Changing patterns of multidisciplinary team treatment, early mortality, and survival in colorectal cancer

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Abstract

Background: This study reports early mortality and survival from colorectal cancer in relation to the pattern of treatments delivered by the multidisciplinary team (MDT) meeting at a high-volume institution in England over 14 years.

Methods: All patients diagnosed with colorectal cancer and discussed during MDT meetings from 2003 to 2016 at a single institution were reviewed. Three time intervals (2003–2007, 2008–2012, and 2013–2016) were compared regarding initial surgical management (resection, local excision, non-resection surgery, and no surgery), initial oncological therapy, 90-day mortality, and crude 2-year survival for the whole cohort. Sub-analyses were performed according to age greater or less than 80 years.

Results: The MDT managed 4617 patients over 14 years (1496 in the first interval and 1389 in the last). Over this time, there was a reduction in emergency resections from 15.5 per cent to 9.0 per cent (P < 0.0001); use of oncological therapies increased from 34.6 per cent to 41.6 per cent (P < 0.0001). The 90-day mortality after diagnosis of colorectal cancer dropped from 14.8 per cent to 10.7 per cent (P < 0.001) and 2-year survival improved from 58.6 per cent to 65 per cent (P < 0.001). Among patients aged 80 years or older (425 and 446, in the first and last intervals respectively) there was, in addition, a progressive increase in ‘no surgery’ rate from 33.6 per cent to 50.2 per cent (P < 0.001) and 2-year survival improved from 58.6 per cent to 65 per cent (P < 0.001). The 90-day mortality after elective resection fell from 10.0 per cent (18 of 180) to 3.3 per cent (5 of 151; P < 0.013).

Conclusions: Survival from colorectal cancer improved significantly over 14 years. Among patients aged ≥80 years, major changes in the type of treatment delivered were associated with a decrease in postoperative mortality.

Introduction

Multidisciplinary teams (MDTs) manage patients with colorectal cancer1 in light of advances in staging, oncological therapies and surgery, to ensure that individual patients receive appropriate advice regarding treatment. Surgical resection offers the best chance of cure for most patients with colorectal cancer. For the old, the frail, or those with advanced disease, best management can be more difficult to determine. Patients presenting as emergencies often share these characteristics3,4. In these groups, local expertise and the evolution of therapies will strongly influence the choice of treatment advised. Many high-risk patients may be managed by local excision, radiotherapy (if rectal cancer), de-functioning stomas, or stenting5.

MDTs have the potential to increase survival from cancer by improving results from existing treatments, by developing or adopting more effective surgical strategies and by offering more patients more appropriate management. Most reports on the treatment of colorectal cancer focus on the effects and outcomes of particular interventions for selected patients such as minimally invasive surgery, chemotherapy, and stenting6,7. There has been little consideration of the types and distribution of treatments recommended by an MDT as potentially significant determinants of clinical outcomes or of survival from the disease among its catchment population.

In this unit, changes in the type of treatment delivered, implemented to reduce post-resection mortality, were evaluated to assess this important function in the cancer pathway.

This study investigated the potential relationship between major changes in the treatments delivered by the MDT and early mortality and crude survival in a cohort of patients with colorectal cancer managed by the MDT and among patients aged 80 years or older.

Methods

Study design and setting

All NHS patients diagnosed with colorectal cancer between January 2003 and December 2016 at Portsmouth Hospitals University NHS Trust were reviewed. Patients with metachronous colorectal cancers were included and analysed in relation to management of their first cancer. Patients with features typical of colorectal cancer (such as on CT) but not biopsy proven, were also included.
All data used for this study had been approved for use by, and previously submitted to, the National Bowel Cancer Audit. Data collected prospectively included patient demographics, date of presentation, cancer stage, treatment received, and postoperative outcomes, including perioperative mortality and long-term survival. Deaths and date of death were identified from ‘patient administration systems’ records (updated regularly) up to August 2020. This study followed STROBE guidelines and has been conducted in compliance with the Declaration of Helsinki.20

Of note, Portsmouth Hospitals University NHS Trust is a large District General Hospital providing acute and elective services to a catchment population of approximately 650,000. The colorectal cancer MDT meets weekly to discuss all newly diagnosed elective patients and to recommend management. For patients admitted acutely and needing urgent management, discussions take place between relevant MDT specialists and the patient to decide the best care. Such patients are notified to the full MDT later. Ultimately, choice of treatment is decided in discussion between the responsible clinician and the patient. Deaths and complications are discussed weekly in a surgical quality assurance meeting. Portsmouth joined the National Bowel Cancer Screening Programme in 2009. Advances in the MDT’s care of patients with colorectal cancer during the course of this study included a policy that a consultant be scrubbed for all colorectal cancer resections (2001), improved surgical risk assessment (2001), better cross-sectional imaging (2003), stenting (2003), laparoscopic resection (2003), increasing use of radiological TNM staging (2004), 24-h availability of CT (2005), provision of a surgical high care unit (2008), enhanced recovery programme (2008), cardiopulmonary exercise testing (2013), robotic resection (2013), better and more personalized oncological therapy (including cetuximab and panitumumab for RAS wild-type metastatic cancers, 2017), and routine assessment of resected specimens for mismatch-repair deficiency (2019).

### Outcomes of interest

The proportions of patients in the different treatment groups were compared over three time intervals (2003–2007, 2008–2012, and 2013–2016) to assess outcomes associated with the MDT’s approach to reducing postoperative mortality. Principal endpoints were 90-day mortality and crude 2-year survival of the whole colorectal cancer cohort and separately, according to age (under 80 years or 80 years or older). Secondary endpoints were 90-day mortality and 2-year survival by treatment group. Postoperative mortality was counted from the date of surgery. Survival was measured from the date of operation, or the date of diagnosis if there was no surgery. Crude survival was chosen over other indices because it was the measure used in the National Bowel Cancer Audit over this time interval. Reasons for not operating were recorded prospectively. For patients undergoing excision or resection, Dukes’ classification is reported as TNM was not used in the first years of the study.

### Statistical analysis

Categorical variables were analysed with Pearson’s chi-squared test or Fishers exact test as appropriate. Grouped continuous data (such as age at presentation) were compared with the non-parametric Mann–Whitney U Test (binary) or Kruskal–Wallis test (more than two categories). Missing data were handled as follows: CT staging was excluded because the data were not consistently entered on the database; also, patients who did not

### Table 1 Patient demographics and patterns of treatment delivered by the multidisciplinary team over time

| Time interval | 2003–2007 | 2008–2012 | 2013–2016 | P 2003–2007 versus 2013–2016 |
|---------------|-----------|-----------|-----------|-----------------------------|
| Number diagnosed in time interval |           |           |           |                             |
| Number diagnosed per year (mean) | 1496 | 1732 | 1389 | – |
| Screening detected per year (mean) | 299 | 346 | 347 | – |
| Age (years), median (i.q.r) | 74 (23–102) | 74 (24–100) | 74 (27–58) | 0.033 |
| Sex ratio (M:F) | 784:712 | 960:772 | 783:606 | – |
| Distribution of surgical treatments over time |       |           |           |                             |
| Resections | 1046 (70.2) | 1195 (69) | 849 (61.1) | <0.0001 |
| Elective | 814 (54.4) | 997 (57.6) | 724 (52.1) | <0.001 |
| Emergency | 232 (15.5) | 198 (11.4) | 125 (9.0) | <0.0001 |
| Non-resectional surgery | 98 (6.2) | 79 (4.6) | 64 (4.6) | – |
| Local excision | 40 (2.7) | 56 (3.2) | 79 (5.7) | <0.001 |
| No surgery | 317 (21.2) | 402 (23.2) | 397 (28.6) | <0.0001 |
| Distribution of oncological treatments over time |       |           |           |                             |
| Chemotherapy with/without radiotherapy | 517 (34.6) | 721 (41.6) | 578 (41.6) | <0.0001 |

Values are n (%) unless otherwise indicated. Comparisons between groups were conducted with a chi-squared test.
have surgery for unknown reasons were excluded from the relevant analysis. Kaplan–Meier analysis was used to plot survival curves. Survival is presented as median (95 per cent c.i.) absolute survival or as per cent survival at 2 years. Comparisons between groups were assessed with log rank tests with significance set at less than 0.05. Statistical analysis was performed with SPSS® version 26.0 (IBM, Armonk, New York, USA) and GraphPad Prism® version 7.0 (GraphPad Software, San Diego, California, USA).

### Table 2 Results in the whole cohort according to surgical treatment received over time, all ages

| Surgical treatment group       | Time interval                  | 2003–2007 | 2008–2012 | P versus 2003–2007 | 2013–2016 | P versus 2003–2007 |
|--------------------------------|--------------------------------|-----------|-----------|--------------------|-----------|--------------------|
| **Resections**                  |                                |           |           |                    |           |                    |
| 90-day mortality               | 36 (4.4)                       | 19 (1.9)  | –         | 11 (1.5)           | <0.001    |
| 2-year survival                | 685 (84.2)                     | 883 (88.6)| –         | 664 (91.7)         | <0.001    |
| **Emergency resection**        |                                |           |           |                    |           |                    |
| 90-day mortality               | 44 (18.9)                      | 34 (17.2) | –         | 18 (14.4)          |           |
| 2-year survival                | 118 (50.9)                     | 101 (51.0)| –         | 70 (56.0)          |           |
| **All resections**             |                                |           |           |                    |           |                    |
| 90-day mortality               | 80 (7.6)                       | 53 (4.4)  | –         | 29 (3.4)           | <0.001    |
| 2-year survival                | 803 (76.8)                     | 984 (82.3)| –         | 734 (86.5)         | <0.0001   |
| **Non-resectional surgery**    |                                |           |           |                    |           |                    |
| 90-day mortality               | 26 (28)                        | 17 (21.5) | –         | 6 (9.4)            | 0.005     |
| 2-year survival                | 9 (9.7)                        | 14 (17.7) | –         | 12 (18.8)          |           |
| **Local excision**             |                                |           |           |                    |           |                    |
| 90-day mortality               | 0 (0)                          | 0 (0)     | –         | 0 (0)              |           |
| 2-year survival                | 38 (95)                        | 51 (91.1)| –         | 75 (94.9)          |           |
| **No surgery**                 |                                |           |           |                    |           |                    |
| 90-day mortality               | 115 (36.3)                     | 126 (31.3)| –         | 113 (28.5)         | 0.026     |
| Median survival (95% c.i.) (months) | 5.3 (4.0–6.5)            | 7.3 (5.9–8.6)| – | 8.0 (6.4–9.5)     | <0.001    |
| 2-year survival                | 26 (8.2)                       | 59 (14.7) | –         | 84 (21.2)          | <0.001    |
| **All patients, all treatments**|                                |           |           |                    |           |                    |
| 90-day mortality               | 221 (14.8)                     | 196 (11.3)| –         | 148 (10.7)         | <0.001    |
| Median survival (95% c.i.) (years) | 3.4 (2.9–3.9)            | 4.8 (4.0–5.6)| <0.0001 | 4.9 (4.0–5.7)     | <0.001    |
| 2-year survival                | 876 (58.6)                     | 1108 (64)| –         | 905 (65.2)         | <0.001    |

Values are n (%) unless otherwise indicated. *Survival presented as median (95 per cent c.i.) or as per cent survival at 2 years. Comparisons between groups assessed using log rank tests with significance set at less than 0.05 or chi-squared test as appropriate.

### Table 3 Results among patients aged under 80 years according to surgical treatment received over time

| Surgical treatment group       | Time interval                  | 2003–2007 | 2008–2012 | P versus 2003–2007 | 2013–2016 | P versus 2003–2007 |
|--------------------------------|--------------------------------|-----------|-----------|--------------------|-----------|--------------------|
| **Resections**                  |                                |           |           |                    |           |                    |
| 90-day mortality               | 809 (75.5)                     | 950 (76.5)| –         | 667 (70.7)         | 0.015     |
| 2-year survival                | 44 (5.4)                       | 27 (2.8)  | 0.006     | 19 (2.8)           | 0.014     |
| **Elective**                   |                                |           |           |                    |           |                    |
| 90-day mortality               | 654 (80.8)                     | 803 (84.5)| 0.041     | 592 (88.8)         | <0.001    |
| 2-year survival                | 634 (59.2)                     | 802 (64.6)| 0.008     | 573 (60.8)         | –         |
| **Emergency**                  |                                |           |           |                    |           |                    |
| 90-day mortality               | 175 (16.3)                     | 148 (11.9)| 0.002     | 94 (10.0)          | <0.001    |
| 2-year survival                | 18 (2.8)                       | 8 (1.0)   | 0.009     | 6 (1.0)            | <0.001    |
| **Non-resectional surgery**    |                                |           |           |                    |           |                    |
| 90-day mortality               | 26 (14.9)                      | 19 (12.8) | –         | 13 (13.8)          | –         |
| 2-year survival                | 97 (55.4)                      | 78 (52.7) | –         | 53 (56.4)          | –         |
| **Local excision**             |                                |           |           |                    |           |                    |
| 90-day mortality               | 63 (5.9)                       | 52 (4.2)  | –         | 51 (5.4)           | –         |
| 2-year survival                | 13 (1.2)                       | 11 (0.8)  | –         | 14 (1.5)           | –         |
| **No surgery**                 |                                |           |           |                    |           |                    |
| 90-day mortality               | 25 (2.3)                       | 40 (3.2)  | –         | 52 (5.5)           | <0.001    |
| 2-year survival                | 24 (96)                        | 38 (95)   | –         | 50 (96.2)          | –         |
| **Total patients in time interval** | 174 (16.2)                     | 200 (16.1)| –         | 173 (18.3)         | –         |
| 2-year survival                | 13 (7.5)                       | 26 (13)   | –         | 37 (21.4)          | 0.0001    |
| **Oncology**                   |                                |           |           |                    |           |                    |
| Radiotherapy                   | 38 (3.6)                       | 45 (3.6)  | –         | 27 (2.9)           | –         |
| Chemotherapy                   | 335 (31.3)                     | 434 (34.9)| –         | 351 (37.2)         | 0.005     |
| Chemoradiotherapy              | 89 (8.3)                       | 146 (11.8)| –         | 105 (11.1)         | 0.032     |
| Chemo/radiotherapy             | 462 (43.1)                     | 625 (50.3)| <0.001    | 483 (51.2)         | <0.001    |

Values are n (%) unless otherwise indicated. Patients were grouped according to nature of surgery or none. Use of oncological therapy is not broken down according to surgical treatment group. *Survival presented as median (95 per cent c.i.) or as per cent survival at 2 years. Comparisons between groups assessed using log rank tests with significance set at less than 0.05 or chi-squared test as appropriate.
Overall, 4617 patients were diagnosed with colorectal cancer and discussed in the colorectal MDT meetings over 14 years, including 33 metachronous cancers and 265 detected by screening (Table 1). Although there was no change in median age over time, the proportion of patients aged 80 years or older increased from 28.4 per cent (425 of 1496) in 2003–2007 to 32.2 per cent (446 of 1389) in 2013–2016 (P = 0.031).

**Overall cohort**
Over time, there was a reduction in overall resections from 70.2 per cent to 61.1 per cent (P < 0.0001) and in emergency resections from...
15.5 per cent to 9.0 per cent ($P < 0.0001$) (Table 1). Use of oncolgical therapies increased from 34.6 per cent to 41.6 per cent ($P < 0.0001$). The 90-day mortality improved significantly for the whole cohort (however treated) as well as for each treatment group (Table 2). Among patients undergoing elective resection, 90-day mortality fell from 4.4 per cent to 1.5 per cent, ($P < 0.001$). There was a relative increase in median survival of 44 per cent (3.4 (2.9–3.9) to 4.9 (4.0–5.7) years; $P < 0.0001$). The 2-year survival increased from 58.6 per cent to 65 per cent ($P < 0.001$). Most of the survival improvement occurred between the first and second time intervals (Table 2).

**Patients aged under 80 years**

Between 2009 and 2016 a mean 36 patients per annum (range 9–63) aged 60–69 years, were referred to the MDT from the Bowel Cancer Screening Programme (mean 21 per annum in 2008–2012, increasing to a mean 48 per annum in 2013–2016). Among patients undergoing elective local excision or resection, there was an increase in Dukes’ class A over time from 21.4 per cent (141 of 258) to 32.3 per cent (202 of 625) ($P < 0.001$), a drop in Dukes’ B and an increase in the proportion of patients with metastatic disease (4.7 per cent (31 of 658) rising to 8.0 per cent (50 of 625); $P = 0.015$). Table 3 shows changes in the pattern of treatments delivered, 90-day mortality, and 2-year survival according to treatment. Compared with the first time interval, there was a significant reduction in the proportion of patients undergoing emergency resection in periods two (11.9 per cent versus 16.3 per cent; $P = 0.002$) and three (10 per cent; $P < 0.001$). The elective resection rate remained unchanged overall—an increased rate in patients aged under 70 years from 57.9 per cent to 65.8 per cent (in keeping with screening), balanced by a drop among patients aged 70–80 years. There was no significant increase in patients allocated to ‘no surgery’ among those aged under 80 years. Use of chemotherapy or radiotherapy increased from 43.1 per cent to 51.2 per cent, mostly between the first and second time intervals ($P < 0.001$).

The 90-day mortality after diagnosis of colorectal cancer in patients aged under 80 years dropped steadily, with significant improvement in the second time interval (Table 3). The 90-day mortality after elective or any resection dropped significantly by the second time interval and did not drop further. Median and 2-year survival for all patients increased early (significantly by the second time interval). The relative increase in 2-year survival was 12 per cent (Table 3 and Fig. 1).

**Patients aged 80 years or more**

Among those undergoing elective resection or local excision in this age group, there was no significant change in resection pathology (Dukes’ classification) over time. Table 4 shows changes in the pattern of treatments delivered, and in survival outcomes, according to treatment. As with the younger patients, there was a reduction in emergency resections and an increase in oncological treatment. In contrast with younger patients, however, there was a significant increase in ‘no surgery’ and a drop in elective resections among those aged 80 years or older. An increasing proportion of patients (41.2 per cent in second interval, $P = 0.018$; 50.2 per cent in the third; $P < 0.0001$) were not offered any surgery (Table 4). Advanced disease accounted for 13.9 per cent (59 of 425) patients not being offered surgery in the first interval and 15.2 per cent (68 of 446) in the last. Patient unfitness was the reason given for not operating in 14.4 per cent (61 of 425) in the first interval increasing to 26.9 per cent (120 of 446) ($P < 0.00001$) in the last. For patients not undergoing surgery in this age group, use of chemotherapy or radiotherapy decreased from 25.9 per cent (37 of 143) to 21.4 per cent (48 of 224; $P < 0.001$). Median survival following diagnosis increased from 16.1 to 21.2 months mostly between the first and second time intervals ($P = 0.013$). The relative increase in 2-year survival for all diagnosed patients was 18 per cent by the third time interval (Table 4, Fig. 1). After resections (overall or electively), the 2-year survival increased significantly by the second time interval concomitant with significant increases in ‘no surgery’ and in oncological treatment but before significant changes in overall resection rate or in elective resections (Table 4).

The 90-day mortality after diagnosis of colorectal cancer or following resection (elective/any) showed no significant change in the second time interval. In the third interval, there was a significant improvement in 90-day mortality after elective resection from 10 per cent to 3.3 per cent ($P = 0.017$) accompanying a drop in the elective resection rate from 42.4 per cent to 33.9 per cent ($P = 0.010$) and increase in ‘no surgery’ to 50.2 per cent.

**Discussion**

Management of colorectal cancer varies across England with regard to treatment delivered and clinical outcomes. At a local level, MDT meetings recommend treatment for individual patients based on the evidence presented. Team-working within MDT meetings has been extensively studied. The relationship between the choice of treatment it delivers, and clinical outcomes has received less attention. This study examined survival outcomes over a 14-year interval during which, major changes in treatment distribution had been instituted to reduce post-resection mortality among the old. The report highlights the possibility that, in addition to improving results from individual treatments, an MDT may improve its survival outcomes by doing more or less of particular treatments.

The 90-day mortality and 2-year survival by age group and for the whole colorectal cancer cohort were selected as primary endpoints for the study. The 2-year rather than 5-year survival was chosen to provide earlier insights into MDT outcomes that might prompt remedial action if needed. The potential for discrepancy between management recommended versus that provided was dealt with by regarding the latter as the care for which the MDT was accountable. Changes in outcomes for individual treatment groups over the course of the study were not primary endpoints because the treatment groups were potentially influenced by selection bias as well as by treatment improvements over time.

During the study, 2-year crude survival from colorectal cancer increased from 59 per cent to 65 per cent. Much of this improvement took place in the early part of the study and likely reflected progress in care across the UK. Among patients aged less than 80 years, there was an early reduction in emergency resections and increase in oncological therapy but no change in the elective resection rate and no increase in ‘no surgery’. The observed improvements in overall 90-day mortality and 2-year survival in this age group are consistent with the impacts of screening, advances in surgical and perioperative care, and increased use of more effective oncological therapies reported during this time.

Patients aged 80 years or more also benefitted from improving cancer care in the NHS: 2-year survival after resection increased early in this study, alongside increased use of oncological therapies (before any substantial change in resection rates);
however, significant improvement in 90-day mortality for all diagnosed patients and for those undergoing resection in this age group followed major changes in the distribution of treatments delivered by the MDT, notably a drop in the elective (as well as emergency) resection rate and a substantial increase in the proportion of patients receiving no surgery. Reducing the elective resection rate by a quarter was associated with a two-thirds fall in 90-day mortality after resection. Contrasted with patients aged under 80 years, the proportion of those aged 80 years or more turned down for resection on account of impaired fitness almost doubled as the team raised the threshold for surgery to reduce post-resection mortality. Improved 90-day mortality for all diagnosed patients in this age group occurred despite decreasing use of resections. Improving survival was also seen among the growing group of patients aged 80 years or more allocated to ‘no surgery’. This could not be explained by increased use of oncological therapies (which decreased in the ‘no surgery’ group) and likely reflects inclusion of patients who would previously have undergone surgery but lived a little longer without it.

Survival from bowel cancer in England is poorer than in many countries, arguably due to lower resection rates especially among the old; however, high rates of resection are only appropriate if postoperative mortality rates are acceptable. Reducing the resection rate to improve outcomes was an initial response while the MDT simultaneously worked to improve perioperative and operative care in the old and for all its patients. Recent excellent 90-day mortality following elective resection suggests that this unit should now offer resection more readily. Advances in individual risk assessment, selection, and surgical care may make resection appropriate for some patients who would previously have been turned down.

This study has several limitations. Although all data were entered prospectively, the analysis was retrospective and covered a time during which there were substantial changes in referral pathways, quality of imaging, and improvements in individual treatments, which likely contributed to the better outcomes observed and which are acknowledged (above). Discussion of detailed radiological staging, which would have been valuable in comparing groups over time, was not possible because data collection was not consistent over the study interval. The MDT did not set criteria for determining who would be offered resection (as is reflected in the gradual changes in numbers resected) and this study does not attempt to advise on this. The study focused mostly on surgical rather than oncological management because it arose from an attempt to deal with post-resection (surgical) mortality. Finally, although we had reliable data concerning urgency of operation, we did not have data on urgency of presentation. Some of the drop in emergency operating could have been attributable to reduced emergency presentations after screening and public awareness campaigns.

Despite working to provide optimal treatment for each patient, MDT outcomes for particular subgroups may be less satisfactory. These insights are most easily gained by regular review of treatments delivered by the MDT in relation to results among the whole managed population and in patient subgroups. For some MDTs, changing the type of treatment provided to particular groups of patients has the potential to improve outcomes.

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Data availability
Data underlying this article will be shared on reasonable request to the corresponding author.

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