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1 TITLE PAGE

2 Puking less per pound, for acute wheezers: quality improvement in a Pediatric

3 Emergency Department

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44ABSTRACT

45**Background & local problem:** Acute wheezing attacks are a leading cause of Pediatric
46Emergency Department (PED) attendances and inpatient admissions and are a
47considerable burden on the healthcare providers. Almost one-third of children vomit
48prednisolone in the PED, requiring anti-emetics and repeat dosing.

49**Aim:** This quality improvement (QI) intervention aimed to improve oral corticosteroid
50(OCS) tolerability, reduce emergency department length of stay (LOS), and reduce OCS
51drug costs for acute wheeze attendances in a PED, while not adversely affecting
52admissions, re-attendance, or mortality rates.

53**QI Interventions:** Included (1) a departmental protocol and (2) modification of the OCS
54type and dosage from prednisolone (3-day course of 1 mg/kg) to dexamethasone (600
55mcg/kg, then single dose 300 mcg/kg).

56**Methods:** The study team reviewed the evidence and implemented the interventions.
57To assess the scale of improvement, we retrospectively collected data on attendance
58records for patients aged 2-14 years with acute wheeze requiring OCS. We collected
59data on 100 children who attended the PED between October and December for each
60year (2016, 2017, and 2018). We then assessed OCS tolerability, LOS, OCS drug costs,
61and, admission, re-attendance, and mortality rates.

62**Results:** Over a 48-month period, we increased OCS tolerability by 67.2% and achieved
63an 85.8% reduction in OCS drug costs (saving £41,553.14). There was no change in the
64LOS, admission, re-attendance, and mortality rates.

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65**Conclusions:** Improved tolerability and substantial cost savings can be achieved by
66implementing a structured acute pediatric wheeze protocol and modifying the OCS to
67single-dose dexamethasone (300 mcg/kg).

68**Registration:** Not registered

69BACKGROUND

70Acute wheezing attacks are a leading cause of Pediatric Emergency Department (PED)
71attendances ¹. Children presenting in respiratory distress, due to asthma attacks or
72episodic viral wheeze (EVW), often require recurrent bronchodilator therapy, oral
73corticosteroids (OCSs), continuous assessment, and occasionally, inpatient admission ²⁻⁴.
74Treatments can be unpleasant, and while early administration of prednisolone, an OCS,
75is recommended, 28% of children vomit, causing further distress and necessitating
76repeated dosing or an intravenous or intramuscular alternative ⁵.
77Achieving patient flow is a persistent challenge for PEDs, especially in autumn and
78winter when viruses are prevalent ^{6,7}. Vomiting of OCS medications, in addition to being
79unpleasant, may delay the therapeutic effects, in turn, prolonging the length of stay
80(LOS) in PEDs. Quality improvement (QI) interventions to tackle these challenges are
81crucial for improving patient outcomes, experiences, and cost-effectiveness, and are
82commonly underpinned by research ^{8,9}.
83Recently, progress has been made on manufacturing flavoured prednisolone
84preparations; although more palatable, they would come at extra cost to the healthcare
85providers ⁵. Dexamethasone, a corticosteroid routinely used in childhood croup, has
86been trialled in acute wheeze as an alternative to prednisolone. Dexamethasone is a
87glucocorticoid with anti-inflammatory potency seven times greater than prednisolone. It
88is enterally well absorbed, yielding peak serum concentrations within 1-2 hours, yet, in
89comparison to prednisolone, dexamethasone has a relatively long half-life of 36-52
90hours ¹⁰⁻¹². A systematic review and meta-analysis found a single dose (or 2-dose regime)
91of dexamethasone to be a viable alternative to a 5-day course of prednisolone, noting
92patients were less likely to vomit ¹³. Cronin *et al.* ¹⁴ compared a single-dose oral

93dexamethasone (300 mcg/kg) to a three-day course of oral prednisolone (1 mg/kg);
94reporting no difference in admission rates, unscheduled attendances, or patients'
95pediatric respiratory assessment measure (PRAM) scores. Dexamethasone may also be
96more cost-effective, although actual savings are unclear ^{15,16}. And while structured
97asthma management protocols have shown some reduction in medication delivery
98times and hospital LOS ¹⁶⁻¹⁸, less is known about the scale of improvement in designated
99PEDs.

100This quality improvement project (QIP) aimed to improve OCS tolerability, reduce LOS in
101the PED, and reduce OCS drug costs for patients with acute pediatric wheeze in a UK
102PED, while maintaining admission, re-attendance, and mortality rates. To achieve this,
103we used a quality improvement framework endorsed by The Institute of Healthcare
104Improvement (IHI) ¹⁹, and based on current evidence, implemented a structured
105protocol and changed the OCS from prednisolone to dexamethasone.

106METHODS

107We undertook this QIP in the PED within the Royal Belfast Hospital for Sick Children
108(RBHSC) over a three-year period (2016-18). We report the study information in-line
109with the Standards for Quality Improvement Reporting Excellence (SQUIRE) 2.0 and
110include a checklist (in supplementary file 1) ²⁰.

111Contextual information

112The RBHSC is a teaching hospital that provides local secondary care to Belfast and
113tertiary care throughout Northern Ireland (UK). The RBHSC PED provides acute care for
114over 45,000 children annually, from birth up to 14 years.

115The QIP team & procedures

116The QIP team included two PED consultants (BB & SM), three PED trainees (GD, BK, &
117RM), two pediatric respiratory consultants (HS & MDS), two medical students (JR & MT),
118a pediatric clinical pharmacist (JM), and a statistician (VO).

119The aims

120This QIP aimed to:

- 121 1. Increase the number of children who tolerated OCSs by 70% within 12-months
122 (and maintain this increase for a further 12-months). OCS tolerability was
123 defined as successful OCS ingestion without 'spitting out' during administration
124 or vomiting post-administration while in the department.
- 125 2. Reduce LOS by 20% within 12-months (and maintain for a further 12-months).
126 LOS was defined as time from arrival at the PED, to discharge home (and
127 excluded those admitted).

- 128 3. No change in hospital admissions, re-attendances, or mortality rates at 12-
129 months (and at 24-months). Hospital admissions (to a medical ward or short-stay
130 unit) and re-attendances (unscheduled, to primary or secondary care)
131 throughout Northern Ireland within seven days of initial attendance were
132 included.
- 133 4. Reduce annual OCS drug costs by 80% within 12-months (and maintain for a
134 further 12-months).

135 Outcome measures (aims 1, 2, and 4) examined the desired impact on patient
136 outcomes, while balancing measures (aim 3) assessed for unintended consequences.
137 Specific targets of 70%, 20%, and 80%, for aims 1,2, and 4, respectively, were based on
138 published evidence and national drug costings ^{13,17,21}.

139 **QI interventions: structured protocol & OSC drug type/dose**

140 In 2016, prior to QI implementation and consistent with national guidance ^{3,4,21}, 1-2
141 mg/kg of oral prednisolone (maximum 40 mg), in the form of 5 mg soluble tablets, were
142 administered to patients in the PED with the first bronchodilator, with further doses
143 given at home for two consecutive days. In September 2017, we implemented the
144 'Acute asthma and viral-induced wheeze management protocol' (supplementary file 2),
145 recommending a single 600 mcg/kg dose of oral dexamethasone with the first
146 bronchodilator in the PED. We used an oral solution dexamethasone preparation of 2
147 mg/5ml concentration, with maximum 16 mg equating to 40 mls. In September 2018,
148 following recent evidence ¹⁴, we amended the protocol (supplementary file 3) to a single
149 300 mcg/kg dose of oral dexamethasone with a repeat 300 mcg/kg dose at admission
150 (16 mg maximum combined). Home dosages of dexamethasone were not prescribed.

151 Protocols were uploaded onto the PED intranet and project updates were relayed to
152 clinical staff at weekly departmental teaching.

153 **Record identification and selection**

154 Record identification

155 To identify relevant records, GD used the PED electronic management system
156 (Symphony ®) to extract all records for patients aged 2-14 years who attended for
157 medical assessment with respiratory complaints in October, November, and December
158 of years 2016, 2017, and 2018. Records were randomised using Microsoft Excel prior to
159 formal selection.

160 Record selection

161 In October 2019, MT and JR reviewed the randomised list and selected 100 attendance
162 records for each year (2016, 2017, and 2018 (total n=300)) based on the eligibility
163 criteria.

164 *Inclusion Criteria:*

- 165 1. Attended the RBHSC PED between 1st October 2016 and 31st December 2016, 1st
166 October 2017 to 31st December 2017, or 1st October 2018 to 31st December 2018.
167 2. Patients aged 2 years and over (to avoid bronchiolitis diagnoses), up to 14 years
168 (the upper departmental age limit).
169 3. Received OCS treatment for an asthmatic attack or EVW (defined as wheeze
170 caused by a viral infection).

171 *Exclusion Criteria:*

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172 1. Discharged home at triage (therefore, not requiring medication).

173 2. Left the department prior to medical assessment.

174**Data collection**

175We extracted information from the PED electronic database (Symphony ®) (MT & JR)

176and the Northern Ireland Electronic Care Record (NIECR) (BK, GD, & RM) using a bespoke

177data collection spreadsheet on Microsoft Excel. Symphony ® provided accurate arrival

178and departure times and complete clinical documentation, whilst the NIECR provided

179clinical records from primary and other secondary care units throughout Northern

180Ireland where children may have re-attended.

181We assigned patients to moderate or severe/life-threatening severity based on the

182National Institute Clinical Excellence (NICE) and Scottish Intercollegiate Guideline

183Network (SIGN) guidelines ^{3,4}; patients' vital signs (i.e. SpO₂, RR, and HR) and assessment

184findings (i.e. presence of difficulty speaking in full sentences, confusion, and silent chest)

185were used to grade severity based on the age-appropriate ranges. As a quality assurance

186measure, GD reviewed 10% of selected records and resolved any anomalies (such as

187missing documentation or data irregularities) by accessing original paperwork.

188**Statistical analysis**

189OCS tolerability, LOS, admission, re-attendance, and mortality data (aims 1, 2, & 3)

190*Analysis by year*

191We undertook two distinct approaches to the analysis. To assess changes in measures

192over time, we initially analysed data based on the years of attendance, grouping records

193according to 3 years (2016, 2017, and 2018). We calculated adherence, defined as

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194administration of the appropriate steroid type (and dosage in respect to
195dexamethasone) as per protocol for the respective period; for dexamethasone, we
196allowed for a +/- 10% dosage discrepancy (which is consistent with published literature)
197²², and accepted maximum doses of 16 mg as per protocol.

198*Analysis by OCS drug type and dosage*

199We noted that some patients in 2016 received dexamethasone, while some patients in
2002017 and 2018 received prednisolone. Therefore, to specifically assess the effects of the
201OCS regime, we grouped records for a second time, based on OCS drug type and dosage.
202Due to dosage heterogeneity (e.g., 450 mcg/kg of dexamethasone), we selected records
203within specific limits, which allowed for direct comparisons. All groupings are defined in
204Box 1.

205For brevity purposes, groupings by year were labelled accordingly: 2016, 2017, and
2062018. In addition, groupings by actual OCS type and dosage administered, irrespective of
207year, were labelled: PRED, DEX(600), and DEX(300).

208Data was analysed initially (by JR and RM) using Microsoft Excel. Further statistical
209analysis was completed (by VO) using IBM SPSS Statistics for Windows software, version
21026 (IBM Corp., Armonk, N.Y., USA). Results are reported using either mean \pm Standard
211Deviation (SD), or median and Interquartile range, where appropriate. For comparisons
212between two or more means, one-way analysis of variance (ANOVA) was used. Where
213data violated the ANOVA assumptions, the non-parametric equivalent Kruskal-Wallis H
214Test was used. Independent samples t-tests were used to compare means of two
215continuous variables. A Chi-squared test or Chi-squared test for trend was used to

216compare the distribution of categorical variables. For all tests, a p-value < 0.05 was
217considered to be statistically significant.

218Annual OCS drug-cost analysis (aim 4)

219We calculated the departmental savings incurred by OCSs (dexamethasone and
220prednisolone) in 2017 and 2018, compared to 2016. To calculate the number of patients
221receiving OCSs for acute wheeze annually, we searched Symphony ®, identified all
222patients presenting with respiratory symptoms in 2019 (January to December), and
223reviewed clinical documentation of 10 attendances for each month to determine the
224proportion receiving OSCs. Further detail is available in supplementary file 4.

225**Study approvals**

226The Belfast Health and Social Care Trust Audit Offices approved this as a QI intervention
227and research ethics was not required ²³.

228**Patient and public involvement**

229Patients and the public were not formally consulted in conceptualisation or design.

230RESULTS

231Demographics, severity, OCS used, and diagnoses

232Three-hundred records, 100 from October to December each year (2016, 2017, and
2332018), were randomly selected out of 2,435 potentially eligible (12.3%) children
234attending the RBHSC PED with respiratory complaints. Two records from 2016 referred
235to second attendances and were merged to avoid duplication. Data on demographics,
236severity, OCSs, and diagnoses for 2016, 2017, and 2018 are displayed in Table 1.

237The mean age was 4.4 years (range 2 to 14 years), 64% were male, and the most
238common diagnosis was EVW (57.0%). There was no statistical difference in severity
239between groups.

240Adherence to protocol

241There was adherence to the protocol in 80.0% and 90.0% of cases for 2017 and 2018,
242respectively. Non-adherence was mainly due to the alternative OCS being administered
243(prednisolone or dexamethasone) and underdosing of dexamethasone. Two cases in
2442016 received both prednisolone and dexamethasone. Figure 1 shows OCS treatment
245administered during 2016, 2017, and 2018.

246Outcomes for groupings by year and by OCS drug type and dosage are presented in
247table 2A and 2B, respectively.

248OCS tolerability (aim 1): improved tolerability, aim partly achieved

249In 2016, intolerability occurred in 12.2% of patients, compared to 3.0% in 2017, and
2505.0% in 2018, ($p = 0.057$); representing a 75.4% increase in OCS tolerability in the first
25112-months and 59.0% in the second 12-months using 2017 and 2018 protocols,

252respectively, compared to 2016. Therefore, the target (70% improvement in tolerability)
253was achieved at 12 months, but not sustained at 24 months. On comparing OCS type
254and dosages (irrespective of year), vomiting occurred in 10.1% of cases administered
255PRED compared to 3.6% and 5.8% in DEX(600) and DEX(300), respectively ($p = 0.101$).
256Anti-emetic usage was highest in 2017 at 5.0%.

257LOS in PED (aim 2): no change, aim not achieved

258Mean LOS in the PED was 3.6 hours in 2016, 4.2 hours in 2017, and 3.8 hours in 2018.
259The target (20% reduction to an average of 2.9 hours) was not achieved for either 2017
260and 2018, or for DEX(300) and DEX(600).

261Figure 2 illustrates OCS intolerability and LOS in PED, between groupings.

262Admission, re-attendance, and mortality rates (aim 3): no change, aim achieved

263The lowest admission rate (32.7%) was seen in 2016, followed by 2018 (35.0%), with the
264highest rate in 2017 (42.0%) ($p = 0.363$). Admission rates for patients receiving PRED and
265DEX(300) were similar, at 32.3% and 31.7% respectively; patients given DEX(600) had an
266admission rate of 45.2% ($p = 0.104$). There was no significant difference in re-attendance
267rates between the three years ($p = 0.149$). The lowest re-attendance rate occurred in
2682017; 3 patients reattended and 1 patient required admission at re-attendance. Nine
269patients reattended in 2016 and 2018; 7 and 0 respectively were admitted at re-
270attendance. The highest re-attendance rate (10.1%) was for those administered PRED,
271followed by DEX(300) (8.7%). Patients receiving DEX(600) had the lowest re-attendance
272rate of 2.4% ($p = 0.109$). Eight (80.0%) of the re-attending patients that were initially

273given PRED required admission, whereas none of the patients given DEX(300) or
274DEX(600) were admitted after re-attendance. No deaths occurred.

275**OCS drug costings (aim 4): reduced costs, aim achieved**

276The PRED regime was most expensive, followed by DEX(600) and DEX(300). For a 10kg
277child in September 2017, PRED at 1mg/kg (once per day for 3 days) costs £10.68,
278compared with £1.88 for DEX(600) and £0.94 for DEX(300). Every year, approximately
2792,096 patients aged 2-14 present to the PED with respiratory symptoms; approximately
28044.16% (n=926) receive OCSs for acute wheeze. Figure 3 illustrates annual departmental
281OCS drug costs over the 3 years. In 2016, the annual OCS cost of prednisolone,
282dexamethasone, and both OCSs for acute wheeze amounted to £23,966.02, £190.35,
283and £24,156.37, respectively. In 2017, the respective cost of prednisolone,
284dexamethasone, and both OCSs equated to £1,876.18, £2,778.91, and £4,655.09,
285producing savings of £19,501.28 (80.7%) and therefore achieving the 80% target. In
2862018, prednisolone, dexamethasone and both OCSs cost the department £277.40,
287£1,900.11, and £2,177.51, respectively, resulting in £21,978.87 (90.9%) savings, and
288achieving the 24-month target for OCS cost.

289**QIP feedback and challenges**

290Informal feedback from PED staff was positive; subjectively, juniors found the protocol
291useful and dexamethasone administration was less arduous for staff and patients. In
2922017, general practitioners (GPs), who received patient discharge plans as standard
293practice, noted the change in OCS treatment and contacted the RBHSC respiratory
294consultants for general advice.

295DISCUSSION

296Key Findings

297By implementing QI interventions, we increased OCS tolerability by 75.4% within the
298first 12 months and 59.0% within the second 12 months and achieved accumulative cost
299savings of £41,480.15 (85.8%) within 24 months. There was no significant difference in
300the LOS in the PED and low admission, re-attendance, and mortality rates were
301maintained.

302Interpretation

303Our findings are consistent with recent evidence, supporting dexamethasone as an
304alternative to prednisolone for acute wheezing attacks, with better tolerability and cost-
305effectiveness ¹⁴. Despite our hypothesis, better tolerability did not reduce the LOS;
306possibly because other factors (i.e., the time of day, day of the week, no. daily
307attendances, staffing levels, and seniority, etc.) also have an impact on patient flow,
308which we did not include in our analysis ²⁴. Non-adherence was mainly due to
309unfamiliarity with the departmental protocol, rounding up or down of liquid
310formulations, and confusing the dose with that used for croup (150 mg/kg).

311Strengths and limitations

312Our study adds to current evidence, because it provides a comprehensive and practical
313account of evidence-based QI interventions, delivering better tolerability and substantial
314cost-savings within a PED, and includes direct comparisons on OCS regimes. We took
315additional steps to ensure data accuracy yet acknowledge the limitations of
316retrospective data collection on data completeness and continual quality improvement

317efforts. Although GPs received patient discharge plans, we acknowledge that direct
318contact outlining prospective changes should have been made prior to implementation.
319Furthermore, surveys or similar formal feedback methods would have been informative
320to the change process, and therefore, should be the focus of future QI efforts.

321**Recommendations for future practice**

322We suggest, based on our findings, that PEDs and general EDs could improve tolerability
323and make significant cost savings using a departmental protocol directing the use of
324single-dose dexamethasone (300 mg/kg) for children with wheezing attacks; drug tariffs
325are nationally applicable, although medication costs vary internationally and are subject
326to pharmaceutical bids. Departments should liaise with ward staff and GPs to create a
327longer-term management protocol post ED discharge or admission, prior to QI
328implementation.

329CONCLUSION

330By implementing a structured acute pediatric wheeze protocol and modifying the OCS
331type and dose, we reduced vomiting by 67.2% over 24 months and achieved cost savings
332of £19,501.28 (80.7%) at 12 months and £41,480.15 (90.9%) at 24 months; whilst
333maintaining LOS, admission, re-attendance, and mortality rates. We would recommend
334that other departments implement similar QIPs to improve tolerability and reduce costs.

335 **List of abbreviations**

336cBNF- British National Formulary for Children

337ED- Emergency department

338EVW- Episodic viral wheeze

339GP- General Practitioner

340LOS- Length of stay

341NICE- National Institute of Clinical Excellence

342NIECR- Northern Ireland Electronic Care Record

343PED- Pediatric Emergency Department

344PRAM- Pediatric Respiratory Assessment Measure

345RBHSC- Royal Belfast Hospital for Sick Children

346SIGN- Scottish Intercollegiate Guidelines Network

347UK- United Kingdom

348 **Ethics approval and consent to participate**

349Not required as per Health Research Authority regulations.

350 **Consent for publication**

351Not applicable.

352 **Availability of data and materials**

353Data available upon request.

354**Authors' contributions**

355GD, BB, JM, & HS conceptualised the project. BB, JM, HS, & SM oversaw project
 356execution. GD, JR, MT, RM, & BK collected data and completed provisional analysis. VO
 357completed further statistical analysis. GD & SM supervised data collection and analysis.
 358GD & JR wrote the first draft and all authors contributed to the write-up. SM & MDS
 359provided critical revision. All authors read and approved the final manuscript.

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443FIGURE LEGENDS

444Figure 1 Oral corticosteroid medication prescribed

445Figure 2 Oral corticosteroid (OCS) intolerability (A) and length of stay (LOS) in the
446Pediatric Emergency Department (PED) (B) for groupings by year, and OCS type and
447dosage. OCS intolerability, spat-out OSC or vomited after administration while in the
448PED; LOS in PED, time from arrival until discharge home (excluding admissions).

449**Figure 3** Annual OCS drug costs (£) for acute wheeze, in the Pediatric Emergency
450Department in 2016, 2017, and 2018.