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1   The association between delays in screening for and assessing dysphagia after

2   acute stroke and the risk of stroke-associated pneumonia

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4

1 Abstract

2 Background

3 There is no robust evidence that screening patients with acute stroke for dysphagia reduces the risk

4 of stroke-associated pneumonia (SAP), or of how quickly it should be done after admission. We

5 aimed to identify if delays in bedside dysphagia screening and comprehensive dysphagia

6 assessments by a speech and language therapist (SALT) were associated with patients' risk of SAP.

7 Methods

8 Nationwide registry based prospective cohort study of patients admitted with acute stroke in

9 England and Wales. Multilevel multivariable logistic regression models were fitted, adjusting for

10 patient variables and stroke severity. The exposures were time from 1) admission to bedside

11 dysphagia screen, and 2) admission to comprehensive dysphagia assessment.

12 Results

13 Of 63650 patients admitted with acute stroke, 55838 (88%) had a dysphagia screen and 24542 (39%)

14 a comprehensive dysphagia assessment. Patients with the longest delays in dysphagia screening (4th

15 quartile adjusted OR 1.14, 1.03-1.24) and SALT dysphagia assessment (4th quartile adjusted OR 2.01,

16 1.76-2.30) had a higher risk of SAP. The risk of SAP increased in a dose-response manner with

17 delays in SALT dysphagia assessment, with an absolute increase of pneumonia incidence of 1% per

18 day of delay.

19 Conclusion

20 Delays in screening for and assessing dysphagia after stroke, are associated with higher risk of

21 stroke-associated pneumonia. Since stroke-associated pneumonia is one of the main causes of

22 mortality after acute stroke, early dysphagia assessment may contribute to preventing deaths from

1   acute stroke and could be implemented even in settings without access to high technology specialist

2   stroke care.

3

1 Introduction

2   Stroke-associated pneumonia (SAP) is a common complication of acute stroke, affecting 6-10% of

3   patients[1]. SAP independently increases the risk of mortality and is one of the main causes of death

4   in the first few days and weeks after stroke[2]. It is also associated with worse functional outcomes,

5   longer length of stay and increased healthcare costs[3,4,5,6,7,8]. One of the main risk factors for SAP is

6   dysphagia, which affects 37-55% of patients[9] after stroke. Dysphagia screening using a brief bedside

7   screening tool (such as a water swallow test), and comprehensive clinical assessments of aspiration

8   risk by speech and language therapists (SALT), are therefore performed commonly in stroke care.

9   Typically, all appropriate patients are screened for dysphagia and those in whom dysphagia is

10   suspected go on to receive a comprehensive assessment. Despite being well established in clinical

11   practice, there is however very little evidence of effectiveness for these interventions. Previous

12   studies have generally used weak designs and provided no information to guide recommendations

13   on how quickly dysphagia assessments after stroke should occur[10,11]. Swallowing assessment prior to

14   eating and drinking is recommended in European[12], United States[13] and United Kingdom clinical

15   guidelines but none recommend a specific approach to assessment or treatment of dysphagia.

16   As SAP develops most frequently within the first 7 days of stroke[1], the timing of both dysphagia

17   screening and SALT assessment after admission are likely to be of importance. In England and Wales,

18   dysphagia screening is recommended within 4 h of admission for acute stroke, and comprehensive

19   assessment by a SALT (if required) within 72 h of admission [14] . The aim of this study was to

20   investigate the association between delays in dysphagia screening and SALT assessment and the

21   incidence SAP within the first 7 days after admission. The hypothesis was that delays in these

22   assessments would be associated with an increased incidence of SAP.

23

24

Methods

Data were from the Sentinel Stroke National Audit Programme (SSNAP), the national register of stroke in England and Wales, of patients aged ≥16 years admitted with acute stroke (ischaemic stroke or primary intracerebral haemorrhage) between April 2013 and March 2014. SSNAP is a prospective continuous register with participation from all hospitals admitting adults with stroke in England and Wales and is estimated to include 90-95% of all stroke admissions[15]. Ethical approval of SSNAP was granted by the Ethics and Confidentiality Committee of the National Information Governance Board. Mortality data were obtained through data linkage with the statutory register of deaths. Data linkage was carried out by a third party and the investigators used an anonymised dataset with all patient identifiers removed.

Dysphagia screening was defined as use of a bedside swallow screening test by an appropriately trained clinician (typically a trained nurse). The exact dysphagia screening protocol was not specified by SSNAP. The times from admission to documented dysphagia screen and comprehensive assessment by SALT were recorded to the nearest minute for all patients in whom these were carried out. For patients who had a stroke as an inpatient, the time from stroke onset was used instead of time of admission. All patients without clinical exclusions (e.g. being treated palliatively only) were eligible for dysphagia screening, and eligibility for comprehensive dysphagia assessment was determined by clinical indications, such as a positive dysphagia screen or clinical suspicion of dysphagia. Patients (n=965) admitted directly to an intensive care unit (ICU) on admission were excluded from the primary analysis, since most of these patients would have been intubated.

SAP was defined as the administration of antibiotics for a new clinical diagnosis of pneumonia in the first seven days after admission and was determined by the treating physician.

Statistical analysis

1 The adjusted odds of SAP were estimated by fitting multivariable logistic regression models. Time

2 from admission to dysphagia screen and SALT dysphagia assessment were analysed both as

3 continuous variables and by division into quartiles. When included as a continuous variable they

4 were fitted as restricted cubic regression splines using the POSTRCSPLINE module[16]. Spline

5 coefficients cannot be interpreted directly and so the models were displayed graphically, showing

6 the modelled association (and 95% confidence interval) between time to dysphagia screen or

7 comprehensive dysphagia assessment and estimated adjusted SAP incidence. Multilevel

8 multivariable logistic regression models were also fitted using quartiles of these times to enable

9 quantification of the study results into odds ratios and also to account for clustering at the hospital

10 level. These models were specified as two level models with hospital level random intercepts.

11 All models included age, sex, stroke subtype (ischaemic, primary intracererbral haemorrhage, or

12 undetermined), pre stroke functional level using the modified Rankin Scale (mRS), place of stroke

13 (out of hospital or inpatient), vascular comorbidity (heart failure, hypertension, atrial fibrillation,

14 diabetes mellitus, previous stroke or transient ischaemic attack) and either NIH Stroke Scale (NIHSS)

15 or level of consciousness on admission.

16 Data were complete for all data items apart from the NIHSS on admission, which was available for

17 73% of patients. Models were therefore also fitted using level of consciousness on admission as a

18 proxy for severity (available for 100% of patients) and the results of the models compared to explore

19 the effect of these missing data on the results.

20 We carried out several sensitivity analyses. Firstly, competing risk from early mortality was explored

21 by excluding patients dying or starting palliative care in the first 3 days. Secondly, models were fitted

22 including a variable indicating change (increase, no change, decrease) in the level of consciousness in

23 the first seven days after stroke, to explore the possible confounding effect of changing

24 consciousness level. Thirdly, ICU patients excluded in the main analysis were included in a complete

25 data set analysis. Fourthly, time from admission to stroke unit admission was included in the models

1   as a possible confounder. Finally, we fitted models of 30 day all cause mortality, excluding patients

2   dying or starting palliative care in the first 3 days (on the grounds that death in these latter patients

3   is more likely due directly to brain injury from the stroke rather than stroke associated pneumonia).

4   These models explored whether delays in dysphagia screening and assessment were associated with

5   mortality after stroke.

Results

There were 63650 patients with acute stroke included in the cohort, admitted to 199 hospitals. Of these, 55838 (87.7%) had a dysphagia screen, and 24542 (38.6%) proceeded to a comprehensive assessment. The characteristics of the whole cohort, and the subgroups of patients according to receipt of dysphagia screening and comprehensive dysphagia assessment are described in Table 1. Patients in whom a dysphagia screen was not performed had a greater incidence of inpatient stroke and primary intracerebral haemorrhage, lower level of consciousness on admission and were less likely to have NIHSS on admission completed.

	Cohort	Dysphagia screening not performed	Dysphagia screening performed	Comprehensive dysphagia assessment
n (%)	*63650*	*7812 (12.3)*	*55838 (87.8)*	*24542 (38.6)*
Median age(IQR)	77 (67-85)	80 (70-86)	77 (67-85)	80 (70-87)
Female (n, %)	32054 (50.4)	4264 (54.5)	27790 (49.8)	13160 (53.6)
Stroke type (n, %)				
Ischaemic	56167 (88.2)	5948 (76.1)	50219 (91.0)	21751 (89.6)
Primary intracerebral haemorrhage	6575 (10.3)	1592 (20.4)	4983 (8.9)	2523 (10.4)
Undetermined	908 (1.4)	272 (3.5)	636 (1.1)	268 (1.1)
Inpatient stroke (n, %)	3155 (5.0)	974 (12.5)	2181 (3.9)	1599 (6.5)
Pre-Stroke mRS (n,%)				
0	36208 (57.9)	3808 (48.8)	32400 (58.0)	12174 (49.6)
1	9726 (15.3)	1109 (14.2)	8617 (15.4)	3919 (16.0)
2	6036 (9.5)	851 (10.9)	5185 (9.3)	2614 (10.7)
3	6708 (10.5)	1073 (13.7)	5635 (10.1)	3163 (12.9)
4	3734 (5.9)	670 (8.6)	3064 (5.5)	1988 (8.1)
5	1248 (2.0)	301 (3.9)	937 (1.7)	684 (2.8)
Admission NIHSS complete (n,%)	46447 (73.0)	3792 (48.4)	42655 (76.4)	17041 (69.4)
Median admission NIHSS (IQR)	4 (2-9)	5 (2-19)	4 (2-9)	7 (3-14)
Level of consciousness on admission				
0 (Alert)	53433 (84.0)	4961 (63.5)	48472 (86.8)	19156 (78.1)
1 (Not alert: Responds to voice)	6032 (9.5)	1069 (13.7)	4963 (8.9)	3697 (15.1)
2 (Not alert: Responds to pain)	2498 (3.9)	828 (10.6)	1670 (3.0)	1267 (5.2)
3 (Totally unresponsive)	1687 (2.7)	954 (12.2)	733 (1.3)	422 (1.7)
Co-morbidity (n,%)				
Heart failure	3463 (5.4)	491 (6.3)	2972 (5.3)	1625 (6.6)
Hypertension	34212 (53.9)	3930 (50.3)	30382 (54.4)	13323 (54.3)
Atrial fibrillation	13159 (20.7)	1801 (23.1)	11358 (20.3)	5929 (24.2)
Diabetes mellitus	12372 (19.4)	1482 (18.9)	10890 (19.5)	4720 (19.2)
Previous stroke/TIA	17626 (27.7)	2144 (27.4)	15482 (27.7)	7109 (29.0)
Time from onset to admission (n, %)				
Unknown (e.g. wake up stroke)	24668 (38.8)	4233 (54.2)	20435 (36.6)	9631 (39.2)
0-179 minutes	21504 (33.8)	2113 (27.1)	19391 (34.7)	9086 (37.0)
180-359 minutes	6144 (9.7)	539 (6.9)	5605 (10.0)	2289 (85.6)
360+ minutes	11334 (17.8)	927 (11.9)	10407 (18.6)	3536 (14.4)
Thrombolysis (n, %)	7087 (11.1)	417 (5.3)	6670 (12.0)	3415 (3.1)
SAP (n,%)	5533 (8.7)	1077 (13.8)	4456 (8.0)	3592 (14.6)
30 day mortality (n,%)	8397 (13.2)	2701 (34.6)	5696 (10.2)	3599 (14.7)

1

2   **Table 1**. *Characteristics of the study cohort*

1

2   The overall incidence of SAP was 8.7%. SAP incidence was highest in the dysphagic group referred

3   for comprehensive dysphagia assessment (14.6%). Thirty day mortality was 13.2% overall, 10.2% in

4   patients screened for dysphagia, 14.7% in patients referred for SALT assessment, and 34.6% in

5   patients in whom a dysphagia screen was not carried out.

6   The median time from admission to dysphagia screening was 2.9 hours (IQR 1.3-5.7 hours) and for

7   comprehensive dysphagia assessment was 22.9 hours (IQR 6.2-49.4 hours). In unadjusted analyses

8   there was a strong association between time from admission to dysphagia screen and incidence of

9   SAP, rising from 7-8% from 0-8 hours and increasing to 15% by 72 hours after admission. Although

10   the association was attenuated after adjusting for patient characteristics, there was still a modest

11   association (equating to approximately 1% absolute increase in the incidence of SAP) between

12   delays in dysphagia screening and incidence of SAP (Figure 1). After adjustment, patients in the

13   fourth quartile (i.e. those with the longest delays in dysphagia screening) had 36% higher odds of

14   SAP compared to those in the first quartile (aOR 1.36, 1.20-1.53) (Table 2).

15

		Time (mins)	OR	95% CI	p
Univariable (n=55838)	1st quartile	0-79	REF		
	2nd quartile	80-176	0.89	0.81-0.98	0.016
	3rd quartile	177-344	0.85	0.77-0.94	0.001
	4th quartile	≥345	1.33	1.21-1.46	<0.0001
Multivariable, including NIHSS (n=42655)	1st quartile	0-79	REF		
	2nd quartile	80-176	0.94	0.83-1.05	0.27
	3rd quartile	177-344	1.06	0.94-1.20	0.36
	4th quartile	≥345	1.36	1.20-1.53	<0.0001
Multivariable, including level of consciousness (n=55838)	1st quartile	0-79	REF		
	2nd quartile	80-176	0.92	0.83-1.01	0.08
	3rd quartile	177-344	0.89	0.81-0.99	0.03
	4th quartile	≥345	1.14	1.03-1.24	0.008

1

2 **Table 2.** *Odds ratio for SAP in univariable and multivariable models of time from admission to*
3 *dysphagia screening.* All multivariable models were also adjusted for age, sex, stroke type, pre-
4 stroke functional level, place of stroke and comorbidity, and measure of stroke severity (NIHSS or
5 level of consciousness)

6

7 There was a strong relationship between delays in comprehensive dysphagia assessment and

8 incidence of SAP, and delays in comprehensive dysphagia assessment were associated with an

9 absolute increase in the risk of SAP of 3% over the first 24 hours (Figure 2). Delays in SALT dysphagia

10 assessment beyond 24 hours were associated with an additional 4% absolute increase in the

11 incidence of SAP (approximately 3-fold increase in the relative incidence). Patients in the slowest

12 quartile had 1.98 (1.67-2.35) the odds of SAP compared to patients receiving the quickest SALT

13 dysphagia assessments (Table 3). Findings were similar in the sensitivity analyses (Supplementary

14 material). The secondary analysis of 30 day mortality broadly supported these findings: there was

15 very weak evidence that delays in dysphagia assessment were associated with an increase in 30 day

16 mortality (aOR 1.14, 0.99-1.30 in the slowest quarter). There was moderately strong evidence that

17 delays in comprehensive SLT assessment were associated with an increase in mortality risk in the

18 second (aOR 1.22, 1.02-1.47), third (aOR 1.55, 1.29-1.85) and fourth (aOR 1.35, 1.12-1.63) quarters

1   of time to assessment respectively. Unlike pneumonia, a dose-response relationship was not

2   demonstrated for the association with mortality (Supplementary material)

1

		Time (mins)	OR	95% CI	p
Univariable (n=24542)	1st quartile	0-369	REF		
	2nd quartile	370-1371	1.53	1.34-1.74	<0.0001
	3rd quartile	1372-2961	1.95	1.71-2.22	<0.0001
	4th quartile	≥2962	2.65	2.33-3.01	<0.0001
Multivariable, including NIHSS (n=17041)	1st quartile	0-369	REF		
	2nd quartile	370-1371	1.35	1.15-1.60	<0.0001
	3rd quartile	1372-2961	1.61	1.37-1.91	<0.0001
	4th quartile	≥2962	1.98	1.67-2.35	<0.0001
Multivariable, including level of consciousness (n=24542)	1st quartile	0-369	REF		
	2nd quartile	370-1371	1.40	1.22-1.60	<0.0001
	3rd quartile	1372-2961	1.60	1.41-1.84	<0.0001
	4th quartile	≥2962	2.01	1.76-2.30	<0.0001

2

3   **Table 3.** *Odds ratio for SAP in univariable and multivariable models of time from admission to SALT*
4   *dysphagia assessment.* All multivariable models were also adjusted for age, sex, stroke type, pre-
5   stroke functional level, place of stroke and comorbidity, and measure of stroke severity (NIHSS or
6   level of consciousness)

7

8

1 Discussion

2 In this national cohort of unselected stroke patients, we found that there was evidence of a modest

3 association between delays in performing dysphagia screening and the risk of SAP. There was

4 stronger evidence for an association between the risk of SAP and delays in carrying out a

5 comprehensive dysphagia assessment. Although limited by the risk of residual confounding, these

6 findings provide the first evidence from a large multicentre cohort that prompt dysphagia screening

7 and comprehensive dysphagia assessment stroke are associated with clinically significant reductions

8 in the risk of SAP, one of the principal causes of early death after stroke.

9 Detecting dysphagia through the use of bedside screening assessments and comprehensive

10 dysphagia assessments carried out by a SALT is widely recommended in clinical guidelines[12-14].

11 However, these recommendations are largely based on consensus and there is little direct evidence

12 for dysphagia screening or assessment after stroke[17]. Previous studies have been limited to

13 ecological studies demonstrating an association between site level rates of screening assessment

14 and SAP rates after stroke[8,10,11]. Several observational studies have described an association between

15 dysphagia screening at any time after stroke and reduced post stroke mortality[10,18,19,20], and

16 dysphagia screening was a component of a stroke care bundle found in a cluster randomised

17 controlled trial to reduce death and dependency after stroke[21]. By contrast, an analysis of the "Get

18 with the Guidelines – Stroke" registry data from the USA found that dysphagia screening was

19 associated with a higher risk of SAP, although the results suggest that confounding by stroke

20 severity contributed to the observed association[22]. There is no current evidence of how quickly

21 dysphagia assessment should occur after stroke or good quality evidence of whether dysphagia

22 assessment reduces the risk of SAP. As a result of this lack of evidence, dysphagia screening has been

23 dropped from the list of stroke quality indicators used in the USA[23], although they remain part of

24 quality indicators used in the UK[24].

1 If our findings represent causal effects, then they imply that dysphagia screening and assessment is

2 effective in reducing the risk of SAP. Since SAP is one of the main causes of death in acute stroke,

3 reducing the risk of SAP would be expected to lead to reduced mortality after stroke. The secondary

4 analyses of mortality in this study provide supporting evidence that this might be the case. We

5 would however emphasise caution in interpreting the mortality findings - there are many causes of

6 death in acute stroke and so reductions in SAP will only prevent a proportion of deaths (in keeping

7 with the reduced effect sizes for mortality we observed) and we did not observe a dose-response

8 relationship between delays in comprehensive dysphagia assessment and mortality, suggesting that

9 there might be additional confounding or bias not accounted for in the analysis.

10 There are several possible mechanisms for why delays in dysphagia assessment might lead to an

11 increased risk of SAP and further studies would be required to test these hypotheses and confirm

12 (or refute) a causal relationship: early screening may reduce the risk of inappropriate administration

13 of oral fluid or food, prompt measures to reduce aspiration risk through positioning, nursing care

14 and appropriate feeding strategies, and avoid unnecessary nasogastric tube insertions. As well as

15 exploring mechanisms, further research might usefully also explore organisational aspects of

16 dysphagia assessment, such as the use of specific assessment and treatment protocols and the

17 relationship between specialist SALT provision on stroke units and patient outcomes.

18 These data are strengthened by being drawn from a national register of unselected patients,

19 reducing the risk of selection bias. Similarly, the study used clinical rather than administrative data,

20 providing more detail than would be available from routine administrative data alone. There are

21 however a number of limitations. Firstly SAP was not defined by specific criteria but was based on

22 the judgement of the treating physician, and we did not have information on the date of diagnosis of

23 SAP and whether it occurred before or after dysphagia screening and assessment. Nonetheless, the

24 overall rate of SAP observed in this study is consistent with other studies, suggesting that differences

25 in ascertainment between centres was not a significant source of bias. In addition, although data

completeness was high, NIHSS data were not available for one quarter of patients. We used level of consciousness as a proxy for this and found that the findings were similar, but having complete data on stroke severity may have strengthened the study. The dataset lacked information on the nature of the bedside dysphagia screening tools used, the details of the comprehensive assessment (e.g. videofluroscopy or fibre-optic evaluation of swallowing), and the results of these assessments. Further studies should aim to capture in more detail the components of these interventions. The main limitation of this study is the risk of residual confounding. The hypothesis that early dysphagia screening and SALT assessment reduce the risk of SAP could be tested in a cluster randomised controlled trial of a protocol of expedited comprehensive dysphagia assessment and this would help guide clinical practice in an important area of stroke care which currently has a poor evidence base.

Implementing faster dysphagia assessments in clinical practice is principally a matter of training healthcare professionals appropriately and in most instances does not require expensive medical equipment. Dysphagia screening has been identified by the World Stroke Organisation as being achievable even in health economies with the lowest level of resources[25]; ensuring that all stroke patients receive rapid dysphagia assessments could therefore be a part of global efforts to improve the outcomes of acute stroke, even in settings without advanced specialist stroke care.

Summary

Delays in screening for dysphagia and carrying out SALT dysphagia assessments after stroke are associated with an increased risk of SAP. This hypothesis that expedited dysphagia screening and assessments reduce the risk of SAP would be testable in an appropriately designed trial or controlled evaluation. In the meantime, these findings suggest that reducing delays in screening and assessing for dysphagia in people with acute stroke should be a focus of quality improvement in stroke care.

Acknowledgements

Funding

Contributions

BDB - study design, analysis, writing

CJS - study design, writing

GCC - writing

PE - writing

MJ - writing

LP - analysis, writing

PJT - writing

CDAW - writing

AGR - writing

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1   **Figure 1**. *Modelled relationship between estimated incidence of SAP in the first seven days of*

2   *admission and time to dysphagia screening.* (A): Multivariable model including NIHSS. (B):

3   Multivariable model including level on consciousness

4   **Figure 2**. *Modelled relationship between estimated incidence of SAP in the first seven days of*

5   *admission and the time to SALT dysphagia assessment.* (A): Multivariable model including NIHSS.

6   (B): Multivariable model including level on consciousness