



Barriers to an early switch from intravenous to oral antibiotic therapy in hospitalised patients with CAP

Madelon F. Engel*, Douwe F. Postma*, Marlies E.J.L. Hulscher[#], Ferdinand Teding van Berkhout[†], Marielle H. Emmelot-Vonk⁺, Sanjay Sankatsing[§], Carlo A.J.M. Gaillard^{†,**,†}, Anke H.W. Bruns^{*,†}, Andy I.M. Hoepelman* and Jan Jelrik Oosterheert*

ABSTRACT: Do physicians apply an early-switch strategy (from intravenous to oral antibiotics) in clinically stable patients hospitalised with community-acquired pneumonia (CAP)? If not, why not?

In a multicentre prospective cohort study, adult patients admitted for *i.v.* CAP treatment were included. On day 3 of antibiotic treatment, clinical stability was assessed and treating resident physicians were interviewed on their switch strategies. Additionally, treating physicians were interviewed to evaluate their knowledge of and adherence to guideline advice.

149 (92%) out of 162 patients were included and 97 (91%) out of 107 physicians were interviewed. A switch to oral antibiotics was possible in 68 (46%) out of 149 patients on day 3 of treatment but not performed in 27 (40%) out of 68. Patient factors delaying the switch were high CURB-65 (confusion of new onset, urea >7 mmol·L⁻¹, respiratory rate of ≥30 breaths·min⁻¹, blood pressure <90 mmHg or diastolic blood pressure ≤60 mmHg, and age ≥65 yrs) score (on admission) (p=0.04) and oxygen treatment (p=0.04), high temperature (p=0.00) and high respiration rate (p=0.04) (day 3). Physicians' barriers to an early switch in clinically stable patients included misconceptions (26 (55%) out of 47), practical considerations (13 (28%) out of 47) and organisational factors (eight (17%) out of 47). Strikingly, 91 (94%) out of 97 interviewed physicians were not aware of guideline advice.

The switch from *i.v.* to oral antibiotics is often unnecessarily delayed in patients hospitalised with CAP due to different types of barriers.

KEYWORDS: Antibiotic treatment, community-acquired pneumonia

Community-acquired pneumonia (CAP) is the leading cause of death from infectious diseases and a substantial burden on healthcare resources in western countries [1]. Hospital admission is usually required because of intravenous antibiotic treatment, which ensures optimal tissue penetration. The *i.v.* therapy is often continued until patients have definitely overcome the infection [2]. A recent meta-analysis showed that an early switch to oral therapy in clinically stable patients on day 2–4 of hospital admission, as opposed to prolonged *i.v.* treatment, resulted in an estimated decrease in length of hospital stay of 3.4 days and fewer medication side-effects [2]. The treatment effect, the number of recurrent infections and mortality

were comparable. Even in severe forms of CAP, an early-switch strategy is usually possible on day 3 of admission [3]. Large-scale implementation of early-switch strategies would lead to a substantial reduction in healthcare costs [3, 4]. Therefore, (inter-)national guidelines advocate an early switch to oral treatment for hospitalised CAP patients [1, 5, 6].

Despite its obvious and proven benefits, an early switch to oral therapy has not been implemented in routine clinical practice for all admitted CAP patients to date. It has been shown that healthcare staff experience barriers towards implementation of an early-switch strategy in lower respiratory tract infections due to several factors, including a

AFFILIATIONS

*Dept of Internal Medicine and Infectious Diseases, Utrecht University Medical Centre, †Dept of Pulmonary Diseases, Utrecht University Medical Centre, †Dept of Geriatric Diseases, Utrecht University Medical Centre, and †Dept of Internal Medicine, Diaconessen Hospital, Utrecht, #Scientific Institute for Quality of Healthcare, Radboud University Medical Centre, Nijmegen, †Dept of Internal Medicine, Meander Medical Centre, Amersfoort, and **Dept of Nephrology, VU University Medical Centre, Amsterdam, The Netherlands.

CORRESPONDENCE

M.F. Engel
Utrecht University Medical Centre
Dept of Internal Medicine and Infectious Disease
Internal mail F02.126
PO Box 85500
Utrecht
The Netherlands
E-mail: m.f.engel-2@umcutrecht.nl

Received:

Feb 20 2012
Accepted after revision:
April 13 2012
First published online:
May 31 2012

For editorial comments see page 5.

This article has supplementary material available from www.erj.ersjournals.com

perceived lack of clear guideline recommendations and a lack of outcome expectancy [7, 8]. The level of adherence to an early-switch strategy (*i.e.* patients being switched to oral agents as soon as clinical stability is reached) in CAP is 58% on average, but varies greatly between different hospitals (22–94%) [9]. This suggests considerable room for improvement of the adherence to an early-switch strategy.

Merely publishing results of randomised controlled trials and issuing of guidelines seems to be insufficient for implementation of an early-switch strategy in daily practice [10]. Generally, implementation strategies are ideally preceded by a proper analysis of barriers to implementation on different levels [11–13]. Therefore, we aimed to evaluate whether physicians base the conversion to oral treatment on guideline advice, what patient and physician characteristics influence the timing of the switch to oral treatment and whether physicians perceive barriers to an early-switch strategy. Identification of these barriers may form the basis of a targeted implementation strategy.

METHODS

Design and setting

In a prospective, observational, multicentre study, we evaluated the timing of the switch (*i.e.* cessation of *i.v.* antibiotics with or without conversion to oral antibiotics) and possible perceived barriers to an early-switch strategy in CAP patients admitted to internal and pulmonary medicine wards. Two teaching hospitals in the Netherlands (the Diaconessen Hospital, Utrecht, with 627 beds and the Meander Medical Centre, Amersfoort, with 982 beds) and one university hospital (Utrecht University Medical Centre, Utrecht, with 1,042 beds) participated in the study. The study protocol was approved by the local medical ethics committees. As the objective of this study was to describe physician behaviour and the main measurements were performed through physician interviews, obtaining patient informed consent was not required.

Patients and procedures

All consecutive adult patients admitted to one of the participating hospitals for *i.v.* CAP treatment were eligible for analysis (online supplementary appendix I). Patients were excluded if they were admitted to the intensive care unit before the switch to oral antibiotics was made or if they had a history of cystic fibrosis or lung transplantation, because prolonged *i.v.* treatment is often necessary in these patients. Patients with co-infections that needed immediate *i.v.* antibiotic treatment were excluded because of presumed interference with the duration of *i.v.* therapy for CAP.

In each participating centre, an investigator identified cases eligible for inclusion by screening the admission lists twice weekly and by attending the daily reports where new admissions are discussed. Usually, in the Netherlands, CAP patients needing hospitalisation are admitted to one of the wards immediately after initial evaluation in the emergency department, usually within 8 h of presentation. After inclusion, severity scores (CURB-65 (confusion of new onset, urea >7 mmol·L⁻¹, respiratory rate of ≥ 30 breaths·min⁻¹, blood pressure <90 mmHg or diastolic blood pressure ≤ 60 mmHg, and age ≥ 65 yrs) score [14] and the Pneumonia Severity Index (PSI)/Fine score [15]) and the antibiotics administered were recorded on the day of admission (day 0). As in routine

practice, calculation of the CURB-65 and Fine scores was based on the available data only.

At five time-points during follow-up (day 3, 6, 14, 21 and 28 after admission), clinical patient data and the route of antibiotic administration (*i.v.*/orally) were recorded. Clinical stability was assessed and defined as: temperature $<37.8^{\circ}\text{C}$, oxygen saturation $>92\%$ without additional administration of oxygen, stable blood pressure without the need for saline infusion or vasopressive medication, cardiac frequency <100 beats·min⁻¹, respiratory rate <25 breaths·min⁻¹ and absence of mental confusion that arose after the onset of infection [16–18]. If patients were able to swallow and were free of nausea or vomiting they were marked as “able to take oral medication”. Parameters not recorded in the medical chart were assumed to be normal. Recording of clinical data ceased after a patient was switched to oral antibiotics, was discharged or died. Visits to the outpatient clinic, re-admissions and mortality were noted retrospectively 28 days after admission.

Physician interviews

Residents or senior students functioning as such, treating included patients on the third day of admission were labelled as “the treating physician” and were asked to fill out a case-specific questionnaire or were interviewed using this questionnaire (online supplementary appendix II). Generally, the resident treating a patient admitted to a ward on a specific time-point is responsible for that patient during the entire admission. We did not record if a change in treating residents occurred during the treatment of one of the included patients because in practice this seldom occurs (*e.g.* when residents rotate after 6 months on a specific ward). Treating physicians were informed about the study purpose in general terms, stating that the aim was to record the physicians’ motivation to choose oral or *i.v.* antibiotic administration. Physicians were asked whether they considered the patient to be clinically stable and if they could identify factors that influenced their choice for oral or *i.v.* administration of antibiotics. During the interview the researcher pointed out that factors on different levels might play a role, such as at the organisational or patient level, in order to stimulate the physicians to provide complete answers. Open-ended questions were used, in order to provide the opportunity to identify various barriers. Because the residents received daily supervision by medical specialists, their answers were considered a reflection of the specialist’s opinion.

After the inclusion period, senior medical students, residents and supervising medical specialists involved in the treatment of included patients were invited to participate in a semi-structured, in-depth interview using a physician-specific questionnaire (online supplementary appendix III). This interview addressed the physicians’ general knowledge, clinical experience and opinion on the optimal timing of the switch to oral antibiotics in CAP patients. If the provided answers seemed incomplete, the physicians were encouraged to elucidate their answers. All interviews were conducted by the same investigator (M.F. Engel) in a standardised manner.

Sample size calculation, outcome measurement and data analysis

The primary outcome was the number of patients adequately switched to oral therapy on day 3 of treatment. Based on

previous studies, a baseline adherence to an early-switch therapy in ~58% of cases can be expected [8, 9]. According to the Wilson's score interval, a sample size of 140 patients would produce a two-sided 95% confidence interval with a width equal to 0.16 when the sample proportion is 0.58. The size of the study population was not suitable for a multilevel analysis; instead, physician and hospital characteristics were evaluated at the patient level using logistic regression analysis.

Secondary outcomes included the patient and physician factors associated with an early-switch strategy. Factors possibly associated with continuous *i.v.* treatment on day 3 were assessed in univariate analysis. Ratio variables were evaluated as such, and not per category (*e.g.* Fine score risk classes) to minimise the loss of information. Variables with a *p*-value <0.05 and with <15% missing values were used for multivariate analysis using the forward likelihood ratio model. The goodness of fit of the model was tested with the Hosmer-Lemeshow test. All statistical analyses were performed using SPSS version 15.0 (SPSS Inc., Chicago, IL, USA).

Remaining secondary outcomes were barriers to an early-switch strategy and the optimal time to switch in CAP patients in general, as perceived by the treating physicians. For these qualitative data, the frequency of specific answers provided during the interviews was calculated by counting the number of times a specific answer was provided either per case or per physician.

RESULTS

Patient and physician characteristics

Between October 2010 and May 2011, 162 patients admitted to one of the three participating hospitals were enrolled. Of the 13 (8%) of patients who were excluded, one was excluded because he/she was transferred to another hospital, six had missing data and six were excluded for other reasons. Of the 149 included patients, one patient died before day 3; this left 148 cases for analysis (fig. 1). Overall, the mean \pm SD age was 67 ± 17 yrs and 71 (48%) were female. 95 (64%) patients had one or more comorbidities; the most frequent co-morbidities were lung disease (*n*=49; 33%) and malignancy (*n*=18; 12%). The mean PSI score on admission was 88 ± 32 (table 1). All included patients were treated on the ward to which they were initially admitted on for at least 3 days.

107 physicians were involved in the treatment of the 149 included patients and were invited for a physician-specific interview; 97 (91%) were interviewed (fig. 1). Out of 10 (9%) physicians who were excluded, four refused to participate, two were no longer working at the hospital and four were on long-term leave. Reasons for refusing participation were insufficient time in three cases and unclear in one case. The interviewed physicians consisted of 35 (36%) specialists with 417.5 cumulative years of experience as a medical specialist, 53 (55%) residents with 142.5 cumulative years of clinical experience and nine (9%) senior students. The physicians had gained their experience in >30 Dutch centres and two foreign hospitals.

Timing of the switch

Overall, *i.v.* antibiotics were continued for 4.7 ± 2.8 days and the length of hospital stay was 8.2 ± 5.5 days (table 2). In 68 (46%) out of 148 cases, a switch to oral agents was possible

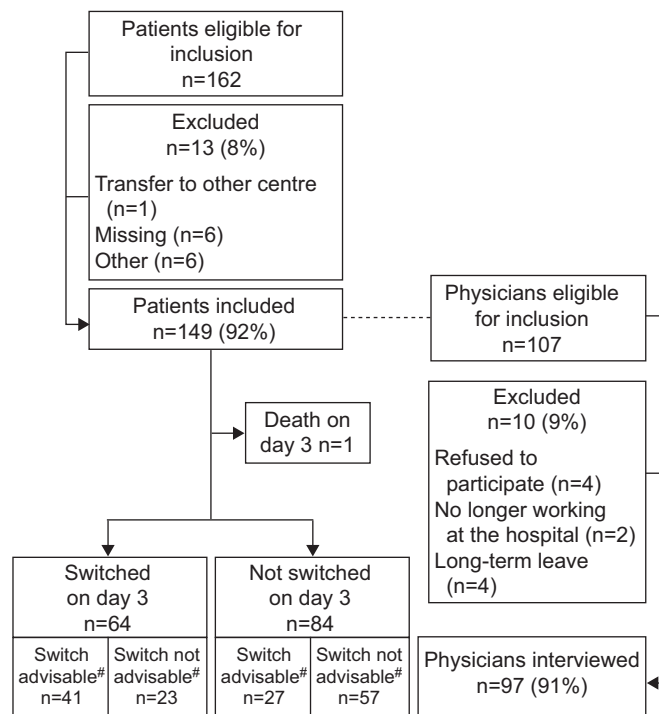


FIGURE 1. Flow chart of patients and physicians during the study. #: patients are clinically stable and able to take medication orally.

based on the predefined clinical criteria (figs 1 and 2). However, in 27 (40%) out of 68 cases, *i.v.* antibiotics were still administered on and beyond day 3 (fig. 1). The case mix on respiratory and nonrespiratory wards was comparable, aside from the higher number of comorbid lung diseases on the respiratory wards (41 (51%) out of 80 *versus* eight (12%) out of 68; *p*<0.00). The percentage of patients switched in a timely fashion was also comparable between these ward types (11 (36%) out of 31 *versus* 16 (43%) out of 37; *p*=0.52). Furthermore, the timing of the switch in the first and second half of the study (4.9 ± 3.2 *versus* 4.5 ± 2.5 ; *p*=0.38) was comparable, which makes a learning effect unlikely.

Adherence to guideline recommendations

Strikingly, 91 (94%) of the 97 treating physicians were not aware of the existence of any clear guidelines on the adequate timing of the switch. Indeed, participating hospitals did not provide these guidelines through antibiotic booklets or their intranet, but there are clear (inter-)national guidelines on this subject accessible online [1]. In the physician-specific interviews, some reported switch criteria that matched the criteria used in this study, such as the absence of fever for 36 (37%) physicians, declining/no need for oxygen for 24 (25%) and haemodynamic stability for 21 (22%) (table 3). Additional factors, not matching our criteria, were a decrease in C-reactive protein for 47 (48%) physicians and signs of clinical improvement (*i.e.* physician's clinical judgment or not specified) for 36 (37%). Notably, 16 (16%) physicians mentioned "the patient feeling better" as a criterion to switch to oral antibiotics.

Of the 116 patients marked as clinically stable by the resident (but not yet discharged) on day 3, 59 (51%) did not meet the

TABLE 1 Patient characteristics[#]

	Total	Timing switch		p-value (95% CI)
		On or before day 3	After day 3	
Demographic data				
Subjects	148 (100)	64 (43)	84 (57)	
Age yrs	67.0 ± 17.3	65.7 ± 19.3	68.0 ± 15.6	0.423 (-7.974–3.364)
Females	71 (48.0)	29 (45.3)	42 (50.0)	0.572
Comorbidity				
Malignancy [†]	18 (12.2)	6 (9.4)	12 (14.3)	0.365
Liver disease [†]	2 (1.4)	1 (1.6)	1 (1.2)	1.000
Congestive heart failure [†]	17 (11.5)	8 (12.5)	8 (10.7)	0.736
Cerebrovascular disease [†]	15 (10.1)	6 (9.4)	9 (10.7)	0.789
Kidney disease [†]	9 (6.1)	6 (9.4)	3 (3.6)	0.143
Lung disease	49 (33.1)	23 (35.9)	26 (31.0)	0.523
≥ 1 comorbidity	95 (64.2)	45 (70.3)	50 (59.5)	0.175
Severity scores and symptoms at presentation⁺				
CURB-65 at presentation	1.5 ± 0.5	1.4 ± 1.1	1.8 ± 1.2	0.046 (-0.759– -0.007)
CURB-65 >2	34 (23.0)	12 (18.8)	22 (26.2)	0.286
Fine score at presentation	88.4 ± 32.0	83.5 ± 27.9	92.2 ± 34.5	0.101 (119.174–1.714)
Fine score IV	57 (38.5)	27 (42.2)	30 (35.7)	0.423
Fine score V	11 (7.4)	0	11 (13.1)	0.003
Cough	107 (73.3)	47 (74.6)	60 (72.3)	0.754
Sputum production	63 (45.7)	30 (51.7)	33 (41.3)	0.223
Dyspnoea	112 (78.9)	51 (81.0)	61 (77.2)	0.588
Temperature >38°C or <36°C	92 (62.2)	39 (60.9)	53 (63.1)	0.789
Pneumonia on auscultation	110 (74.8)	54 (84.4)	56 (67.5)	0.019
Leukocytosis	102 (69.4)	44 (68.8)	58 (69.9)	0.883
CRP > 30 mg·L ⁻¹	133 (89.9)	55 (85.9)	78 (92.9)	0.167

Data are presented as n (%) or mean ± SD, unless otherwise stated. Independent-sample t-tests were used for ratio variables and the Chi-squared test or Fisher's exact test for nominal variables. CURB-65: confusion of new onset, urea >7 mmol·L⁻¹, respiratory rate of ≥30 breaths·min⁻¹, blood pressure <90 mmHg or diastolic blood pressure ≤60 mmHg, and age ≥65 yrs; CRP: C-reactive protein. [#]: n=148; [†]: as defined in the Fine score; ⁺: missing data were not used in the calculation. Bold indicates statistical significance.

objective criteria derived from research findings. For example, some patients with high cardiac frequency (up to 182 beats·min⁻¹), fever (up to 39.2°C) or still needing oxygen were marked as clinically stable. In contrast, some physicians noted that they did not apply an early-switch strategy in specific patients because of "fever" (36.7°C) or "tachycardia" (72 beats·min⁻¹).

Barriers to an early-switch strategy

To evaluate factors that influence the decision to convert to oral antibiotics, characteristics of the 84 (56%) patients still on *i.v.* antibiotics on day 3 were compared with the 65 (44%) patients on oral antibiotics (tables 1 and 2). Variables univariately associated with a prolonged duration of *i.v.* administration of antibiotics ($p < 0.05$) were abnormalities on auscultation suggesting pneumonia ($p = 0.02$) and a high CURB-65 score ($p = 0.046$; 95% CI -0.76– -0.017) on admission; admission to a secondary care hospital, as opposed to a university medical centre ($p = 0.03$); a high respiratory frequency ($p = 0.04$; 95% CI -5.38– -0.08); high temperature ($p = 0.00$; 95% CI -0.70– -0.21); oxygen administration ($p = 0.01$) and clinical instability according to the resident ($p = 0.00$) on day 3. The respiratory frequency was not recorded in 103 (69%) patients, therefore

this variable was excluded, and the remaining variables were used in multivariate analysis. In this analysis, nine (6%) out of 148 patients were excluded because they were discharged on day 3 before measurements could be taken, and an additional 10 (7%) had missing values because of incomplete recording. Variables that tested significant in multivariate analysis were high temperature (odds ratio 2.9, 95% CI 1.5–5.4) and oxygen administration (OR 2.5, 95% CI 1.8–5.5) on day 3.

As shown, patients still on *i.v.* antibiotics on day 3 had a higher temperature and higher respiratory frequency, and oxygen was administered more often as compared with the remaining patients. The observed mortality (due to pulmonary disease and all causes) was significantly higher in the group still receiving *i.v.* therapy on day 3 (8% versus 0%, $p = 0.02$, and 10.7% versus 0%, $p = 0.01$). These data suggest that patients still on *i.v.* antibiotics on day 3 were more severely ill. Therefore, we performed an additional analysis of patients in whom a switch was advisable on day 3 and compared the characteristics of the 27 patients who were not switched with the characteristics of the 41 patients who were adequately switched (online supplementary appendix IV). The aforementioned 27 patients

TABLE 2 Outcome measures[#]

	Total	Timing switch		p-value (95% CI)
		On or before day 3	After day 3	
Follow-up day 3				
Subjects	148 (100)	64 (43)	84 (57)	
Clinically stable [¶]				
According to protocol [†]	21 (21.2)	9 (28.1)	12 (17.9)	0.245
According to physician [§]	124 (84.4)	61 (95.3)	63 (75.9)	0.001
Clinical parameters [¶]				
Temperature °C	37.0±0.7	36.8±0.5	37.2±0.8	0.000 (-0.698- -0.211)
Oxygen saturation %	95±3	94.4±3.7	95.4±3.0	0.086 (-2.166-0.147)
Oxygen administered yes	66 (50.8)	18 (36.0)	48 (60.0)	0.008
Respiratory rate breaths·min ⁻¹	20±4	18±2	21±4	0.044 (-5.380- -0.081)
Blood pressure mmHg				
Diastolic	75±12	76±14	74±11	0.343 (-2.212-6.314)
Systolic	137±21	138±21	136±22	0.574 (-5.346-9.604)
Cardiac frequency beats·min ⁻¹	87±19	83±17	89±19	0.070 (-12.290-0.479)
Day 3 in weekend	38 (25.7)	12 (18.8)	26 (31.0)	0.092
Follow-up day 28				
Stop <i>i.v.</i> agents day	4.7±2.8	2.5±0.7	6.4±2.7	0.000 (-4.581- -3.219)
Length of hospital stay days	8.2±5.5	6.0±4.4	10.0±5.6	0.000 (-5.700- -2.362)
Readmission within 28 days				
All	13 (9.3)	8 (12.7)	5 (6.5)	0.208
Pulmonary pathology	5 (3.6)	2 (3.2)	3 (3.9)	1.000
Mortality within 28 days				
All	9 (6.1)	0 (0.0)	9 (10.7)	0.005
Pulmonary pathology	7 (4.7)	0 (0.0)	7 (8.3)	0.019
Physician characteristics				
Clinical experience yrs				
Resident as treating physician	2.2±1.9	2.4±2.2	2.0±1.6	0.320 (-0.309-0.938)
Supervisor as medical specialist	10.9±8.1	12.2±8.6	10.0±7.7	0.128 (-0.647-5.085)
Type of admission				
Lung ward	80 (54.1)	36 (56.3)	44 (52.4)	0.640
University hospital	44 (29.7)	25 (39.1)	19 (22.6)	0.030

Data are presented as n (%) or mean±SD, unless otherwise stated. Independent-sample t-tests were used for ratio variables and the Chi-squared test or Fisher's exact test for nominal variables. [#]: n=148; [¶]: missing data were not used in the calculation; [†]: patients discharged before day 3 were excluded; [§]: patients discharged before day 3 were marked as clinically stable according to the physician. Bold indicates statistical significance.

appeared to be comparable with the patients who were adequately switched; for example, clinical parameters on day 3 and mortality rate were not significantly different.

By means of the case-specific interviews, we evaluated the barriers to a conversion to oral antibiotics. We evaluated the barriers reported by physicians treating the 27 (18%) patients in whom a switch to oral antibiotics was possible but not performed (table 4). The most frequently mentioned barriers were "supervisor's opinion", "day 3 of admission was a Saturday or a Sunday" and "a switch to oral agents was possible, but forgotten"; each scenario was mentioned in five (19%) cases. Overall, the reported barriers can be grouped into three main categories: practical considerations mentioned in 13 (28%) out of 47, organisational factors mentioned in eight (17%) out of 47 and misconceptions mentioned in 28 (55%) out of 47.

Through the physician-specific interviews, additional theoretical barriers to an early-switch strategy were identified (table 4). Prominent discrepancies between the barriers mentioned in practice (patient-specific questionnaire) and in theory (physician-specific questionnaire) were forgetting to switch to oral agents (19% in practice *versus* 2% in theory), absence of an oral variant for the administered *i.v.* antibiotic (18% in theory *versus* 4% practice) and comorbidities delaying the switch (22% theory *versus* 4% practice).

DISCUSSION

In this study, a conversion from *i.v.* to oral antibiotics either before or on day 3 of treatment was possible in but not performed in 27 (40%) out of 68 of patients hospitalised with CAP. 94% of physicians were not aware of current guideline recommendations on the conversion to oral antibiotics.

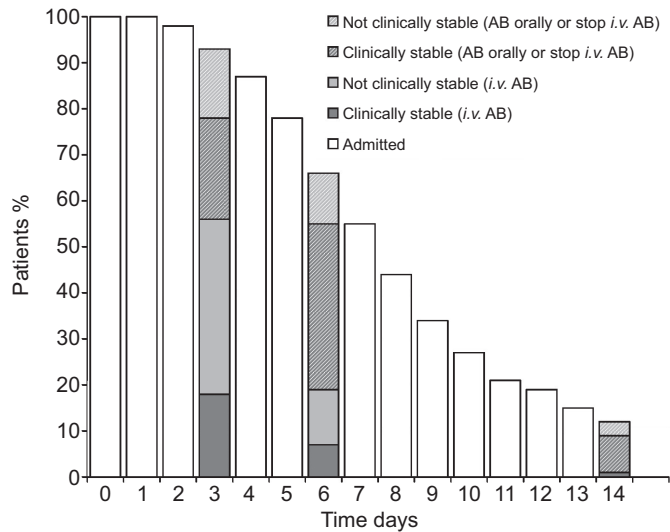


FIGURE 2. Patient stability and the route of antibiotic (AB) administration over time. *i.v.* intravenous.

Perceived barriers to an early-switch strategy included mainly misconceptions, practical considerations and organisational factors. It is therefore likely that the majority of the barriers identified in this study could be reduced by means of an educational intervention and structural organisational changes [7, 19]. This may reduce the duration of *i.v.* antibiotic treatment and, consequently, the length of hospital stay in CAP patients.

This study has several strengths. Through intensive screening, we included all patients consecutively hospitalised with CAP in both teaching and university hospital settings. The age and sex distribution, incidence of comorbidities, severity of presentation and duration of *i.v.* therapy are comparable with other cohorts of CAP patients, which enhances the generalisability of our results [7, 19, 20]. The percentage of patients in which an appropriate switch strategy was applied in this study was comparable with the percentage found in a similar study (60% versus 58%) [9]. Furthermore, physicians with working experience in >32 different hospitals were included and reasons for nonparticipation were unlikely to have influenced study results. Reasonably, the professional experience and expertise of the participating physicians represents those of all physicians working in the same setting in the Netherlands and possibly other countries with a similar healthcare system as well [10].

There are three main drawbacks of the chosen study design. First, measuring clinical parameters on several selected days of follow-up as opposed to each day of admission may have led to an underestimation of the number of patients that were not switched on a timely basis, as several of these patients might have reached clinical stability before day 3 and a switch to oral agents would have been appropriate at that time. Secondly, the type of observation might have influenced study results. By asking physicians about their motivation to administer the antibiotic either *i.v.* or orally in specific cases at set times during the admission, their attention is drawn to the chosen treatment and they are likely to re-evaluate their choices. This phenomenon might be reflected by the notably high percentage of patients that are switched on day 3, the first day of follow-up (fig. 2).

TABLE 3 Criteria applied by the 97 treating physicians when deciding to switch to oral antibiotics

Category	Qualitative answers [#]	Frequency n (%)
Clinical course		
Clinical judgement	Clinical improvement (clinical judgement or not specified)	36 (37.1)
Oral intake	Able to swallow, no nausea, vomiting or diarrhoea	20 (20.6)
Temperature	Fever subsiding	19 (19.6)
	No fever	36 (37.1)
	No fever for ≥ 1 day	7 (7.2)
	No fever for ≥ 2 days	6 (6.2)
	Other	1 (1.0)
Respiratory	Dyspnoea subsiding	16 (16.5)
	Less/no need for oxygen administration	24 (24.7)
Cardiovascular	Haemodynamically stable	21 (21.6)
	No signs of sepsis	4 (4.1)
Other		16 (16.5)
Patient characteristics		
Subjective	Patient feels better	16 (16.5)
Other		1 (1.0)
Laboratory parameters		
Inflammation parameters	Decrease in C-reactive protein	47 (48.4)
	Decrease in leukocytes	12 (12.4)
	Other	2 (2.0)
Culture	Pathogen known/culture results available	11 (11.3)
Practical considerations		
Timeline	<i>i.v.</i> antibiotics, at least until day 2	4 (4.1)
	<i>i.v.</i> antibiotics, at least until day 3	17 (17.5)
	<i>i.v.</i> antibiotics, at least until day 4	16 (16.5)
	Other	3 (3.1)
Choice of antibiotics	A good oral variant is available	6 (6.2)
Other		5 (5.1)

[#]: answers are displayed in detail if they were provided by ≥ 4 physicians.

Conversely, this effect could also be caused by the general awareness of the importance of an early-switch strategy, as shown through the physician-specific questionnaires, in which a majority of physicians state that the ideal timing of the switch is either before or on day 3. However, the potential influence of the latter drawback would lead to an underestimation of the number of patients who are not switched in a timely manner and does not lessen the room for improvement in this field. Lastly, our study results may have been different if carried out in more than three hospitals. However, the number of participating physicians is quite large as compared with other studies [7, 21], and physicians working in a teaching hospital as well as in a university hospital setting were included. Therefore, we feel that our results adequately reflect current practice in the Netherlands.

Optimisation of antibiotic therapy in CAP by implementation of guideline advice has been the focus of attention for several years. HAGAMAN *et al.* [8] evaluated the effect of the implementation of American Thoracic Society guidelines on the switch to

TABLE 4 Barriers to an early-switch strategy by case (case-specific questionnaire) and in theory (physician-specific questionnaire)

Factors	Qualitative answers	Frequency	
		Case specific [#]	Physician specific [†]
Physician factors			
	Opinion of supervisor	5 (19)	13 (13)
	Forgot	5 (19)	2 (2)
	Practice experience/intuitive	0	9 (9)
	Delay due to resident	0	5 (5)
	Other	NA	3 (3)
Patient characteristics			
Patient factors	Absorption orally not secured [‡]	NA	37 (38)
	Comorbidity	1 (4)	21 (22)
	Elderly patient (>75 yrs)	0	4 (4)
	Therapy adherence not secure	0	4 (4)
Clinical course	Patient very ill at admission	1 (4)	7 (7)
	Patient still ill	4 (15)	23 (24)
	Patient feels ill	1 (4)	0
	Fever	3 (11)	6 (6)
	Fever subsided <24 h ago	2 (7)	0
	Dyspnoea/oxygen needed	3 (11)	1 (1)
	Haemodynamically unstable	2 (7)	0
Additional diagnostics			
	Elevated CRP	3 (11)	3 (3)
	High leukocytes	1 (4)	0
	Abnormalities on chest radiography	1 (4)	1 (1)
	Confusion/delirium	0	4 (4)
	Empyema/pleural effusion/abscess	0	8 (8)
	Secondary infection	0	4 (4)
	Other	NA	11 (11)
Other hospital staff factors			
	Other	NA	4 (4)
Organisation factors			
	Weekend	5 (19)	11 (11)
	No supervision available	0	4 (4)
	Other	NA	9 (9)
Other			
Antibiotics	No oral variant for <i>i.v.</i> agent	1 (4)	17 (18)
	Allergy/toxicity oral variant	3 (11)	15 (16)
	Recent change in antibiotic regimen	1 (4)	1 (1)
	Short duration of <i>i.v.</i> therapy	1 (4)	0
	Other	NA	3 (3)
Microbiology	Culture results still unknown	3 (11)	12 (12)
	Causative pathogen is atypical	1 (4)	10 (10)
	Other	NA	6 (6)
Admission related/other	Patient stays admitted/needs <i>i.v.</i> medication for other reason	0	8 (8)
	Other	NA	2 (2)

Data are presented as n (%) of cases in which an answer was provided or n (%) of physicians that provided this answer, unless otherwise stated. Answers mentioned in the physician-specific questionnaires are only displayed in detail if they were provided by ≥ 4 physicians or if they were also provided in the patient-specific questionnaire; the remaining answers are provided elsewhere (supplementary appendix V). CRP: C-reactive protein; NA: not applicable. [#]: n=27; [†]: n=97; [‡]: e.g. not able to swallow, nausea or vomiting. Bold indicates percentages that differ by >10%.

oral antibiotics. Unfortunately, besides assessing the guideline knowledge of physicians, practical barriers to an early-switch strategy were not studied. The implementation effect could have been greater if aimed at specifically identified barriers [11–13]. SCHOUTEN *et al.* [7] and HALM *et al.* [22] assessed barriers to an early-switch strategy in CAP patients as perceived by physicians through semi-structured interviews and written surveys,

respectively. Perceived barriers to a timely switch strategy identified in these studies are also reflected by our study results, and include unfamiliarity with guideline recommendations, lack of outcome expectancy and misconceptions. In contrast to the latter studies, we are the first to compare the barriers mentioned in theory with the ones experienced in practice, completing the analysis of barriers to a timely switch strategy as

a whole. In addition, little is currently known about the quality and quantity of the influence of patients' knowledge, behaviour and assertiveness on the antibiotic treatment regimen in hospital care [10]. Our study shows a modest role for these patient-related factors; the switch to oral antibiotics was delayed because the patient felt ill in only one (4%) out of 27 cases.

The results of this study stress the need for a tailored intervention aimed at the identified barriers in order to stimulate an early switch in CAP patients. The effectiveness of such an intervention needs to be evaluated in a study setting. If effective, the implementation strategy can be applied in other centres and possibly also to other types of infection. Ideally, maximal results are obtained and all patients are switched to oral antibiotics in a timely manner after implementation. Based on our study results, theoretical implementation of an early-switch strategy in all CAP patients in the Netherlands would lead to reduction of length of hospital stay of 3.4 days [2] in an additional 18% of patients and, consequently, to a reduction of healthcare costs of nearly €9.6 million (based on 31,000 admissions annually and €505 per admission day) [23, 24].

In conclusion, the switch from *i.v.* to oral antibiotics is often unnecessarily delayed in patients hospitalised with CAP; this is mostly based on misconceptions and practical and organisational issues. A tailored intervention aimed at these factors is most likely to reduce the duration of treatment with *i.v.* antibiotics and, consequently, length of hospital stay.

SUPPORT STATEMENT

Funding was obtained from the Netherlands Organisation for Health Research and Development (ZonMw), grant no. 171103003.

STATEMENT OF INTEREST

None declared.

ACKNOWLEDGEMENTS

We thank all participating physicians and patients.

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