1	Physician Perspectives on Vaccination and Diagnostic Testing in Children with
2	Gastroenteritis: A Primary Care Physician Survey
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36	
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45 Introduction

Acute gastroenteritis remains a common cause of childhood mortality worldwide, claiming
approximately 578000 lives annually(1). Although mortality is very low in Canada, morbidity is
significant with an estimated 240000 children seeking emergency department (ED) care annually
on account of gastroenteritis(2). Societal costs are enormous as each ED visit costs ~\$800 and
hospitalized child ~\$2700(3).

51

52 Implementation of rotavirus vaccination programs has successfully resulted in a reduction in the 53 incidence of rotavirus-related ED visits, hospitalizations, and healthcare costs(4-7). However, a 54 universal rotavirus vaccination program had not been included in Alberta's publicly-funded 55 health-care system at the time of this study, despite endorsement by Canada's National Advisory 56 Committee on Immunization in 2010(8) (note: a vaccination program was implemented June 1, 57 2015). The delayed implementation of a program in Alberta could have been due to a lack of 58 knowledge of the local pathogen-specific gastroenteritis epidemiology and disease burden(9). 59 60 Identifying the pathogens responsible for gastroenteritis symptoms has traditionally been 61 challenging. The majority of affected patients do not seek medical care, stool samples are not

62 always requested from those who seek care, and compliance with specimen collection is

63 suboptimal(10). Diagnostic capabilities of routine laboratory testing and physician understanding

of these test results, is limited. These issues contribute to underreporting and a misrepresentationof disease burden.

66

67	Our objective was to ascertain physician understanding of the diagnostic tests used to identify
68	various enteric pathogens and the ease of sample collection methods. Such knowledge is needed
69	to drive changes in the diagnostic approach to gastroenteritis. Additionally, we sought to
70	measure physician support for the implementation of an enteric pathogen vaccination program.
71	

72 Methods

73 Survey development

74 Physicians and researchers with clinical and academic expertise from the areas of pediatrics (SF, 75 BL), family medicine (JD), emergency medicine (SF), and microbiology/virology/infectious 76 disease (ML, LC, XP, BL), collaborated to develop survey content. The survey was designed to 77 capture clinician knowledge and awareness of existing diagnostic tests used in Alberta and also 78 knowledge of "optimal" testing options available. Additionally, the survey aimed to obtain 79 perspectives on the integration of an enteric vaccine into the existing local vaccine schedule. The 80 final version of the survey was restricted to 30 questions and was designed for completion in 5 to 81 7 minutes. Before distribution, the survey was tested among 10 Alberta Provincial Pediatric 82 EnTeric Infection TEam (APPETITE) members for face validity, content, relevance, and time 83 required for completion. Based on the feedback provided, the survey was revised accordingly.

84

85 Survey setting and population

The province of Alberta has a population of 3.8 million and is divided into five healthcare zones
with two-thirds of the population residing in two urban zones (Calgary and Edmonton). The
survey was administered to physicians across all five healthcare zones
(http://www.albertahealthservices.ca/ahs-map-ahs-zones.pdf) who were members of Alberta's
Primary Care Networks (PCN), The Alberta Recording and ReseArch NeTwork (TARRANT),

91 and The Society of General Pediatricians of Greater Edmonton. These groups were selected as

92 they represent discrete target end-user physician groups who directly treat children with

93 gastroenteritis. They were also felt to potentially have higher response rates than from unselected

94 populations. Eligible participants had an email address and were members of one of the95 aforementioned networks.

96

97 Forty-six PCNs were listed on the Primary Care Initiative website when it was accessed in 98 October 2013. Seven of the PCNs were listed as "under development" or did not provide contact 99 information online and were excluded from this study. Therefore, 39 PCNs representing 2492 100 physicians were invited to participate. Of these 39 PCNs, 18 PCNs representing 1106 physicians 101 agreed to participate and confirmed sending out the survey to their physicians. Twenty-one PCNs 102 declined participation or did not confirm sending out the surveys. Four of the PCNs that declined 103 participation suggested that we contact their encompassing medical clinics directly, which 104 resulted in 35 physicians from 9 clinics being sent the survey link. Thus 1141 physicians, or 105 45.8% of the eligible PCN physician members, were contacted by these methods. 106 107 TARRANT is a province-wide influenza illness surveillance program and is composed of 108 approximately 50 voluntary participating primary care physicians. The Society of General 109 Pediatricians of Greater Edmonton is comprised of approximately 60 pediatricians. 110 111 **Survey distribution** 112 A study team member called each PCN and explained the purpose of the survey to administrative 113 personnel. PCNs that agreed to participate distributed the survey to their physicians with an 114 initial email containing the survey link, followed by three reminder emails spaced ~1 week apart. 115 PCN administrative staff was subsequently contacted to ensure that survey links were

116 distributed. Two participating PCNs distributed the link within their newsletter. In keeping with

117	the administrative infrastructure required by these PCNs, survey distribution was performed by
118	an administrator within each network, thereby prohibiting our group's ability to track emails and
119	reminders sent.

120

121	The head of the TARRANT network (JD) is a member of our research group and agreed to		
122	distribute the survey link and three reminder emails to all members. The chair of The Society of		
123	General Pediatricians of Greater Edmonton is affiliated with a member of our research group		
124	(BL), and agreed to distribute the survey link and three reminder emails to all members. Again,		
125	survey distribution was performed by an administrator within each network, thereby prohibiting		
126	our group's ability to track emails and reminders sent.		
127			
128	The final 30-question survey was administered between October 2013 and January 2014 using		

The final 30-question survey was administered between October 2013 and January 2014 using
 REDCapTM (Research Electronic Data Capture).

130

131 Data Analysis

132 Qualitative analysis of the data was performed. The results presented include respondent 133 demographics and perspectives on three domains: knowledge needs surrounding diagnostic tests 134 to detect pathogens, stool sample collection methods, and implementation of enteric virus 135 vaccination programs. Responses are summarized as frequencies. Likert scale responses 136 (strongly agree, agree, neither agree nor disagree, disagree, strongly disagree) were offered to 137 collect respondents' perspectives. While the raw data is presented in tabular format, manuscript 138 text grouped together "strongly agree" with "agree," and "strongly disagree" with "disagree" to 139 facilitate and clarify the interpretation of the responses received.

140

141 **Results**

142 **Demographics**

- 143 Of the 1251 physicians who were provided the survey link, 92 (7.4%) completed the survey,
- 144 including 78 family physicians, 9 pediatricians, 3 physicians who indicated their field of training
- as "other," and 2 physicians who abstained from answering this question. Surveys were collected
- 146 from 18 physicians (19.6%) from the North zone, nine (9.8%) from the Edmonton zone, five
- 147 (5.4%) from the Central zone, 52 (56.5%) from the Calgary zone, and eight (8.7%) from the
- 148 South zone. Fifty-eight percent (53/91) of respondents described their practice as "all urban" or
- 149 "mostly urban".
- 150

151 Knowledge needs

152 To identify self-perceived gaps related to diagnostic testing modalities, respondents evaluated

their comfort with ordering tests to identify bacteria, viruses, and parasites in stool (Table 1).

154 Ninety-one percent (81/89) of respondents claimed they are aware of the tests available to

155 identify bacterial and parasitic pathogens. Fifty-one percent (44/87) reported they possess

156 sufficient knowledge to order tests to identify viruses. Seventy-six percent (68/90) of

157 respondents believe that enhancing their knowledge of viral pathogens would enable them to

158 provide better care and guidance.

159

160 Respondent physicians were asked to identify diagnostic tests that a) are currently used and b)

are optimal, to identify stool pathogens (Table 2). Although EM and EIA were used to evaluate

162 viruses in stool samples in Alberta during the study period (11), only 20% and 48% of

163 respondents identified EM and EIA respectively as tests they could request for these purposes.

164 Culture was correctly identified by 99% of respondents as the available test to identify bacterial 165 stool pathogens, however, culture was also incorrectly chosen as a modality to diagnose viruses 166 (74%), parasites (47%), and C. difficile (67%) in stool. Additionally, PCR is the diagnostic 167 modality with the greatest ability to identify all discussed stool pathogens, but was only selected 168 as the best modality to identify bacteria, viruses, parasites, and C. difficile by 22%, 40%, 8%, and 169 29% of respondents respectively. 170 171 **Stool sample collection** 172 Though 77% (69/90) of respondents agreed that children with diarrhea and vomiting can be seen 173 on the same day in their practice, 62% (55/89) believe that current methods of stool sample 174 collection for diagnostic testing are inconvenient, and 82% (72/88) indicated that specimen 175 submission rates would improve if rectal swab samples could be submitted for testing. 176 Furthermore, 40% (36/90) of physicians agreed that current stool sample testing methods lack 177 sensitivity and do not provide results in a timely manner (Table 3). 178 179 Vaccination perceptions (Table 4) 180 Seventy-three percent (66/90) of physician support an enteric vaccine program however, 82% 181 (71/87) stated that a requirement for the addition of such a vaccine into the provincial 182 vaccination schedule should include evidence of cost-effectiveness.

183

185 The survey indicated there is room for improvement in the understanding of available and 186 optimal diagnostic test options to detect enteric pathogens. Although the vast majority of 187 respondents claim that they possess adequate knowledge of the tests available to identify 188 bacterial and parasitic pathogens, only half claim they are aware of tests to identify viruses. This 189 disparity uncovers a need for increased knowledge of viral pathogens and the methods by which 190 they can be diagnosed. Limited testing and identification of fecal viral pathogens in Alberta may 191 be connected to knowledge gaps we identified regarding diagnostic tests. 192 193 Diagnostic testing based on stool collection impedes pathogen identification in community 194 practice. Our survey identified that physicians believe that rectal swab use would increase 195 convenience and sample collection rates. Such an approach has been used to identify enteric 196 pathogens and it may be as sensitive as stool(12). Rectal swab use could therefore improve our 197 understanding of the pathogen-specific burden of disease. 198 199 Another target for improvement relates to microbiology requisitions which often require 200 physicians to select tests from a list. Such a process does not directly ask what organism 201 physicians are seeking or what the clinical context is – such approaches could better link clinical 202 needs and microbiologic tests. Given that most clinicians have limited knowledge of likely 203 pathogens and recent advances in microbiologic testing procedures, a gap is created between 204 clinical needs and the tests requested. This challenge has been identified(13) and an alternative 205 to changing the approach to test ordering may be the use of assays that test for a broad range of 206 the most common disease causing pathogens. One such example is the

207 The Luminex Gastrointestinal Pathogen Panel (xTAG(®) GPP) that detects in one assay 208 adenovirus 40/41, norovirus genogroup I/II, rotavirus A, *Clostridium difficile* toxin A/B, 209 Campylobacter sp., Escherichia coli O157, Enterotoxigenic E. coli heat-labile enterotoxin/heat-210 stable enterotoxin, Salmonella sp., Shiga-toxin producing E. coli, Shiga-like toxin (Stx)1/2, 211 Shigella sp., Vibrio cholerae, Yersinia enterocolitica, Cryptosporidium sp., Entamoeba 212 histolytica and Giardia sp. However, such approaches also present challenges for clinicians with 213 respect to interpretation. 214 215 We found that 73% of respondent physicians support the addition of a gastroenteritis vaccine

216 into Alberta's vaccination schedule. Our survey adds to the existing literature which reported that 217 only 53% of Canadian pediatricians recommend the administration of a rotavirus vaccine and 218 59% endorse the implementation of a publically funded vaccination program(14). A recent study 219 reported that Canadian physician endorsement of rotavirus vaccination is the lowest of the seven 220 vaccines evaluated (15). Our findings demonstrate that physician endorsement of a gastroenteritis 221 vaccine is far from universal. This knowledge is important since physician opinion plays a major 222 role in vaccine uptake(16). One possible explanation for the low endorsement rate may relate to 223 the limited ability to identify a pathogen and hence a lack of direct connection between episodes 224 of rotavirus and severe gastroenteritis symptoms. Additionally in Canada, rotavirus infections are 225 associated with very low mortality rates compared to elsewhere in the world.

226

227 Limitations of this study include a very low response rate, which may have resulted in

respondent bias. Although physicians across all of Alberta's five healthcare zones participated,

selection bias must always be considered as physicians with strong opinions might have been

230	more likely to participate. Family physicians and pediatricians were invited to participate in the
231	survey in an attempt to promote the generalizability however this approach also introduced
232	heterogeneity. In Alberta these two groups of physicians along with emergency room physicians
233	directly treat children with gastroenteritis, and therefore combining the groups is appropriate.
234	Lastly, sub-analyses by group were planned, but due to the low response rate they were not
235	practical.
226	

236

238 Conclusions

- 239 Challenges with stool collection and test ordering could minimize the perception of the
- 240 pathogen-specific burden of disease and may contribute to limited vaccine support. Our data
- support the notion that simplified stool testing and collection procedures may improve our
- knowledge and understanding of the value of immunization, while enhancing the ability to
- 243 identify enteric infections in children.

244

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