

# Pharmacists' Interventions in Prescribing Errors at Hospital Discharge

## An Observational Study in the Context of an Electronic Prescribing System in a UK Teaching Hospital

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### Abstract

**Background:** Pharmacists have an essential role in improving drug usage and preventing prescribing errors (PEs). PEs at the interface of care are common, sometimes leading to adverse drug events (ADEs). This was the first study to investigate, using a computerized search method, the number, types, severity, pharmacists' impact on PEs and predictors of PEs in the context of electronic prescribing (e-prescribing) at hospital discharge.

**Method:** This was a retrospective, observational, 4-week study, carried out in 2008 in the Medical and Elderly Care wards of a 904-bed teaching hospital in the northwest of England, operating an e-prescribing system at discharge. Details were obtained, using a systematic computerized search of the system, of medication orders either entered by doctors and discontinued by pharmacists or entered by pharmacists. Meetings were conducted within 5 days of data extraction with pharmacists doing their routine clinical work, who categorized the occurrence, type and severity of their interventions using a scale. An independent senior pharmacist retrospectively rated the severity and potential impact, and subjectively judged, based on experience, whether any error was a computer-related error (CRE). Discrepancies were resolved by multidisciplinary discussion. The Statistical Package for Social Sciences was used for descriptive data analysis. For the PE predictors, a multivariate logistic regression was performed using STATA<sup>®</sup> 7. Nine predictors were selected *a priori* from available prescribers', patients' and drug data.

**Results:** There were 7920 medication orders entered for 1038 patients (doctors entered 7712 orders; pharmacists entered 208 omitted orders). There were 675 (8.5% of 7920) interventions by pharmacists; 11 were not associated with PEs. Incidences of erroneous orders and patients with error were 8.0% (95% CI 7.4, 8.5 [n = 630/7920]) and 20.4% (95% CI 18.1, 22.9 [n = 212/1038]), respectively. The PE incidence was 8.4% (95% CI 7.8, 9.0 [n = 664/7920]). The top three medications associated with PEs were paracetamol (acetaminophen; 30 [4.8%]), salbutamol (albuterol; 28 [4.4%]) and omeprazole (25 [4.0%]).

Pharmacists intercepted 524 (83.2%) erroneous orders without referring to doctors, and 70% of erroneous orders within 24 hours. Omission (31.0%), drug selection (29.4%) and dosage regimen (18.1%) error types accounted for >75% of PEs. There were 18 (2.9%) serious, 481 (76.3%) significant and 131 (20.8%) minor erroneous orders. Most erroneous orders (469 [74.4%]) were rated as of significant severity and significant impact of pharmacists on PEs. CREs (n=279) accounted for 44.3% of erroneous orders. There was a significant difference in severity between CREs and non-CREs ( $\chi^2=38.88$ ;  $df=4$ ;  $p<0.001$ ), with CREs being less severe than non-CREs. Drugs with multiple oral formulations (odds ratio [OR] 2.1; 95% CI 1.25, 3.37;  $p=0.004$ ) and prescribing by junior doctors (OR 2.54; 95% CI 1.08, 5.99;  $p=0.03$ ) were significant predictors of PEs.

**Conclusions:** PEs commonly occur at hospital discharge, even with the use of an e-prescribing system. User and computer factors both appeared to contribute to the high error rate. The e-prescribing system facilitated the systematic extraction of data to investigate PEs in hospital practice. Pharmacists play an important role in rapidly documenting and preventing PEs before they reach and possibly harm patients. Pharmacists should understand CREs, so they complement, rather than duplicate, the e-prescribing system's strengths.

## Background

Pharmacists have an essential role in improving drug usage and preventing prescribing errors (PEs).<sup>[1,2]</sup> It has been reported that not all PEs are intercepted by pharmacists' interventions and, similarly, not all pharmacists' interventions are in response to PEs.<sup>[3]</sup> The relationship between pharmacists' interventions and PEs is poorly studied. PEs are very common in US<sup>[4,5]</sup> and UK<sup>[3,6]</sup> secondary care.<sup>[7]</sup> PEs at the interface of care, the point of hospital admission or discharge, are common and can lead to preventable adverse drug events (pADEs).<sup>[8]</sup> Errors attributable to prescribing make up the largest share of pADEs in hospitals.<sup>[9]</sup>

PEs occur at hospital discharge, during which patients' medication orders require verification and clarification from multiple sources of information, including the best possible medication history, medications prescribed during the inpatient stage (newly prescribed, adjusted or discontinued), unchanged patient's own medication, and the newly started medications at discharge.<sup>[10-13]</sup> PEs and ADEs after hospital discharge arise from insufficient medication reconciliation processes

at discharge and inadequate information to patients regarding their pharmacotherapy.<sup>[14-18]</sup> Some studies<sup>[5,19-21]</sup> have focused on pharmacists' interventions with PEs during or after hospital discharge of adults. However, none were specifically aimed at studying the rate, nature, potential severity and predictors of PEs in the context of electronic prescribing (e-prescribing) at hospital discharge. In addition, none of the studies investigated the relationship between pharmacists' interventions and PEs at the care interface.

Information technology, in the form of e-prescribing systems, has been proposed to reduce PEs in hospitals.<sup>[22]</sup> Although they have many benefits, e-prescribing systems are not fully protected against PEs; they have even facilitated the introduction of new types of PEs.<sup>[2]</sup> Most of the evidence-based findings of the strengths and weaknesses of e-prescribing systems have emerged from the US,<sup>[23-26]</sup> where clinical practice and e-prescribing systems are different from the UK.<sup>[27]</sup> No previous study has investigated pharmacists' interventions and PEs in the context of e-prescribing systems at hospital discharge using a computerized search method.

Studies have used a number of techniques and information sources to collect data for the identification of PEs, such as manual chart review, computerized search of electronic records, and trigger tool studies. Most PE studies have utilized paper-based records to detect and document error rate and types; moreover, data were manually collected, mostly by pharmacists, which may have been less efficient and more expensive<sup>[28]</sup> than a computerized search. Unsurprisingly, although these methods enjoy a number of advantages, all of them are inherently fraught with inaccuracies and systematic bias.<sup>[29]</sup>

The aim of this study was to characterize and investigate the number, types, severity and predictors of PEs in the context of an e-prescribing system at hospital discharge. Computer-related errors (CREs), high-risk medications, pharmacists' impact on PEs, and the circumstances of independent pharmacists' interventions were also investigated.

## Methods

### Setting and e-Prescribing System

The study hospital was a 904-bed teaching hospital, which cared for an average of 320 000 people a year in the northwest of England. There were more than 400 doctors and 35 pharmacists working in the hospital. A comprehensive pharmaceutical service was provided between 9:00am and 5:00pm on weekdays.

An electronic patient record with an e-prescribing system (iSoft Clinical Manager™; iSoft plc, Manchester, UK) was installed in the hospital in 2000. At the time of the study, the e-prescribing system was enabled at the time of discharge only – inpatient prescribing was still paper-based. When the patient was discharged, the prescriber used the inpatient chart to select the required medication from the e-prescribing system's drug catalogue and prepare the discharge prescription. The drug catalogue was accessed in a medicines 'look-up' list providing suggested default dosage regimens and formulations for most medications, which appeared upon typing the first letters of a medication's approved name. Free-text could be

entered by the prescriber if the medication or one of its characteristics did not exist in the look-up list. No further explicit clinical decision support was enabled in the system. Any pharmacist or doctor discontinuing a medication order had to select (from a pull-down menu) or type in (as free-text) a reason for discontinuation. Only one reason could be entered into the field for each discontinued medication order.

Each weekday the pharmacists routinely checked the discharge medication orders electronically prescribed by doctors and compared them with the inpatient paper-based drug chart to ensure that all orders were clear, legal, complete and clinically appropriate. Weekend prescriptions were checked by the dispensary pharmacist working on Saturdays and Sundays, supported by an on-call service. If any discrepancy, ambiguity or error existed in the discharge electronic order, the pharmacist might call the prescriber, discuss the matter and correct it (or let the prescriber correct it). Alternatively, if the pharmacist covered the ward on which the patient was located and had a complete picture about the patient's condition, the pharmacist might change the prescription independently without consulting the prescriber.

### Extraction of Data

Data were extracted from the electronic patient records using Microsoft Access® (Microsoft Access® 2003; Microsoft Corporation, Stateline, NV, USA) via SQL Server (Microsoft SQL Server, Microsoft Corporation, Redmond, WA, USA). Only medication orders prescribed by doctors and discontinued by pharmacists were included in the study. Patient characteristics (date of admission, date of discharge, age, sex, number of discharge medications and ward), medication order characteristics (drug name, strength, frequency, route and grade of prescriber) and any changes made in the subsequent discharge order (drug name, strength, frequency, route and person who made changes) stored in the system were manually collected on preset data entry forms by one of the authors (DAQ). The study was deemed service evaluation by a local Research Ethics Committee and formal review was not needed.

## Study Design

This was a recently retrospective, observational, cross-sectional, process-based<sup>[30]</sup> study. For practical reasons, the busiest wards (Medical and Care of the Elderly) were chosen. PEs were identified either during routine drug chart review (paper or electronic) or dispensing of a medication. Therefore, PEs could be intercepted and addressed with or without the knowledge of the prescriber, to avoid patient harm.

All discharge medication orders, discontinued by pharmacists and generated from the study wards by doctors, were included in the study, during 4 purposively selected weeks: the second week of May, first week of June, and first and fourth weeks of October 2008. A week was chosen because it included any potential errors occurring both in discharge and weekend leave medication orders. At the time of the study, junior doctors changed their rotation (within and outside the hospital) in February and August every year; hence, the authors selected 3 months in which doctors will presumably have had time to settle into the hospital's context and e-prescribing system.

To investigate the potential predictors of PEs, a retrospective case-control study was conducted in parallel with the observational study.

## Definitions

### Medication Order

A medication order was defined as any prescribed medication along with its characteristics (strength, unit of measurement, frequency, dosage form, route and administration information) intended to be administered to or by a patient after discharge from hospital.

### High-Risk Medication

A high-risk medication was defined as a medication that, if involved in an erroneous medication order, carried a greater risk of significant harm or death.<sup>[31]</sup> High-risk medications were compiled from the Institute for Safer Medication Practices' high-alert medication list,<sup>[32]</sup> as well as from articles that reported on high-risk medications (figure 1).<sup>[1,31,33]</sup>

1. Insulin
2. Anticoagulants
3. Opioids
4. Muscle relaxants
5. Chemotherapeutic agents
6. Intravenous potassium chloride/phosphate
7. Intravenous antibacterials
8. Antiepileptics
9. Cardiac glycosides
10. Respiratory stimulants
11. Antidepressants and mood stabilizers
12. Centrally acting antihypertensives
13. Antiarrhythmics
14. Adrenergic neurone blocker
15.  $\alpha$ -Blockers

**Fig. 1.** High-risk medications list – *British National Formulary* categories.<sup>[1,31-34]</sup>

### Prescribing Error (PE) Types and Severity

Dean and colleagues<sup>[35]</sup> definition of PE was adopted: “a clinically meaningful prescribing error occurs when, as a result of a prescribing decision or prescription [ordering] process, there is an unintentional significant (a) reduction in the probability of treatment being timely and effective or (b) increase in the risk of harm when compared with generally accepted practice.” According to this definition, prescribing without taking into account the patient's clinical status, failure to include essential information and errors in transcribing (i.e. from inpatient chart to electronic discharge prescription) are all considered PEs. However, failure to adhere to standards, such as prescribing guidelines or the drug's product licence, was not considered a PE where these reflected accepted practice. The *British National Formulary*<sup>[34]</sup> (BNF) was used as a reference for the use of medicines. All errors in regular, when-required (prn) and once-only medications were included. One of the authors (DAQ) classified all PE types. One erroneous medication order might have more than one PE type. PE types in the context of e-prescribing at discharge are shown in figure 2. Severity of error was classified using the scale of Overhage and Lukes,<sup>[36]</sup> with some modifications (table I). Control medication orders were those where no PEs were identified by the pharmacists.

### Types of Incidence

Six types of incidence were calculated. To calculate the incidence of PEs, all errors identified

during the study weeks were divided by all medication orders made (and those orders that should have been made but were omitted in error) during the study weeks. All errors were included when more than one error existed in the same order. The incidence of patients with error was calculated by dividing the number of patients prescribed an erroneous order by all discharged patients on the day of data collection. The incidence of erroneous orders was the number of medication orders with an error divided by the total number

of orders made plus those omitted in error. The incidence of omission errors (number of omission errors divided by orders made), incidence of commission errors (number of commission errors [i.e. PEs without omission errors] divided by orders made) and incidence of PEs relative to the opportunities for error (PE incidence divided by four) were also calculated. Each correctly indicated medication order was judged to have four opportunities for error (drug name, dose, frequency and formulation).

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| <p>1. Unintentional omission</p> <ul style="list-style-type: none"> <li>• Unintentional omission of drug from discharge prescription</li> <li>• A drug is not prescribed for a clinical condition for which one is indicated<sup>1</sup></li> <li>• Product, route or formulation not specified</li> <li>• Strength or dose not specified</li> <li>• Failure to specify the maximum dose for a drug prescribed to be given as required</li> <li>• Administration times not specified</li> <li>• Start date/end date not specified</li> <li>• Controlled drugs' prescription requirements</li> </ul>  |
| <p>2. Drug selection error</p> <ul style="list-style-type: none"> <li>• No indication for a drug prescribed when patient reaches discharge stage</li> <li>• Duplication of therapy (same drug or different drug with similar effect) when only one is clinically required</li> <li>• Prescription of drug to which patient has a history of significant allergy</li> <li>• Prescription of drug to which patient has clinical contraindication (drug-disease interaction)</li> <li>• Continuing a drug in the event of a clinically significant adverse drug event or sudden adverse drug reaction</li> <li>• Prescription of drug that is contraindicated due to drug interaction (drug-drug interaction)</li> <li>• Prescribing a new medication or continuing with a medication that is inappropriate for a discharged patient</li> <li>• Prescription of a different low therapeutic index brand name</li> <li>• Prescription of a drug that was not that intended (because of any reason, such as picking error, typing error, etc.)</li> </ul> |
| <p>3. Dosage regimen (strength, frequency, unit of measurement) selection error</p> <ul style="list-style-type: none"> <li>• Over-dosage for not taking into account drug-drug interaction</li> <li>• Over-dosage due to hepatic/renal impairment</li> <li>• Drug's strength correct, but frequency is incorrect</li> <li>• Total daily dose correct but divided into doses incorrectly, and it matters clinically<sup>1</sup></li> <li>• Over-dosage for a specific patient<sup>1</sup></li> <li>• Under-dosage</li> <li>• Dosage adjustment related to current disease severity/patient's condition</li> <li>• Prescription of a dose that was not that intended (because of any reason, such as picking error, typing error, etc.)</li> </ul>   |
| <p>4. Dosage form/route selection error</p> <ul style="list-style-type: none"> <li>• Wrong or inappropriate route</li> <li>• Wrong or inappropriate formulation</li> <li>• Prescription of a form/route that was not that intended (because of any reason, such as picking error, typing error, etc.)</li> </ul>   |
| <p>5. Patient selection error</p> <ul style="list-style-type: none"> <li>• Prescription of a drug for the wrong patient by unintentionally choosing the wrong patient's page on the system</li> </ul>  |
| <p>6. Therapeutic timing error</p> <ul style="list-style-type: none"> <li>• Premature discontinuation of drug on discharge (failure to renew treatment on discharge)</li> <li>• Continuation of drug for a longer duration than necessary where that is of clinical significance to the patient<sup>1</sup></li> <li>• Administration times incorrect</li> <li>• Start date/end date incorrect</li> </ul>  |

**Fig. 2.** Error types of electronic prescribing at discharge. <sup>1</sup> May be prescribing error, taking into account the patient's clinical condition.

**Table 1.** Instrument for characterizing pharmacists' clinical interventions (adapted from Overhage and Lukes<sup>[36]</sup>)

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**Severity of error in medication order: assess the inappropriateness of the order or its deviation from the standard of practice**

A. Potentially lethal

- High potential for life-threatening adverse effects/reactions
- Potentially lifesaving drug at a dosage too low for the disease being treated
- High dosage (>10 times normal) of drug with low therapeutic index

B. Serious

- Route of administration could lead to severe toxicity
- Low dosage of drug for serious disease in patient with acute distress
- High dosage (4–10 times normal) of drug with low therapeutic index
- Dosage resulted in serum drug concentration in potentially toxic range
- Drug could exacerbate the patient's condition (related to warnings or contraindications)
- Misspelling or mix-up in medication order could lead to dispensing of wrong drug
- Documented allergy to drug
- High dosage (>10 times normal) of drug without low therapeutic index

C. Significant

- High dosage (1.5–4 times normal) of drug with low therapeutic index
- Drug dosage too low for patient's condition
- High dosage (1.5–10 times normal) of drug without low therapeutic index
- Errant dual-drug therapy for single condition
- Inappropriate dosage interval
- Omission from medication order

D. Minor

- Incomplete information in medication order
- Unavailable or inappropriate dosage form
- Non-formulary drug
- Noncompliance with standard formulations and hospital policies

E. No error

- Information or clarification requested by physician or other healthcare professional from pharmacist
- Cost savings only

**Potential impact:<sup>a</sup> assess the potential impact of the pharmacist's recommendation on patient care**

1. *Extremely significant*: Recommendation qualified by extremely serious consequences or potential life-and-death situation
2. *Very significant*: Recommendation qualified by a potential or existing dysfunction in a major organ. Avoidance of serious adverse drug interaction or contraindication to use
3. *Significant*: Recommendation would bring patient care to a more acceptable, appropriate level (i.e. standard of practice), including quality-of-life issues with evidence from the patient or documentation elsewhere, as well as issues of cost and convenience. (Standard of practice is defined by institutional guidelines and protocols, and supported by acceptable references to the literature)
4. *Somewhat significant*: Patient's benefit from the recommendation could be neutral depending on professional interpretation (to distinguish this rank from rank 3, where a standard of practice would support the recommendation). More information or clarification must be obtained by the pharmacist from the physician, nurse or other appropriate healthcare professional before an order can be processed
5. *Not significant*: Information only. Recommendation not patient specific
6. *Adversely significant*: Recommendation inappropriate; its implementation may lead to adverse outcomes

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a Described in the original paper of Overhage and Lukes<sup>[36]</sup> as "value of service".

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### **Pharmacists' Clinical Intervention**

This is defined as "any reactive (in response to an erroneous medication order) activity undertaken to suggest changes in one medication order that might involve contacting medical staff."<sup>[3]</sup> Clinical interventions were classified as those associated or not associated with PEs.

### **Junior Doctors**

They were defined as doctors who were doing their first 2 years of postgraduate training in hospital, termed foundation years 1 and 2.

### **Identification and Classification of Pharmacists' Interventions**

All pharmacists' interventions were identified and validated by several members of the research team (the authors and the previously trained 29 pharmacists working in the hospital). Validation happened when the pharmacists utilized their 'insider' knowledge and experience, which is part of the pharmacist's routine clinical role, to decide initially upon the occurrence and severity of any PE recorded. Meetings among the research team members were held as soon as possible (within 1 week) after each intervention, to finalize judgments about PE occurrence and severity. One of the authors (DAQ) independently rated the severity and discussed any discrepancy with the pharmacist who made the intervention until reaching agreement. An open-ended question was posed to pharmacists as to whether they intervened independently or after having spoken to doctors. Doctors were unaware of the study in order to avoid changes in behaviour.

### **Severity of Errors and Impact of Pharmacists' Interventions**

When a medication order had more than one PE, the most severe error was considered in the severity, impact and CRE rating. Following the rating performed by the research team, a senior clinical pharmacist (LH), independent of the research team, retrospectively rated the severity and potential impact of PEs using a published scale.<sup>[36]</sup> The senior pharmacist was blinded as to whether interventions were pharmacist-led in order to

minimize any opportunity for rating bias. Any discrepancy in rating between the senior pharmacist and the research team was resolved by discussion and agreement between LH and one of the authors (DAQ).

### **Computer-Related Error**

Any error was subjectively classified as a CRE when it was more likely to have happened because of the presence of the e-prescribing system and would have been less likely to happen had the traditional handwritten prescribing been in place. The independent pharmacist and one of the authors (DAQ) independently judged whether any error was a CRE.

### **Data Analysis**

#### **Pharmacists' Interventions**

The statistical package SPSS<sup>®</sup> v.15 for Windows (SPSS Inc., Chicago, IL, USA) was used for this analysis. Nominal data were compared using the  $\chi^2$  test and continuous data by the unpaired *t*-test; 95% confidence intervals (CI) were calculated for parametric differences. An interrater reliability analysis using the weighted kappa ( $\kappa$ ) statistic was performed to determine consistency between raters of severity, pharmacist's impact and CREs. Values of kappa from 0.40 to 0.59 were considered moderate, 0.60 to 0.79 substantial, and 0.80 outstanding. For the open-ended question about independent pharmacists' interventions, the conversation was transcribed verbatim, coded and grouped into emerging themes.

#### **PE Predictors**

To investigate potential PE predictors, a retrospective, case-control, 4-week study was conducted. For a minimum detectable odds ratio (OR)  $\geq 1.2$ ,  $p=0.05$  and power = 0.8, a randomly selected total number of sample size required (assuming a population of 10% cases and 90% control) was 2624 medication orders, which were randomly drawn from medication orders, stratified by study week. The authors purposively selected 5% ( $n=30$  orders per week) of the controls (i.e. non-erroneous orders on which pharmacists did not intervene) to confirm that PEs did not exist.

*A priori* potential predictors of PE compiled from the literature and selected based on the available information from the electronic system were used as independent variables. These were grouped into:

1. Prescriber characteristics: doctor's grade, specialty and number of discharged patients prescribed for during the day of data collection (as a surrogate for workload).
2. Patient characteristics: age, sex, length of stay and number of co-prescribed drugs.
3. Drug characteristics: dosage form and BNF therapeutic category.

A multivariate logistic regression was performed based on a forced entry method using Stata<sup>®</sup> 7 (Stata Corp., College Station, TX, USA). Patients were clustered by the patient ID number. ORs, significance levels and 95% CIs were calculated.

## Results

### Overview of the Study Data

Doctors entered 7712 electronic discharge medication orders for 1038 patients (540 females, 498 males) on the study wards. There was no statistically significant difference between the numbers of erroneous or accurate orders among the 4 study weeks; hence, the data were combined. Pharmacists entered 208 omitted medication orders; therefore, in total there were 7920 medication orders entered by doctors and pharmacists during the study period. Demographics of patients associated with one or more PEs are shown in table II.

### Pharmacists' Interventions and PEs

There were 675 pharmacists' interventions recorded in the e-prescribing system; 11 were not associated with PEs. There were 630 erroneous medication orders, responsible for 664 PEs, for 212 patients; hence, the incidence of PEs was 8.4% (95% CI 7.8, 9.0) [table III].

### Erroneous Medication Orders and PEs per Patient

During the study period, pharmacists intercepted 630 erroneous medication orders in the e-prescribing system for 212 patients; hence, there were 3.0 erroneous orders per patient for all pa-

**Table II.** Demographics of patients prescribed medication orders with errors

| Characteristics  | Value           |
|--|-----------------|
| Total number of patients with prescribing errors                         | 212             |
| Mean age in years (SD)   | 65.8 (19)       |
| Median length of stay in days (SD, range)                                | 6.0 (8.2, 59.5) |
| Female (%)   | 114 (53.8)      |
| Mean patients' number of medications (SD, range)                         | 9.2 (4.0, 21.0) |
| Number of patients on erroneous high-risk medications (%)                | 62 (29.2)       |
| Co-morbidities (%)   | 188 (88.7)      |
| Speciality   |                 |
| medical wards (%)  | 150 (70.8)      |
| care of the elderly (%)  | 62 (29.2)       |
| Prescribed by junior doctors (%)   | 185 (87.3)      |
| Median number of patients prescribed for by a doctor per day (SD, range) | 3.0 (1.2, 5.0)  |

tients with error. There were 664 PEs detected for the same patients, with 3.1 PEs per patient for all patients with error.

### Specialities and Therapeutic Categories of Erroneous Medication Orders

Most erroneous medication orders were prescribed on the General Medicine (139 [22.0%]), Cardiology (113 [17.9%]) and Elderly Care (101 [16.0%]) wards. Eighty-one (12.9%) erroneous orders were for medications with only one oral formulation, 428 (67.9%) had multiple oral formulations and 69 (11.0%) orders were for inhalation formulations. Table IV classifies the erroneous orders according to the BNF therapeutic categories. A closer look at the BNF sub-therapeutic categories showed that antihypertensive classes had the largest share (94 [14.9%]), followed by analgesics and antirheumatics (87 [13.8%]), single and combined inhalation bronchodilators (62 [9.8%]), and antibacterials and antifungals (39 [6.2%]). The most frequently recorded individual medications associated with an error were paracetamol (acetaminophen; 30 [4.8%]), salbutamol (albuterol; 28 [4.4%]), omeprazole (25 [4.0%]), aspirin (21 [3.3%]), codeine (18 [2.9%]), Seretide<sup>®</sup> (salmeterol/fluticasone propionate; 15 [2.4%]), senna (14 [2.2%]) and prednisolone (13 [2.1%]). Of the total prescribed items (n = 7920) during the study period, paracetamol



had 9.0% (713), salbutamol 1.6% (127), omeprazole 2.6% (206), aspirin 3.6% (285), codeine 4.5% (356), Seretide® 1.4% (111), senna 2.5% (198) and prednisolone 1.5% (119).

### PE Types

The ratio between commission (n=456) and omission errors (n=208) was 2.2:1.0. Unintentional omission, selection of drug and dosage regimen error types accounted for more than three-quarters of PEs recorded (table V).

### PE Severity and Impact Ratings

Kappa statistics showed substantial agreement between raters (range 0.70–0.72). There were 18 (2.9%) serious, 481 (76.3%) significant and 131 (20.8%) minor erroneous orders. Most interventions on erroneous orders (506 [80.3%]) were rated as of significant impact. Around three-quarters (469 [74.4%]) of the corrected erroneous orders were rated as C3 (significant severity with significant impact on pharmacists) and 95 (15.1%) as D4 (minor severity with somewhat significant impact on pharmacists).

Severity of the errors appeared to differ according to the error types (table VI). There was a statistically significant difference between these error types versus their severity rating ( $\chi^2=388.97$ ;  $df=10$ ;  $p<0.001$ ). Only the unintentional omission, drug selection and dosage regimen error types were rated as serious. Examples of serious PEs detected are shown in table VII.

### Computer-Related Errors (CREs)

There were 279 (44.3%) erroneous orders rated as CREs. Erroneous orders were rated as serious in 7 (2.5%) and significant in 183 (65.6%) of CREs detected, in comparison with 11 serious (3.1%) and 288 (82.1%) significant erroneous orders not related to the computer system. There was a statistically significant difference in severity between CREs and those not related to the computer system ( $\chi^2=38.88$ ;  $df<4$ ;  $p<0.001$ ), with CREs being less severe than non-CREs. Table VIII lists some examples of whether PE types were computer related (CRE) or not. Clinical examples of PEs and their interpretation are shown in table IX.

**Table III.** Prescribing errors (PEs) and interventions incidence during the study period

| Code | Parameter   | Equation              | Value (95% CI)    |
|------|---|-----------------------|-------------------|
| A    | Total number of patients during study   |                       | 1038              |
| B    | Total number of electronic discharge medication orders prescribed by doctors  |                       | 7712              |
| C    | Total number of omission errors entered by pharmacists                        |                       | 208               |
| D    | Total number of patients with PEs   |                       | 212               |
|      | Total number of medication orders with:                                       |                       |                   |
| E    | 1 PE  |                       | 598               |
| F    | 2 PEs   |                       | 30                |
| G    | 3 PEs   |                       | 2                 |
| H    | Total number of erroneous medication orders intercepted by pharmacists        | $E + F + G$           | 630               |
| I    | Total number of PEs <sup>a</sup>  | $E + 2F + 3G$         | 664               |
| J    | Total number of interventions not associated with PE                          |                       | 11                |
| K    | Total number of clinical interventions  | $I + J$               | 675               |
| L    | Incidence of patients with error [%]  | $(D/A) * 100\%$       | 20.4 (18.1, 22.9) |
| M    | Incidence of PEs [%]  | $I/(B + C) * 100\%$   | 8.4 (7.8, 9.0)    |
| N    | Incidence of erroneous orders [%]   | $H/(B + C) * 100\%$   | 8.0 (7.4, 8.5)    |
| O    | Incidence of omission errors [%]  | $(C/B) * 100\%$       | 2.7 (2.4, 3.0)    |
| P    | Incidence of PEs without omission errors (incidence of commission errors) [%] | $[(I - C)/B] * 100\%$ | 5.9 (5.4, 6.4)    |
| Q    | Incidence of PEs vs opportunities for error <sup>b</sup> [%]                  | $I/4(B + C) * 100\%$  | 2.1 (2.0, 2.3)    |

a Seventeen medication orders were judged to be erroneous by the authors only, because the intervening pharmacists had left employment before meeting with them.

b Each correctly indicated medication order has four opportunities for error (drug name, dose, frequency and formulation).

**Table IV.** British National Formulary (BNF)<sup>[34]</sup> classification of the erroneous medication orders

| BNF category  | N (%)      |
|---|------------|
| Cardiovascular system                               | 160 (25.4) |
| Central nervous system                              | 128 (20.3) |
| Gastrointestinal system                             | 80 (12.7)  |
| Respiratory system                                  | 76 (12.1)  |
| Nutrition and blood                                 | 45 (7.1)   |
| Endocrine system                                    | 43 (6.8)   |
| Infections  | 39 (6.2)   |
| Musculoskeletal and joint diseases                  | 22 (3.5)   |
| Skin  | 12 (1.9)   |
| Obstetrics, gynaecology and urinary tract disorders | 10 (1.6)   |
| Eye   | 7 (1.1)    |
| Ear, nose and oropharynx                            | 6 (1.0)    |
| Malignant diseases and immunosuppression            | 2 (0.4)    |
| Total   | 630 (100)  |

#### Characteristics of Pharmacists' Interventions

Pharmacists' interventions were performed independently (i.e. without consulting doctors) in 524 (83.2%) erroneous medication orders. The elapsed time to make interventions ranged between 0.05 and 552 hours (median 6 hours, SD 116.3). Pharmacists intervened upon 70% (441) of erroneous orders within 24 hours.

#### Erroneous High-Risk Medication Orders

There were 81 erroneous high-risk medication orders for 62 patients; this accounted for 12.9% of the erroneous orders (n = 630). These high-risk orders were associated with 86 PEs; in other words, 13.0% of total PEs detected (n = 664). The most frequent (top ten) high-risk medications were codeine (18 [22.2%]), morphine (7 [8.6%]), doxazosin (6 [7.4%]), amitriptyline (5 [6.2%]), citalopram (5 [6.2%]), oxycodone (4 [4.9%]), warfarin (4 [4.9%]), tramadol (4 [4.9%]), insulin (3 [3.7%]) and digoxin (3 [3.7%]). The severity of five (6.2%) medication orders was rated as serious, 68 (84.0%) significant and eight (9.8%) minor. There was a statistically significant difference in CRE occurrence between high-risk medications and non-high-risk medications ( $\chi^2 = 11.073$ ;  $df = 2$ ;  $p = 0.004$ ), where the proportion of high-risk medications

was higher in the non-CREs (16.86%) than CREs (7.89%).

#### Predictors of PEs

None of the sample of control orders was associated with PEs and they were judged to be valid controls. Demographics of cases and controls are shown in table X.

Table XI gives information about the contribution of the predictor variables. Only two predictors (multiple oral formulations and junior doctor) contributed significantly to the predictive ability of the model.

#### Circumstances Where Pharmacists Independently Changed Erroneous Medication Orders

Changes to erroneous medication orders were done independently by pharmacists if they had access to patients' clinical data at the time of decision making, and had sufficient time to check and correct PEs.

*"It depends on where you're based, so I would usually do all my TTOs [discharge medication orders, i.e. to take out] on the ward in front of the patient. The ward round's just gone on, so I would normally be quite up-to-date with where [the medical team are] up to."* Ph1

Pharmacists would independently intervene if they had a good relationship with the prescriber. Pharmacists noted that they would not bother doctors with trivial mistakes, which at the same time could cause problems in a patient's treatment plan. Pharmacists also discussed some characteristics of the 'minor' erroneous medication orders

**Table V.** Prescribing error types classification

| Type                              | N (%)      |
|-----------------------------------|------------|
| Unintentional omission            | 206 (31.0) |
| Drug selection error              | 195 (29.4) |
| Dosage regimen selection error    | 120 (18.1) |
| Dosage form/route selection error | 105 (15.8) |
| Therapeutic timing error          | 7 (1.1)    |
| Patient selection error           | 31 (4.6)   |
| Total                             | 664 (100)  |

**Table VI.** Severity of erroneous orders cross-tabulated with their types<sup>a</sup>

| Type                              | Serious | Significant | Minor | Total |
|-----------------------------------|---------|-------------|-------|-------|
| Unintentional omission            | 6       | 187         | 9     | 202   |
| Drug selection error              | 6       | 167         | 20    | 193   |
| Dosage regimen selection error    | 6       | 109         | 4     | 119   |
| Dosage form/route selection error | 0       | 6           | 78    | 84    |
| Patient selection error           | 0       | 7           | 0     | 7     |
| Therapeutic timing error          | 0       | 5           | 20    | 25    |
| Total                             | 18      | 481         | 131   | 630   |

a Only the most severe error was rated, when more than one error was found in an erroneous order.

that they would change independently. For example, if a medication was not relevant to the patient's reason for admission or any illness they had in their admission, pharmacists would change the medication themselves. Interventions would also be done independently if changes in an 'erroneous' medication order did not adversely affect a patient's blood results.

*"I'll change it independently if it's minor and less problematic, for example, if it's a PRN, Beconase<sup>®</sup> nasal spray they only use it in the summer, or any drug for symptomatic relief, or*

*not relevant to their reason for admission."* Ph10

On the other hand, the majority of pharmacists claimed that wrong dosage, contraindicated therapy, starting new medication at discharge and serious health conditions were all reasons why they would contact the prescriber before they would do any clinical intervention.

*"Definitely it depends on how clear I am and what's going on. It depends on how serious and how potentially problematic the medication is. Something like aspirin, warfarin or an anti-hypertensive, I would not just say, oh it's obvious. I would check it more than something like an inhaler or something topical. So it depends on the reason for admission, the indication for that drug and the drug itself."* Ph4

## Discussion

This was the first study to investigate pharmacists' interventions and PEs through a systematic computerized search followed by structured meetings held within 1 week with the pharmacists who made the clinical interventions. This method does not replace other identification methods, but could

**Table VII.** Examples of serious prescribing errors

| Serious prescribing error                                      | Type   | Intervention   |
|--|--|--|
| Amiodarone 200 mg tab tid                                      | Unintentional omission of drug from TTO  | Added to patient's TTO medications                         |
| Bendroflumethiazide 2.5 mg tab morning                         | Duplication of therapy (in the presence of furosemide 20 mg tab morning)   | Removed from patient's TTO prescription                    |
| Codeine 30–60 mg tab q4–6h                                     | Duplication of therapy (in presence of tramadol 100 mg cap tid)  | Removed from patient's TTO prescription                    |
| Codeine 30 mg tab q4–6h  | Continuing a drug in the event of a clinically significant ADE or sudden ADR (nausea, vomiting and constipation) | Replaced with tramadol 50 mg cap q6h                       |
| Dexamethasone 4 mg tab bid                                     | Overdosage   | Dexamethasone 2 mg tab tid, 2 mg bid, 2 mg qd over 10 days |
| Isosorbide mononitrate 10 mg tab morning                       | Underdosage  | Isosorbide mononitrate M/R 120 mg tab morning              |
| Nifedipine (Adalat <sup>®</sup> Retard) 5 mg M/R tab bid       | Duplication of therapy (in presence of nifedipine 5 mg cap bid)  | Removed from patient's TTO prescription                    |
| Nitrofurantoin M/R (Macrobid <sup>®</sup> ) 100 mg cap M/R bid | Prescription of drug to which patient has clinical contraindication (drug-disease interaction)                   | Replaced with trimethoprim 200 mg bid                      |
| Risedronate 6 mg tab morning                                   | No indication for a drug prescribed  | Replaced with risperidone 6 mg tab night                   |

**ADE** = adverse drug event; **ADR** = adverse drug reaction; **bid** = twice daily; **cap** = capsule; **M/R** = modified release; **qd** = once daily; **qxh** = every x hours; **tab** = tablet; **tid** = three times daily; **TTO** = to take out; discharge medication order.

**Table VIII.** Examples of prescribing errors and their relation to computer-related error (CRE)

| Description of error   | Is it a CRE?  |
|--|---------------|
| Drug's strength is correct, but the frequency is incorrect   | Yes           |
| Duplication of therapy because previous medication order is still active in the system   | Yes           |
| Duplication of therapy due to slip or lapse  | No            |
| No indication for a drug prescribed when patients reached discharge  | No/uncertain  |
| Overdosage or underdosage  | Yes/uncertain |
| Prescription of a medication that was not that intended from the system's 'look-up' list   | Yes           |
| Selection of the wrong medication due to picking error from an alphabetical list of medications in the e-prescribing system's drug catalogue               | Yes           |
| Start/end date of therapy is incorrect due to mistake in calculating duration of therapy   | No            |
| Start/end date of therapy is incorrect due to system's default regimen   | Yes           |
| Unintentional omission of a medication because the doctor did not select the 'discharge' option while completing the electronic TTO                        | Yes           |
| Unintentional omission of a medication because the patient has more than one drug chart, or doctor has not checked patient's self-administered medications | No            |
| Wrong dosage regimen for certain medications that require complex prescribing scenarios (e.g. amiodarone, prednisolone)                                    | Yes           |
| Wrong or inappropriate drug formulation  | Yes           |

**TTO** = to take out; discharge medication order.

provide a technique that is relatively consistent, objective, efficient and inexpensive.<sup>[29]</sup> This study employed a 'standard of reference', i.e. the reports given by pharmacists making clinical interventions on erroneous medication orders. One of the authors conducted the entire data extraction and meetings, which provided consistency of data collection and investigation throughout the study. This approach was deemed appropriate, taking into account its validity, efficiency and feasibility relative to the traditional manual chart review.<sup>[37]</sup>

### Study Limitations

The study has three limitations. First, generalizability of the results from a single teaching hospital in the northwest of England, from Medical

and Elderly Care wards, with one version of e-prescribing system at one point in time, is limited.

Second, the incidence of clinical interventions and PEs might have been underestimated since no independent check (by, for example, an external researcher) of all doctors' medication orders was performed; a previous study<sup>[38]</sup> showed that more than one-third of the errors were revealed by the principal investigator following an additional check.

Finally, the substantial agreement between raters could have been different had a doctor taken part in the rating. A study<sup>[1]</sup> reported that an internal medicine specialist rated the severity and impact of pharmaceutical care interventions lower than the hospital pharmacist. In addition, albeit minimal, the hindsight bias might still have had its effect to construct seemingly reasonable justifications for the raters' conclusions. However, the problem of multiple pharmacists rating PE severity differently had minimal effect on the overall rating since one person prospectively met with the pharmacists and resolved any discrepancy.

### Incidence, Types and Severity of PEs

The PE incidence in this study was at least twice as high as the PE incidence (or prevalence) in other studies.<sup>[38-41]</sup> The 'mandatory' documentation of the reason for any medication discontinuation in the e-prescribing system as well as the computerized data collection method might have facilitated the emergence of this high error rate. Furthermore, the discharge stage may have had more opportunities for error than the inpatient stage.<sup>[8]</sup> As with all other methods, the PE incidence calculated here could not reflect the 'true' overall PE rate. The incidences reflected only the recorded clinical interventions made by pharmacists. Errors detected and corrected by other healthcare professionals (e.g. doctors and nurses) were not investigated in this study, which constitute part of the overall PE rate. Moreover, errors that were not recorded in the e-prescribing system (e.g. verbal consultation by pharmacists) were not included in the study.

Whichever type of incidence is used to explain the frequency of PEs, the equation of calculation

should be explicitly stated, along with consistent definitions and methods, to help researchers confidently perform comparisons and draw meaningful conclusions. The authors argue that choosing 'opportunities for error' as the denominator when calculating PE incidence would give a more meaningful ratio than when choosing the total number of medication orders (made plus omitted) as a denominator. In addition, having separate incidences for commission and omission errors, with a common denominator being the number of orders made in the study period, would logically relate each type of error to the total number of orders prescribed by doctors only, rather than including omitted orders prescribed by pharmacists as well.

Anton et al.<sup>[42]</sup> argued, using the number of e-prescribing system's warning messages generated per prescription, that doctors' prescribing improves with time as they become more experienced with

the e-prescribing system. The lack of difference among PE rates between weeks in this study, however, contradicts their finding. Future research is required to explore the reasons why different grades of doctors make PEs in the context of an e-prescribing system both quantitatively and qualitatively.

Unintentional omission of medication orders was the largest type of PEs detected, followed by drug selection error and then dosage regimen errors. This was not consistent with UK studies conducted during the inpatient stage where dosage regimen error ranked the first type.<sup>[38,41]</sup> Collecting data about electronic, rather than paper-based, prescriptions and about discharge prescriptions only, might have caused this difference in PE types.<sup>[38,41]</sup> A recent review<sup>[26]</sup> of the evidence of the impact of e-prescribing systems on medication errors has found that the use of e-prescribing systems was associated with a 66%

**Table IX.** Examples of prescribing errors and their interpretations

| Erroneous medication order  | Prescribing error type                                    | Interpretation by research team  |
|---|---|--|
| Aminophylline (phyllocontin continus <sup>®</sup> ) 225 mg tab bid  | Unintentional omission                                    | Patient had more than one drug chart. Doctor did not check all charts  |
| Amiodarone 200 mg tab tid   | Unintentional omission                                    | Doctor did not select the 'discharge' option, hence the order remained in its default form (inpatient) and no TTO was obtained for patient |
| Clarithromycin 500 mg tab bid, stop date on 13/05 instead of 16/05  | Therapeutic timing error – start/end date incorrect       | Doctor did not enter the correct stop date in the system   |
| Glyceril trinitrate spray one to two puffs prn  | Unintentional omission                                    | Doctor did not enter the self-administered medication the patient was on before admission  |
| Isosorbide mononitrate 60 mg M/R tab bid in the presence of previous similar order  | Drug selection error – duplication                        | Doctor did not realise that a previous similar order had been entered days ago and was still active in the system                          |
| Lisinopril 10 mg tab morning, instead of 20 mg tab morning  | Dosage regimen error – underdosage                        | Doctor picked up the first PDO that comes on the screen without checking the full medication order characteristics                         |
| Metformin 1000 mg tab daily (12 noon), instead of daily (6:00pm)  | Therapeutic timing error – administration times incorrect | Doctor picked up the first PDO that comes on the screen without checking the full medication order characteristics                         |
| Nicotine inhalator (Nicorette <sup>®</sup> ) one inhalation prn   | Unintentional omission                                    | Doctor did not check the prn section in the drug chart   |
| Nitrofurantoin M/R (Macrobid <sup>®</sup> ) 100 mg cap M/R bid  | Drug selection error – contraindication                   | Doctor did not check the patient's renal function at discharge. Patient was renally impaired   |
| Seretide <sup>®</sup> (salmeterol/fluticasone propionate) CFC-free Evohaler two puffs bid, instead of Accuhaler one blister bid | Dosage form error   | Doctor picked up the first PDO that comes on the screen without checking the full medication order characteristics                         |
| Risedronate 6 mg tab morning  | Drug selection error – wrong drug                         | Doctor wrongly picked up the PDO that comes on the screen in alphabetical order (instead of risperidone 6 mg tab morning)                  |

**bid** = twice daily; **cap** = capsule; **CFC** = chlorofluorocarbon; **M/R** = modified release; **PDO** = pre-defined order; **prn** = when required; **tab** = tablet; **tid** = three times daily; **TTO** = discharge medication order (to take out).

**Table X.** Demographics of cases and controls (n=2630 medication orders)

| Predictor  | Control (n=2421)  | Case (n=209)      |
|--|-------------------|-------------------|
| Median length of stay in days (SD, range)                      | 7.0 (4.3, 29.0)   | 7.0 (4.6, 29.0)   |
| Mean age in years (SD, range)                                  | 64.8 (15.1, 73.0) | 64.2 (16.8, 71.0) |
| Female (%)   | 1074 (44.4)       | 134 (64.1)        |
| Mean patient's number of medications (SD, range)               | 9.4 (2.7, 14.0)   | 10.3 (3.8, 14.0)  |
| Form   |                   |                   |
| single form oral (%)   | 455 (18.8)        | 36 (17.2)         |
| multiple form oral (%)   | 1158 (47.8)       | 128 (61.2)        |
| other (%)  | 808 (33.4)        | 45 (21.5)         |
| British National Formulary Category <sup>[34]</sup>            |                   |                   |
| gastrointestinal system (%)                                    | 216 (8.9)         | 15 (7.2)          |
| cardiovascular system (%)                                      | 707 (29.2)        | 49 (23.4)         |
| respiratory system (%)   | 302 (12.5)        | 27 (12.9)         |
| others (%)   | 1196 (49.4)       | 118 (56.6)        |
| Junior doctor (%)  | 1425 (59.0)       | 156 (74.6)        |
| Specialty  |                   |                   |
| medical (%)  | 1813 (74.9)       | 169 (80.9)        |
| elderly care (%)   | 608 (25.1)        | 40 (19.1)         |
| Median number of patients prescribed for by doctor (SD, range) | 3.0 (1.1, 5.0)    | 3.0 (1.7, 5.0)    |

reduction in total PEs in adults. It showed that the rate of prescribing a wrong drug did not decrease after the introduction of an e-prescribing system; moreover, four studies of seven did not show significant changes in wrong dose prescribing. In conclusion, this shows that a considerable risk for PEs and ADEs still exists, which shows the importance of involving clinical pharmacy services in a multidisciplinary intervention strategy.

The types and frequency of intercepted medication classes in this study were consistent with that shown in a meta-analysis on pADEs.<sup>[43]</sup> Interestingly, studies have emphasized that the use of the identified medication classes has been a frequent cause of hospital admission,<sup>[44,45]</sup> or ADEs following discharge.<sup>[46]</sup> In this study, certain commonly prescribed medication classes were more likely to be associated with PEs, in particular analgesics, bronchodilators and aspirin. Therefore, these PE-associated medications (especially the high-risk medications) are good targets to improve an e-prescribing system's decision support or doctor education about good prescribing practice.

This study used a practical, reliable and comprehensive method<sup>[36]</sup> for assessing PE severity

and pharmacists' intervention impact. The distribution of severity rating percentages in this study was closely consistent with the ratings in other studies.<sup>[3]</sup>

The availability of a medication in multiple oral formulations and prescribing by junior doctors were two expected predictors of PEs, and closely concur with another study.<sup>[47]</sup> That junior doctors' prescribing medication orders was shown to be an independent risk factor for PEs, compared with senior doctors, is argued to imply either that medication prescribing improves with experience or that junior doctors exert less intellectual effort on (or spare less time for) prescribing at discharge. Furthermore, both predictors imply that doctors, especially the juniors, should be trained in how to write correct and complete medication orders.<sup>[48,49]</sup> Lack of prescribing training for junior doctors should be looked at as a crucial opportunity for development. Medical students both in the UK<sup>[50]</sup> and abroad<sup>[51]</sup> have reported that they feel that therapeutics teaching is not given enough time. Future research should be directed towards the incidence and causative factors of PEs committed by junior doctors; this is a recommendation of a recent systematic review that

failed to reach meaningful conclusions given the wide range of designs, methods and error rates.<sup>[52]</sup>

### Characteristics and Impact of Pharmacists' Interventions

It is obvious that the number of *recorded* pharmacists' clinical interventions not associated with PEs was astonishingly few (only 11), in comparison with a study<sup>[1]</sup> reporting a rate of 15.2%. Underreporting of this type of pharmacists' intervention is attributed to the inflexibility of the e-prescribing system designed to help pharmacists capture almost all PE types but not all pharmaceutical care activities. It is also important to note that many of the typical pharmacist interventions are done during hospital admission and not at the point of discharge. One study<sup>[3]</sup> reported that only 60% of pharmacists' interventions related to PEs. A French study<sup>[2]</sup> reported a pharmacists' interventions rate similar to the rate in this study.

**Table XI.** Multivariate logistic regression for prescribing error predictors (n=2630)<sup>a</sup>

| Predictor   | OR (95% CI) <sup>b</sup> | p-Value |
|---|--------------------------|---------|
| Median length of stay (days)                        | 1.02 (0.94, 1.11)        | NS      |
| Age (y)   | 0.99 (0.97, 1.02)        | NS      |
| Female (male as Ref)                                | 0.47 (0.21, 1.04)        | NS      |
| Patient's number of medications                     | 1.07 (0.89, 1.28)        | NS      |
| Form  |                          |         |
| single form oral (Ref)                              |                          |         |
| multiple form oral                                  | 2.1 (1.25, 3.37)         | 0.004   |
| other   | 1.46 (0.87, 2.45)        | NS      |
| British National Formulary Category <sup>[34]</sup> |                          |         |
| gastrointestinal system (Ref)                       |                          |         |
| cardiovascular system                               | 0.53 (0.27, 1.04)        | NS      |
| respiratory system                                  | 0.80 (0.49, 1.29)        | NS      |
| other   | 1.08 (0.54, 2.18)        | NS      |
| Junior doctor                                       | 2.54 (1.08, 5.99)        | 0.03    |
| Specialty   |                          |         |
| medical (Ref)                                       |                          |         |
| elderly care  | 0.76 (0.32, 1.83)        | NS      |
| Patients prescribed for by doctor                   | 1.29 (0.84, 2.01)        | NS      |

a Wald  $\chi^2 = 45.61$  ( $p < 0.0001$ ); pseudo  $R^2 = 0.07$ ;  $-2 \log$  likelihood =  $-673.2$ .

b Significant ORs were adjusted for age and sex.

NS = non-significant; OR = odds ratio; Ref = reference variable.

Independent interventions included all interventions independently done by the pharmacists regardless of whether they informed the doctors afterwards or not. The high proportion of these independent interventions and the short time elapsed to do these could fairly reflect how 'competent' the pharmacists were in checking and correcting discharge medication orders, as well as how much they were up-to-date with their patients' clinical status at discharge. Reasons for the wide time lapse for pharmacists to intervene upon a few erroneous orders included (i) when doctors had already entered discharge medication orders for patients whose discharges were delayed because of unpredicted changes in their health status at discharge; (ii) lack of efficient communication between pharmacists and doctors, where pharmacists were sometimes not up-to-date with patients' clinical status and medication changes happened during consultant ward rounds; and (iii) heavy workload, pressure to discharge patients and shortage of pharmacy staff.

The findings of this study contribute to the body of literature that pharmacists provide a crucial 'safety net' to improve the quality of hospital prescribing.<sup>[1,29,39,41,53]</sup> The independent and efficient pharmacists' interventions reported in this study concur with the view that pharmacists are in the best position to transcribe, prescribe and change medication orders at discharge.<sup>[54-56]</sup> Given similarities in error types between inpatient and discharge stages, the latter argument can be extended to pharmacists intervening at the inpatient stage.<sup>[57,58]</sup> Strategically, the findings support the importance of developing a systematic medicines reconciliation process at hospital discharge, which is consistent with findings from other studies.<sup>[10]</sup>

### Implications for Development of e-Prescribing Systems and Clinical Pharmacy Practice

The percentage of CREs in this study was higher than that reported in a UK study,<sup>[59]</sup> but closely similar to one French study.<sup>[2]</sup> Most of the CRE types identified in this study have been reported in the US and UK literature.<sup>[3,23,25]</sup> The information gathered about these errors should be used to improve the structure and facilities of

the e-prescribing system. Suggested system enhancements could include the enabling of a decision-support facility to prevent 'explicit' (e.g. duplication of order) and more complicated (overdosage for a patient with renal impairment) errors, and the ability for the prescriber to view and prescribe medication orders throughout the patient's journey from admission to discharge. Other examples of a system's improvement of features could include the use of Tallman lettering,<sup>1</sup> and showing the part of the dosage regimen and formulation at the beginning of any pre-defined order selected from the 'look-up' list (rather than keeping it after the drug name). Ergonomic improvement of the e-prescribing system cannot completely replace the pharmacist's role in optimizing patient care, but it may allow pharmacists to focus on the most relevant clinical interventions.<sup>[2]</sup> A recent systematic review<sup>[60]</sup> has concluded that the evidence base reporting the effectiveness of e-prescribing to reduce PEs is not compelling.

Koppel commented, "*Evaluation of CPOE [Computerized Prescriber Order Entry] systems, and of all healthcare information technology, is mostly terra incognita.*"<sup>[53]</sup> Any decision to develop the existing e-prescribing system has to be evidence based; this would help estimate the harm avoided by such systems, which could be disproportional to their reduction in errors.<sup>[38]</sup> In addition, changing the clinical practice in response to e-prescribing system implementation should be simultaneously investigated as well, so pharmacists can work to complement, rather than duplicate, an e-prescribing system's benefits.<sup>[3]</sup> Foot and Taylor<sup>[61]</sup> argue that the area that has to change most working practices is pharmacy, which should ensure the safe and effective use of the e-prescribing system within its remit. To date, published data on how e-prescribing has affected pharmacy services in the UK are limited.<sup>[62]</sup>

A multicentre, interrupted time-series study, adopting multi-method approaches, involving a range of qualitative and quantitative techniques, employing experienced study pharmacists and using cost-benefit analysis, needs to be conducted to

investigate the incidence of PEs and ADEs in the context of e-prescribing systems.<sup>[63]</sup>

## Conclusions

It is undeniable that PEs are common and will continue to exist, particularly in a field where complexity is the rule rather than the exception. In this study, four PEs occurred every hour in discharge prescribing on a normal workday, and one-third of the patients with PEs received high-risk medications. The pressing need for updated patient and therapeutic knowledge, the overlapping tasks of medical teams and flaws in the human-machine interface have all contributed to the emergence of PEs in hospitals.<sup>[24,25]</sup> Although challenging, pharmacists play a prominent role in efficiently documenting and preventing PEs before they reach and possibly harm the patient. There is no perfect solution for PEs. Implementing clinical pharmacy services, utilizing an e-prescribing system supported with clinical decision facility, and enhancing the awareness of the PE problem could all minimize the impact of this problem on patient safety. "*Substitution of technology for people is misunderstanding of both.*"<sup>[25]</sup>

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**1** Tallman lettering is the practice of writing part of a drug's name in upper case letters to help distinguish sound-alike, look-alike drugs (e.g., predniSONE/prednisoLONE) from one another in order to avoid prescribing errors.



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