# Swallowing outcomes following surgical and non-surgical treatment for advanced laryngeal cancer

## E BURNIP<sup>1</sup>, S J OWEN<sup>2</sup>, S BARKER<sup>1</sup>, J M PATTERSON<sup>3</sup>

<sup>1</sup>Speech and Language Therapy Department, Sunderland Royal Hospital, <sup>2</sup>Speech and Language Therapy Department, Freeman Hospital, Newcastle upon Tyne, and <sup>3</sup>Institute for Health and Society, Newcastle University, UK

#### Abstract

*Background*: Treatment for advanced laryngeal cancer includes surgery, and/or chemoradiotherapy or radiotherapy. Each of these treatments results in major changes to the swallowing mechanism. Dysphagia is strongly correlated with poorer quality of life. A good understanding of outcomes is needed for well-informed treatment decisions.

*Method*: This study reports on patients' swallowing outcomes following surgical and non-surgical treatments based on the results of three different swallowing tests. A total of 123 data sets were collected in out-patient clinics across two hospitals in North East England.

*Results*: There were no significant differences between treatment groups for patient-reported swallowing outcomes or swallowing performance. However, patients who had undergone chemoradiotherapy or radiotherapy (with or without laryngectomy) had significantly more diet restrictions than other groups.

*Conclusion*: Long-term dysphagia is a common outcome of treatment for advanced laryngeal cancer. Patients treated with chemoradiotherapy and laryngectomy reported the worst overall outcomes. More longitudinal prospective research with large treatment groups is needed to investigate swallowing outcomes following different treatment methods.

Key words: Dysphagia; Chemoradiotherapy; Laryngectomy; Head And Neck Cancer; Larynx; Swallowing

#### Introduction

Treatment for advanced laryngeal cancer has evolved to include larynx-preserving interventions such as chemoradiotherapy or radiotherapy (RT), as well as traditional surgical techniques such as total laryngectomy or pharyngolaryngectomy. Trials have shown that chemoradiotherapy, with careful patient selection, can provide survival rates that are broadly comparable with laryngectomy.<sup>1,2</sup>

Patients need to be made aware of the short- and long-term morbidity risk associated with these treatment options. These treatments have implications for swallowing ability. This is a major issue for head and neck cancer patients.<sup>3,4</sup> Up to 72 per cent of laryngectomees report long-term swallowing problems.<sup>5</sup> Swallowing transit time can become prolonged once the larynx is removed. Some patients may present with specific problems such as fibrosis or strictures, which further hamper functionality. Patients who have undergone pharyngolaryngectomy may have additional problems depending on the extent of resection and the nature of reconstruction.<sup>6</sup>

Studies examining swallowing outcomes of chemoradiotherapy often include patients on the basis of reported swallowing difficulty. Furthermore, tumour sites are often combined, which makes it difficult to extract findings relevant to laryngeal carcinoma specifically. These studies report significant deterioration in swallowing following chemoradiotherapy,<sup>7,8</sup> with side effects such as oedema, reduced sensation and longterm fibrotic changes that markedly affect functioning. Furthermore, rates of aspiration are as high as 84 per cent,<sup>8–10</sup> which can result in complications such as pneumonia. Some patients may depend on a longterm feeding tube, which is a major determinant of poorer quality of life in head and neck cancer patients.<sup>11</sup> Where disease is still present, or has recurred, a salvage laryngectomy may be necessary. Evidence suggests that despite dual modality treatment, the majority of patients are able to manage a soft to normal diet.<sup>12</sup> A small number of post-chemoradiotherapy patients may be offered a laryngectomy because of a severely dysfunctional larynx, in the absence of disease. Patients undergoing this procedure need to

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be carefully counselled about the subsequent impact on function. A single case study suggested that a laryngectomy for a dysfunctional larynx had little effect on the patient's diet.<sup>13</sup>

A comparison of the outcomes reported by laryngeal cancer studies is difficult because swallowing can be measured in different ways, ranging from a description of the pathophysiology based on videofluoroscopy findings, to self-report and quality of life measures. Many of the standardised videofluoroscopy measurements, such as hyolaryngeal elevation, cannot be used with laryngectomy patients because of anatomical differences. This makes comparisons between surgical and non-surgical groups more problematic. Furthermore, clinician-reported and patient-reported dysphagia outcomes can vary;<sup>14</sup> both of these perspectives need to be included in descriptions of swallowing status.

The current study aimed to investigate swallowing outcomes following laryngectomy, and/or chemoradiotherapy or RT for advanced laryngeal cancer.

### **Materials and methods**

#### Patients

Patients were eligible for inclusion in the study if they had been diagnosed with advanced laryngeal cancer and had been treated with one of five interventions: total laryngectomy only; total laryngectomy with RT; radical RT only; chemoradiotherapy; or total laryngectomy and chemoradiotherapy.

Patients were consecutively recruited via multidisciplinary head and neck cancer clinics held at two hospitals in North East England over a five-month period (September 2010–January 2011).

Patients were only included in the study if they had been disease-free for at least six months following their last treatment. Patients were excluded if they had significant cognitive impairment, or had difficulty understanding English and were therefore unable to complete a questionnaire. Patients with conditions that may impact on swallowing such as stroke were also excluded.

## Assessments

Three clinical swallowing assessments were administered as follows.

In the water swallow test,<sup>15</sup> patients were asked to drink 100 ml of water as quickly as was comfortably possible. The test was timed and a measure of swallow capacity (millilitres swallowed divided by time taken) was derived. The test was not performed if the patient reported choking on fluids, in which case a score of zero was given. Patients who choked during the test were asked to stop immediately. Valves were checked for leakage and changed prior to assessment if necessary.

Patients completed the normalcy of diet subsection of the Performance Status Scale, which is a validated, clinician-rated scale of diet for Head and Neck Cancer Patients.<sup>16</sup> Scores range from 0 (nil by mouth) to 100 (indicating a full, unrestricted diet). Patients also completed the M D Anderson Dysphagia Inventory.<sup>17</sup> This 20-item swallowing questionnaire was designed specifically for head and neck cancer patients. A total score is calculated by summing all responses (excluding one global question response), computing the mean score and then multiplying the result by 20. Scores range from 20 to 100. A higher score represents a better self-reported outcome, with 80 points or more suggestive of minimal swallowing difficulties.

#### Statistical analysis

The three measures of swallowing provided continuous data; the treatment groups were therefore compared using one-way analysis of variance. Post-hoc tests were conducted using the Bonferroni correction. Data analyses were performed using the Statistical Package for the Social Sciences version 17.0 software (SPSS Inc, Chicago, Illinois, USA).

## Ethical considerations

Ethical approval was given for the patients receiving non-surgical treatments as part of a separate prospective study. In addition, the outcome measures used in this study are routinely collected, with patient consent, as part of a patient's clinical assessment. These assessments were carried out by speech and language therapists, and students, as part of our ongoing service provision evaluation. Therefore, approval from the local ethics research committee was not required for this particular study.

#### **Results**

In total, 141 patients were eligible for inclusion in the study. Data sets were collected from 124 patients, with one patient completing only the M D Anderson Dysphagia Inventory. Patients who underwent RT alone were treated using three-dimensional conformal RT: 63 Gy in 30 fractions. None of the patients received intensity-modulated RT. Chemoradiotherapy involved sixweekly cycles of cisplatin  $(30-40 \text{ mg/m}^2)$ . Treatment for both the RT plus laryngectomy group and the chemoradiotherapy plus laryngectomy group included salvage laryngectomy, laryngectomy for a dysfunctional larynx, and adjuvant RT or chemoradiotherapy. Overall, eight patients underwent a pharyngolaryngectomy. The groups therefore comprised patients who received different treatments; this was done to ensure adequate group sizes for the purpose of data analysis. Patient details are summarised in Table I.

There were more males than females in the sample (a ratio of 4.6 to 1). A significant age difference was found between the groups (F = 4.325, p < 0.05); the chemoradiotherapy patients were younger (mean = 62 years old) than the laryngectomy patients (mean = 70 years) and the RT patients (mean = 69 years old). There was a significant difference between groups in terms of the duration of disease-free time since the last treatment (F = 12.818, p < 0.05); laryngectomy patients had the longest amount of disease-free time post-treatment

TABLE I PATIENT TREATMENT DATA		
Treatment	Pts ( <i>n</i> )*	Disease-free time <sup>†</sup> (mth (range))
Laryngectomy RT + laryngectomy - Adjuvant RT - Salvage laryngectomy - Laryngectomy for dysfunctional larynx RT CRT CRT + laryngectomy - Adjuvant CRT - Salvage laryngectomy - Laryngectomy for dysfunctional larynx	17 41 20 17 4 26 31 9 2 5 2	84 (7–456) 60 (6–168) 12 (10–41) 12 (6–42) 14 (11–72)

\*Total n = 124 (102 males, 22 females), aged 43–87 years (mean 66 years). <sup>†</sup>Median time post treatment. Pts = patients; mth = months; RT = radiotherapy; CRT = chemoradiotherapy

compared with the chemoradiotherapy group and the RT group (see Table I).

#### Swallow capacity

Norms for swallow capacity are different for males and females.<sup>15</sup> As there were just a small number of females in the total sample, only male patient data were analysed. This revealed a significant difference between the treatment groups (F = 2.7, p = 0.03) (Figure 1). Post-hoc testing indicated that the RT group had significantly better swallow capacity than the chemoradiotherapy plus laryngectomy group (a mean difference of 9.9 ml).

#### Normalcy of diet

Four patients in the sample were nil by mouth (2 per cent) and 40 patients had a full oral diet (32 per



#### FIG. 1

Mean swallow capacity and treatment type error bar plot for males (95 per cent confidence interval). RT = radiotherapy; CRT = chemoradiotherapy; pts = patients cent). There was a significant difference in normalcy of diet scores between groups (F = 5.155, p < 0.01) (Figure 2). Post-hoc tests revealed that the chemoradiotherapy group and the chemoradiotherapy plus laryngectomy group had significantly lower mean scores than the RT group (mean differences of 24 and 36 points respectively). The chemoradiotherapy plus laryngectomy group showed the greatest range of scores and had the smallest number of participants. There was only a small range in scores within both the RT group and the laryngectomy plus RT group, as demonstrated by Figure 2. A large majority of the RT group were managing a full oral diet (84 per cent), with some requiring a drink with meals. Just under half of the chemoradiotherapy patients (45 per cent) were able to manage this level of diet.

#### M D Anderson Dysphagia Inventory

The total score was used to summarise the M D Anderson Dysphagia Inventory results. There was no significant difference between the five treatment groups (F = 2.244, p > 0.05) (Figure 3). Patients in the chemoradiotherapy plus laryngectomy group showed the largest range of scores and the lowest mean score. In order to demonstrate differences in M D Anderson Dysphagia Inventory scores between groups, with a minimum standardised effect size of 0.07 with 90 per cent power, a power analysis estimated that a sample size of 282 patients would be required.

#### **Discussion**

This study aimed to assess swallowing outcomes following different treatments for advanced laryngeal cancer. Three simple clinical swallow tests were administered to patients at least six months after treatment, after which time swallowing function begins to



#### FIG. 2









stabilise following the acute impact of treatment.<sup>4</sup> These tests provide different perspectives on swallowing outcomes, including swallowing performance, dietary restrictions and patient self-report.

The majority of this sample had some degree of swallowing difficulty. Patients treated with chemoradiotherapy combined with laryngectomy consistently had the poorest mean scores across all assessments. There was no clear benefit of surgery or organ preservation treatment with respect to the water swallow test results or patient self-reports. Overall, patients receiving only RT showed better swallowing performance than the other groups in the water swallow test. In addition, the majority of the RT group were able to return to a full oral diet, whereas only half of the chemoradiotherapy group achieved this level of functioning. Patients who had undergone a laryngectomy with or without RT had similar dietary restriction scores. There was a trend for higher M D Anderson Dysphagia Inventory scores in the single modality treatment groups. Approximately half of the patients in the RT group (57 per cent) and in the laryngectomy group (47 per cent) scored themselves over 80 on the M D Anderson Dysphagia Inventory, indicating minimal swallowing problems. This dropped to 35 per cent in the RT plus laryngectomy group and was even lower for patients treated with chemoradiotherapy (19 per cent). There was a large variation in outcomes for patients who underwent a laryngectomy because of a dysfunctional larynx. The patient with the best function in this subgroup had not required a flap reconstruction.

The 100 ml water swallow test is a relatively new test for head and neck cancer patients. Normative data on healthy males of similar ages (with a mean swallow capacity of 18.7 ml),<sup>18</sup> and results of pre-treatment head and neck cancer patients (mean swallow capacity of 16.8 ml),<sup>19</sup> suggest that treatment for advanced laryngeal cancer results in poorer swallowing performance. Swallowing difficulties following chemoradiotherapy have been well described in the literature.<sup>7,10,20–22</sup> These studies have shown biomechanical problems and reduced strength, which affect swallowing performance and safety (i.e. aspiration). Swallowing dysfunction identified on videofluoroscopy is associated with poorer swallow capacity.<sup>23</sup> Therefore, the poorer scores for the chemoradiotherapy patients in the present cohort may be associated with treatment-induced changes to the swallow physiology.

No previous study has reported on the 100 ml water swallow test in total laryngectomy patients. The physiology of swallowing is altered in this group too; the reconstruction of the pharynx results in a smaller lumen, which may account for a reduction in swallow capacity. In addition, lower pharyngeal pressures have been recorded, which reduce bolus propulsion and clearance.<sup>24</sup> However, it has been suggested that surgical patients suffer less from dysphagia than those treated with chemoradiotherapy.25 That particular study did not describe subgroups, so information on laryngectomy as distinguished from organ preservation treatment could not be extrapolated. The swallowing of solid food may be problematic for the total laryngectomy group.<sup>24</sup> This difficulty may not be evident based on the results of a water swallow test, but may be inferred from normalcy of diet scale results.

In line with the findings of the current study, previous papers have reported that RT only patients have minimal dietary restrictions,<sup>10,26,27</sup> and that chemoradiotherapy patients have significantly poorer outcomes in terms of diet.7 Intensity-modulated RT patients are more likely to return to a normal diet than patients receiving other treatments,<sup>28</sup> but research has so far predominantly reported on oropharyngeal cancer patients. For the laryngectomy patients, the addition of RT did not have much impact on diet in the present sample; this supports the findings of our previous study.<sup>29</sup> However, adjuvant RT has been identified as a strong predictor of increased dietary restrictions in a study where patients were divided into three broader groups of diet outcome.<sup>30</sup> Dietary restrictions may not be due to dysphagia alone; dietary changes have also been associated with other issues specific to RT, such as xerostomia and dental clearance.26

In the current study, the chemoradiotherapy group and chemoradiotherapy plus laryngectomy group had lower M D Anderson Dysphagia Inventory scores than the other treatment groups, but there were no statistically significant differences. Gillespie *et al.*<sup>31</sup> also reported a trend for lower M D Anderson Dysphagia Inventory scores in laryngeal cancer patients treated with chemoradiotherapy versus those treated with surgery. In a quality of life study, chemoradiotherapy patients reported more difficulty swallowing than those who had undergone laryngectomy, whereas the laryngectomy group had more problems with communication and shoulder function.<sup>32</sup>

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#### Clinical implications

The outcome measures used in this study are simple, inexpensive and quick to collect in an out-patient review clinic, with excellent patient compliance. These data can provide outcome benchmarks for future comparisons.

This is the first study to report 100 ml water swallow test scores of laryngectomy patients. These results can be used to provide more accurate expectations of eating and drinking abilities following treatment. On the basis of these findings, laryngeal cancer patients referred for RT alone may be given higher expectations regarding the possible impact of treatment on diet, whereas chemoradiotherapy patients will need to be counselled regarding the likelihood of permanent dietary restrictions.

In the current study, patients who underwent chemoradiotherapy plus total laryngectomy had the poorest outcomes. This is perhaps not surprising given that this was the most radical form of treatment. In cases where this treatment is used, there is likely to be a trade-off between long-term functional problems and an increased chance of survival.<sup>13</sup> Eating and drinking ability following laryngectomy for a dysfunctional larynx is hugely variable. Placing expectations on the basis of these results is difficult. Those requiring a flap reconstruction may be particularly vulnerable.

- Surgical and non-surgical treatments for advanced laryngeal cancer have long-term effects on swallowing
- Laryngectomy with chemoradiotherapy is associated with the poorest swallowing outcome
- Laryngectomy with or without radiotherapy, or radiotherapy alone have a minimal effect on diet
- Swallowing outcomes of different head and neck cancer treatments can be assessed using simple clinical tests
- Salvage laryngectomy for chronic dysphagia necessitates careful patient selection and counselling

#### Limitations

Swallowing performance and self-reports are subject to individual variation; therefore, the cross-sectional study design is one limitation of this investigation. A further limitation concerns the fact that patients were at different time points post-treatment, although it had been at least six months since their last intervention. This variation reflected the introduction of chemoradiotherapy as a treatment option for advanced laryngeal cancer. Given the relatively small population and the projected M D Anderson Dysphagia Inventory power calculation, a prospective longitudinal study is likely to be too time-consuming and expensive. Some potential variables were not taken into account as subdividing the groups further would have produced too few patient numbers. For example, swallowing ability in laryngectomy patients may differ according to the type of surgical closure performed or flap reconstruction requirements. These problems indicate the need to collect multi-centre data. Furthermore, patients were not randomly allocated to their treatment group; the type of treatment depended on the nature of the disease and comorbidities.

#### Conclusion

Swallowing outcomes were worse for those patients who received the most radical treatment for advanced laryngeal cancer (i.e. laryngectomy plus chemoradiotherapy). Surgery and organ preservation techniques had no significant swallowing advantage over one another; however, those receiving RT alone were likely to have fewer dietary restrictions and better swallowing performance compared with other treatment groups. Further research on swallowing outcomes following treatment for advanced laryngeal cancer is warranted.

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Address for correspondence: Dr J Patterson, Institute for Health and Society, Newcastle University, The Baddiley-Clark Building, Richardson Road, Newcastle upon Tyne NE2 4AX, UK

E-mail: joanne.patterson@ncl.ac.uk

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