27 - 1.84)	0.95 (0.43 - 2.09)	-
С	24 (1.1)	2.22 (1.44 - 3.44)	0.94 (0.55 - 1.60)	-
В	1 (0.5)	1.12 (0.17 - 7.52)	1.38 (0.44 - 4.36)	-
A	23 (0.3)	0.67 (0.43 - 1.04)	0.87 (0.58 - 1.31)	-

Supplementary table 4. Complications and mortality after surgery including all patients entered onto the International Surgical Outcomes Study database. Data presented as n (%). ARDS, acute respiratory distress syndrome; N/A, category not applicable for this complication. Some patients may have developed more than one complication, and consequently in some cases the denominator is the number complications whilst in the left most column the denominator is the number of patients. The cell at the bottom of the far right column represents the number of deaths divided by the number of patients with at least one complication.

		Severity of complications			Mortality for patients who developed complications
	n = 45,599	Mild	Moderate	Severe	n = 212
Infectious complications					
Superficial surgical site	1366 (3.0)	711/1366 (52.0)	527/1366 (38.6)	128/1366 (9.4)	19/1366 (1.4)
Deep surgical site	585 (1.28)	127/585 (21.7)	256/585 (43.8)	202/585 (34.5)	29/585 (5.0)
Body cavity	348 (0.8)	99/348 (28.4)	138/348 (39.7)	111/348 (31.9)	26/348 (7.5)
Pneumonia	732 (1.6)	246/732 (33.6)	335/732 (45.8)	151/732 (20.6)	57/732 (7.8)
Urinary tract	696 (1.5)	299/696 (43.0)	342/696 (49.1)	55/696 (7.9)	13/696 (1.9)
Bloodstream	432 (0.9)	143/432 (33.1)	169/432 (39.1)	120/432 (27.8)	50/432 (11.6)
Total infectious complications	4159	1625/4159 (39.1)	1767/4159 (42.5)	767/4159 (18.4)	110/4159 (2.6)
Cardiovascular complications					
Myocardial infarction	145 (0.3)	48/145 (33.1)	45/145 (31.0)	52/145 (35.9)	26/145 (17.9)
Arrhythmia	1238 (2.7)	477/1238 (38.5)	571/1238 (46.1)	190/1238 (15.3)	76/1238 (6.1)
Pulmonary oedema	337 (0.7)	131/337 (38.9)	142/337 (42.1)	64/337 (19.0)	33/337 (9.8)
Pulmonary embolism	83 (0.2)	17/83 (20.5)	36/83 (43.4)	30/83 (36.1)	6/83 (7.2)

Stroke	117 (0.3)	33/117 (28.2)	30/117 (25.6)	54/117 (46.2)	18/117 (15.4)
Cardiac arrest	160 (0.4)	N/A	N/A	160/160 (100)	94/160 (58.8)
Total cardiovascular complications	2080	706/2080 (33.9)	824/2080 (39.7)	550/2080 (26.4)	147/2080 (7.1)
Other complications					
Gastro-intestinal bleed	206 (0.5)	98/206 (47.6)	68/206 (33.0)	40/206 (19.4)	25/206 (12.1)
Acute kidney injury	798 (1.8)	433/798 (54.3)	209/798 (26.2)	156/798 (19.5)	80/798 (10.0)
Post-operative bleed	1377 (3.0)	N/A	1162/1377 (84.4)	215/1377 (15.6)	54/1377 (3.9)
ARDS	147 (0.3)	50/147 (34.0)	41/147 (27.9)	56/147 (38.1)	34/147 (23.1)
Anastomotic leak	212 (0.5)	52/212 (24.5)	64/212 (30.2)	96/212 (45.3)	21/212 (9.9)
All others	2978 (6.5)	1371/2978 (46.0)	1221/2978 (41.0)	386/2978 (13.0)	85/2978 (2.9)
Total other complications	5718	2004/5718 (35.0)	2765/5718 (48.4)	949/5718 (16.6)	163/5718 (2.9)
Total number of complications	11957	4335/11957 (36.3)	5356/11957 (44.8)	2266/11957 (18.9)	212/7673 (2.8)

Members of the ISOS group

Steering committee

Rupert M. Pearse (chief investigator)

Scott Beattie

Pierre-Alain Clavien

Nicolas Demartines

Lee A. Fleisher

Mike Grocott (independent, chair)

James Haddow

Andreas Hoeft

Peter Holt

Rui Moreno

Naomi Pritchard (independent, patient representative)

Andrew Rhodes

Duminda Wijeysundera

Matt Wilson (independent)

Writing committee

Rupert M. Pearse (chair)

Tahania Ahmad

Xiangming Fang

Peter Holt

Brennan C. Kahan

Andrew Rhodes

Duminda Wijeysundera

Study management group

Tahania Ahmed (statistician)

Kirsty Everingham

Russell Hewson (study websites and information governance)

Marta Januszewska (study coordinator)

Rupert M. Pearse (chief Investigator)

Mandeep-Kaur Phull

National Co-ordinators

Australia: Richard Halliwell, Mark Shulman and Paul Myles

Austria: Werner Schmid and Michael Hiesmayr

Belgium: Patrick Wouters and Stefan de Hert

Brazil: Suzana Lobo

Canada: Scott Beattie and Duminda Wijeysundera

China: Xiangming Fang

Denmark: Lars Rasmussen

France: Emmanuel Futier, Matthieu Biais, Aurélien Venara and Karem Slim

Germany: Michael Sander

Greece: Despoina Koulenti and Kostoula Arvaniti

Hong Kong: Mathew Chan

India: Atul Kulkarni

Indonesia: Susilo Chandra and Aida Tantri
Iraq: Emad Geddoa and Muntadhar Abbas

Italy: Giorgio Della Rocca

Malaysia: Datin Sivasakthi and Marzida Mansor

Mexico: Pastor Luna

Netherlands: Arthur Bouwman and Wolfgang Buhre

New Zealand: Vanessa Beavis, Douglas Campbell and Tim Short

Nigeria: Tunde Osinaike **Portugal:** Ricardo Matos

Romania: Ioana Grigoras

Russia: Mikhail Kirov and Denis Protsenko

South Africa: Bruce Biccard

Spain: Cesar Aldecoa

Sweden: Michelle Chew

Switzerland: Christoph Hofer and Martin Hubner

Uganda: James Ditai

United Kingdom: Tamas Szakmany

United States of America: Lee Fleisher

Local investigators

*local co-ordinator

Australia

Alfred Hospital

Marissa Ferguson, Michael MacMahon, Mark Shulman*

Blacktown Hospital

Ritchie Cherian, Helen Currow*, Kathirgamanathan Kanathiban,

Coffs Harbour Base Hospital

David Gillespie*, Edward Pathmanathan, Katherine Phillips, Jenifer Reynolds*, Joanne Rowley,

John Hunter Hospital

Jeanene Douglas, Ross Kerridge*

Mount Druitt Hospital

Helen Currow*, Sameer Garg

Prince of Wales Hospital

Michael Bennett*, Megha Jain

Royal Hobart Hospital

David Alcock, Nico Terblanche*

Royal Melbourne Hospital

Rochelle Cotter, Kate Leslie*, Marcelle Stewart, Nicolette Zingerle

Royal North Shore Hospital

Antony Clyde, Oliver Hambidge, Adam Rehak*

Royal Prince Alfred Hospital

Sharon Cotterell, Wilson Binh Quan Huynh, Timothy McCulloch*

St Vincent's Hospital

Erez Ben-Menachem*, Thomas Egan,

Westmead Hospital

Jennifer Cope, Richard Halliwell*

Austria

Allgemeines Krankenhaus Wien

Paul Fellinger, Simone Haselberger, Caroline Holaubek, Paul Lichtenegger, Florian Scherz, Werner Schmid*

Landesklinikum Hollabrunn

Franz Hoffer*

Landesklinikum Horn

Veronika Cakova, Andreas Eichwalder, Norbert Fischbach, Reinhold Klug*, Elisabeth Schneider, Martin Vesely, Reinhart Wickenhauser

Landesklinikum Krems

Karl Gernot Grubmueller, Marion Leitgeb*

Landesklinikum Neunkirchen

Friedrich Lang, Nancy Toro*

Landesklinikum Wiener Neustadt

Marlene Bauer, Friedrich Laengle*

Landesklinikum Zwettl

Thomas Mayrhofer*

Landeskrankenhaus Feldkirch

Christian Buerkle, Karin Forstner, Reinhard Germann*, Harald Rinoesl

Landeskrankenhaus Villach

Elke Schindler, Ernst Trampitsch*

Salzburger Landeskliniken

Gerhard Fritsch*

Belgium

Centre Hospitalier Régional dede Namur

Christian Szabo*

Centre Hospitalier Universitaire Brugmann

Jawad Bidgoli*, Hans Verdoodt

Cliniques Universitaires Saint-Luc

Patrice Forget, David Kahn*, Fernande Lois, Mona Momeni, Caroline Prégardien, Audrey Pospiech, Arnaud Steyaert, Laurent Veevaete

Heilig Hart Ziekenhuis Roeselaere

Dirk De Kegel*

Onze Lieve Vrouw Ziekenhuis Aalst

Karen De Jongh, Luc Foubert*

Universitair Ziekenhuis Antwerpen

Carine Smitz, Marcel Vercauteren*

Universitair Ziekenhuis Brussel

Jan Poelaert*, Veerle Van Mossevelde

Universitair Ziekenhuis Gent

Jacques Abeloos, Stefaan Bouchez, Marc Coppens, Luc De Baerdemaeker, Isabel Deblaere,
Ann De Bruyne*, Stefan De Hert*, Kristine Fonck, Bjorn Heyse, Tom Jacobs, Koen Lapage,
Anneliese Moerman, Martine Neckebroek, Aliaksandra Parashchanka, Nathalie Roels, Nancy
Van Den Eynde, Michael Vandenheuvel, JurgenVan Limmen, AnnVanluchene,
CarolineVanpeteghem, Patrick Wouters*, Piet Wyffels

Universitair Ziekenhuis Leuven

Christel Huygens, PunithaVandenbempt, Marc Van de Velde*

Ziekenhuis Oost Limburg campus St-Jan Genk

Dimitri Dylst*, Bruno Janssen, Evelien Schreurs

Brazil

Faculdade de Medicina e Hospital de Base de São José do Rio preto - SP

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Hospital das Clinicas da faculdade de Medicina da Universidade de Sao Paulo

Jaquelline Jardim, Joao Silva*

Hospital das Clínicas da Faculdade de Medicina de Botucatu

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Zhengyuan Liu*

Lishui People's Hospital of Zhejiang Province

Lihua Fan*

Nanfang Hospital of Southern Medical University

Jing Tang*

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Kaizhi Lu*, Ting Zhao

Affiliated Hospital of Hangzhou Normal University

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Jens Rolighed Larsen*, Søren Hübertz Smith

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Rebecca Monett Oestervig, Lars Rasmussen*

Rigshospitalet, JMC

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Tage Hansen Gade, Aarhus

Birgitte Brandsborg*

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Centre Hospitalier Universitaire de Rennes

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CHU d'Angers

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CHU de Bordeaux, Hôpital Pellegrin, Centre François Xavier Michelet

Matthieu Biais*

CHU de Bordeaux, Hôpital Pellegrin, Hôpital du Haut Lévêque

Alexandre Ouattara*, Marlene Sineus

CHU de Saint-Etienne

Serge Molliex*

CHU-Hospital Saint Louis, Paris

Marie Lim Legouge*

CHU-Lyon Sud, Lyon

Florent Wallet*

Hôpital Cochin, APHP, Paris

Antoine Tesniere*

Hôpital de Nantes

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Hôpital d'instruction des Armées (HIA) du Val-de-Grace

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Hopital Le Mans

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Charité Universitätsklinik Berlin, Campus Virchow Klinikum

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Ulrich Goebel*, Sarah Harris

Vivantes Klinikum Friedrichshain

Siegfried Veit*

Greece

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Evangelia Andreadaki*, Flora Souri

424 Army Hospital, Thessaloniki

Ioannis Katsiadramis*, Anthi Skoufi, Maria Vasileiou

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Agios Pavlos General Hospital, Thessaloniki

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Ahepa University Hospital, Thessaloniki

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Hippokrateion General Hospital of Athens

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Hippokration General Hospital, Thessaloniki

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Mexico

Centro Medico Abc

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Hospital Salubridad Culiacan Sinaloa

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Khalid Al 'Amri, Hafiz Aladin, Mohammed Asif Arshad, James Barraclough, Conor Bentley, Colin Bergin, Ronald Carrera, Aisling Clarkson, Michelle Collins, Lauren Cooper, Samuel Denham, Ewen Griffiths, Peter Ip, Somasundaram Jeyanthan, Kavita Joory, Satwant Kaur, Paul Marriott, Natalie Mitchell, Sukumar Nagaiah, Annette Nilsson, Nilesh Parekh*, Martin Pope, Joseph Seager, Hosam Serag, Alifia Tameem, Anna Thomas, Joanne Thunder, Andrew Torrance, Ravinder Vohra, Arlo whitehouse, Tony Wong

Queen Elizabeth Hospital Kings Lynn

Mark Blunt*, Kate Wong

Queen Victoria Hospital

Julian Giles*, Isabelle Reed*, Debbie Weller

Queens Hospital

Gillian Bell, Julie Birch, Rose Damant, Jane Maiden, Clare Mewies*, Claire Prince, Jane Radford

Robert Jones and Agnes Hunt Orthopaedic Hospital

Birender Balain, Robin Banerjee, Andrew Barnett, Ben Burston, Kirsty Davies, Jayne Edwards, Chris Evans, David Ford, Pete Gallacher, Simon Hill, David Jaffray, Sudheer Karlakki*, Cormac Kelly, Julia Kennedy, Nigel Kiely, Simon Lewthwaite, Chris Marquis, Matthew Ockendon, Stephen Phillips, Simon Pickard, James Richardson, Richard Roach, Tony Smith, Richard Spencer-Jones, Niall Steele, Julie Steen, Marck Van Liefland, Steve White

Rotherham Hospital

Matthew Faulds, Meredyth Harris, Carrie Kelly*, Scott Nicol, Sally Anne Pearson

Royal Blackburn Hospital

Srikanth Chukkambotla*

Royal Cornwall Hospital

Alyson Andrew, Elizabeth Attrill, Graham Campbell, Amanda Datson, Anna Fouracres, Juan Graterol, Lynne Graves, Bosun Hong, Alexander Ishimaru, Arvind Karthikeyan, Helen King, Tom Lawson*, Gregory Lee, Saoirse Lyons, Andrew Macalister Hall, Sophie Mathoulin, Eilidh Mcintyre, Danny Mclaughlin, Kathleen Mulcahy, Anna Ratcliffe, James Robbins, Weilin Sung, Adeoluwa Tayo, Lisa Trembath, Suneetha Venugopal, Robert Walker, Geoffrey Wigmore

Royal Derby Hospital

Catherine Boereboom, Charlotte Downes, Ryan Humphries, Susan Melbourne, Coral Smith, Samson Tou*, Shafa Ullah

Royal Devon and Exeter Hospital

Nick Batchelor, Leigh Boxall, Rupert Broomby, Tariq Deen, Alistair Hellewell, Laurence Helliwell, Melanie Hutchings, David Hutchins, Samantha Keenan, Donna Mackie, Alison Potter*, Frances Smith, Lucy Stone, Kevin Thorpe, Richard Wassall, Andrew Woodgate

Royal Free Hospital

Shelley Baillie, Tara Campbell, Sarah James*, Chris King, Daniela Marques de Araujo, Daniel Martin, Clare Morkane, Julia Neely, Rajkumar Rajendram

Royal Glamorgan Hospital

Megan Burton, Kathryn James, Edward Keevil, Orsolya Minik, Jenna Morgan, Anna Musgrave, Harish Rajanna, Tracey Roberts, Tamas Szakmany*

Royal Gwent Hospital

Michael Adamson*, Sandra Jumbe, Jennie Kendall, Mohan Babu Muthuswamy*

Royal Hallamshire Hospital

Charlotte Anderson, Andrew Cruikshanks, Sailaja Pothuneedi, Rachel Walker, Ian Wrench*, Lisa Zeidan

Royal Hampshire County Hospital

Diane Ardern, Benjamin Harris, Johanna Hellstrom, Jane Martin, Richard Thomas*, Nimu Varsani, Caroline Wrey Brown

Royal Infirmary Edinburgh

Philip Docherty*, Michael Gillies*, Euan McGregor, Helen Usher

Royal Lancaster Infirmary

Jayne Craig, Andrew Smith*

Royal London Hospital

Tahania Ahmad, Phoebe Bodger, Thais Creary, Kirsty Everingham, Alexander Fowler, Russ Hewson, Eke Ijuo, Marta Januszewska, Timothy Jones, Ilya Kantsedikas, Sumitra Lahiri, Aaron Lawson McLean, Edyta Niebrzegowska, Mandeep Phull*, Difei Wang, Nadine Wickboldt

Royal Preston Hospital

Jacqueline Baldwin, Donna Doyle, Sean Mcmullan, Michelle Oladapo, Thomas Owen*, Shiva Tripathi*, Alexandra Williams

Royal Surrey County Hospital

Hull Daniel, Peter Gregory, Tauqeer Husain, Justin Kirk-Bayley*, Edward Mathers, Laura Montague

Royal Sussex County Hospital

Stuart White*

Royal United Hospital Bath

Joanne Avis, Tim Cook*, Lola Dali-Kemmery, Ian Kerslake*, Victoria Lambourne, Annabel Pearson

Royal Victoria Infirmary

Christine Boyd, Mark Callaghan, Cathy Lawson, Roopa McCrossan, Vanessa Nesbitt, Laura O'connor, Julia Scott, Rhona Sinclair*

Russells Hall Hospital

Nahla Farid*, Ciro Morgese

Saint Mary's Hospital

Kailash Bhatia, Swati Karmarkar, Akbar Vohra*

Salisbury District Hospital

Jamil Ahmed*, Graham Branagan*, Monica Hutton, Andrew Swain

Sandwell & West Birmingham Hospitals NHS Trust

Jamie Brookes, Jonathan Cornell, Rachael Dolan, Jonathan Hulme*, Amanda Jansen van Vuuren, Tom Jowitt, Gunasheela Kalashetty, Fran Lloyd, Kiran Patel, Nicholas Sherwood

Scarborough Hospital

Lynne Brown, Ben Chandler*, Kerry Deighton, Temlett Emma, Kirsty Haunch

Scunthorpe General Hospital

Michelle Cheeseman, Kathy Dent, Sanjeev Garg, Carol Gray*, Marion Hood, Dawn Jones, Joanne Juj, Atideb Mitra*, Roshan Rao, Tara Walker

Southend University Hospital

Mashel Al Anizi, Clarissa Cheah, Yushio Cheing, Francisco Coutinho, Prisca Gondo, Bernard Hadebe, Mazvangu Onie Hove, Ahamed khader*, Bobby Krishnachetty*, Karen Rhodes, Jagdish Sokhi

Southmead Hospital

Katie-Anne Baker, Wendy Bertram, Alex Looseley*, Ronelle Mouton

St Mary's Hospital, London

Glenn Arnold, Shobhit Arya, Danai Balfoussia, Linden Baxter, James Harris, Craig Jones, Alison Knaggs, Sheraz Markar, Anisha Perera, Alasdair Scott*, Asako Shida, Ravneet Sirha, Sally Wright

St Peter's Hospital

Victoria Frost, Catherine Gray, Mark MacGregor*

St Thomas' Hospital

Emma Andrews, Lindsay Arrandale, Stephen Barrett, Heena Bidd*, Elna Cifra, Mariese Cooper, Dragos Dragnea, Cifra Elna, Jennifer Maclean, Sonja Meier, Donald Milliken, Christopher Munns, Nadir Ratanshi, Abegail Salvana, Anthony Watson

Sunderland Royal Hospital

Hani Ali, Gill Campbell, Rebecca Critchley*, Catherine Hicks, Alison Liddle, Marc Pass, Charlotte Ritchie, Charlotte Thomas, Lingxi Too*, Sarah Welsh

Royal Oldham Hospital

Talvinder Gill*, Joanne Johnson, Joanne Reed

Royal Orthopaedic Hospital

Edward Davis*, Sam Papadopoullos

Torbay Hospital

Clare Attwood, Andrew Biffen*, Kerenza Boulton, Sophie Gray, David Hay, Sarah Mills, Jane Montgomery*, Rory Riddell, James Simpson

Trafford Hospitals

Neeraj Bhardwaj, Elaine Paul, Nosakhare Uwubamwen, Akbar Vohra*

Tunbridge Wells Hospital

Emer Berntsen, Tracey Nolan*, Angus Turner

University College London

Maini Alexander, James Arrich, Swarna Arumugam, Douglas Blackwood, Victoria Boggiano, Robyn Brown, Yik Lam Chan, Devnandan Chatterjee, Ashok Chhabra, Rachel Christian, Hannah Costelloe, Madeline Coxwell Matthewman, Emma Dalton, Julia Darko, Maria Davari, Tejal Dave, Matthew Deacon, Shantal Deepak, Holly Edmond, Jessica Ellis, Ahmed El-Sayed,

Philip Eneje, Rose English, Renee Ewe, William Foers, John Franklin, Laura Gallego, Emily Garrett, Olivia Goldberg, Harry Goss, Rosanna Greaves, Rudy Harris, Charles Hennings, Eleanor Jones, Nelson Kamali, Naomi Kokkinos, Carys Lewis, Leda Lignos, Evaleen Victoria Malgapo, Rizwana Malik, Andrew Milne, John-Patrick Mulligan, Philippa Nicklin, Natasha Palipane, Thomas Parsons, Rebecca Piper, Rohan Prakash, Byron Ramesh, Sarah Rasip, Jacob Reading, Mariam Rela, Anna Reyes, Stephens Robert*, Martin Rooms, Karishma Shah, Henry Simons, Shalil Solanki, Emma Spowart, Amy Stevens, Christopher Thomas, Helena Waggett, Arrash Yassaee

University Hospital of North Durham

Anthony Kennedy, Sara Scott, Sameer Somanath*

University Hospital of North Tees

Andrew Berg, Miguel Hernandez, Rajesh Nanda*, Ghanshyambhai Tank, Natalie Wilson, Debbie Wilson

University Hospital of Wales

Yassr Al-Soudaine, Matthew Baldwin, Matthew Baldwin, Julie Cornish, Zoe Davies, Leigh Davies*, Marc Edwards, Natasha Frewer, Sian Gallard, James Glasbey, James Glasbey, Rhiannon Harries, Luke Hopkins, Taeyang Kim, Vilavan Koompirochana, Simon Lawson, Megan Lewis, Zaid Makzal, Sarah Scourfield

University Hospital Southampton

Yousra Ahmad, Sarah Bates, Clare Blackwell, Helen Bryant, Suzanne Coulter, Ross Cruickshank, Sonya Daniel, Thomas Daubeny, Mark Edwards*, Kim Golder, Lesley Hawkins, Bryant Helen, Honor Hinxman, Denny Levett, Ben Skinner, Joseph Walsgrove

Wansbeck General Hospital

Mike Bradburn*, Jane Dickson

Watford General

Kathryn Constantin, Markwell Karen, Peter O'Brien*, Lynn O'Donohoe, Hannah Payne, Saul Sundayi, Elaine Walker

West Suffolk Hospital

Jenny Brooke, Jon Cardy, Sally Humphreys, Laura Kessack, Christiane Kubitzek*, Suhas Kumar

Weston General Hospital

Donna Cotterill, Emil Hodzovic, Gurunath Hosdurga*, Edward Miles*, Glenn Saunders

Whipps Cross University Hospital

Marta Campbell, Peter Chan, Kim Jemmett, Ashok Raj*

Whittington Hospital

Aditi Naik, Rajarajan Ramamoorthy*, Nimesh Shah, Axel Sylvan*

Worcestershire Royal Hospital

Katharine Blyth, Andrew Burtenshaw*, David Freeman, Emily Johnson, Philip Lo, Terry Martin, Emma Plunkett, Julie Wollaston*

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York Hospital

Andrew McDonald, Davies Simon*, Vivek Sinha, Thomas Smith

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Beth Israel Deaconess Medical Centre

Valerie Banner-Goodspeed*, Myles Boone*, Kathleen Campbell, Fengxin Lu, Joseph Scannell

Columbia University Medical Center

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Duke University in Durham

Naraida Balajonda, Karen Clemmons, Carlos Conde, Bonita Funk, Roger Hall, Thomas Hopkins*, Omowunmi Olaleye, Omer Omer, Michelle Pender, Angelo Porto, Alice Stevens, Peter Waweru, Erlinda Yeh

Icahn School of Medicine at Mount Sinai

Daniella Bodansky, Adam Evans*, Steven Kleopoulos, Robert Maril, Edward Mathney*, Angela Sanchez, Elizabeth Tinuoye

Massachusetts General Hospital

Brian Bateman*, Kristen Eng, Ning Jiang, Karim Ladha*, Joseph Needleman

Mount Zion

Lee-lynn Chen*

Parnassus/Moffit-Long Hospital

Lee-lynn Chen*, Rondall Lane, David Robinowitz

South Shore Hospital

Neil Ghushe, Mariam Irshad, Samir Patel*

San Francisco Veterans Affairs Medical Center

Steven Takemoto*, Art Wallace*

The University of Maryland School of Medicine

Michael Mazzeffi*, Peter Rock, Karin Wallace, Xiaomao Zhu

University of Pennsylvania

Pandora Chua, Lee Fleisher*, Matthew Mattera

University of Washington Medical Center

Rebecca Sharar, Stephan Thilen*, Miriam Treggiari

UPMC Presbyterian and Montefiore Hospitals

Angela Morgan, Iwan Sofjan, Kathirvel Subramaniam*

Washington University in St Louis

Michael Avidan*, Hannah Maybrier, Maxwell Muench, Troy Wildes