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**Background.** As diarrheal stool samples are the recommended specimen for testing in acute gastroenteritis (AGE), etiological investigations are rarely performed in children presenting with isolated vomiting. This study identifies enteropathogens in children with AGE presenting with isolated vomiting.

**Methods.** Children <18 years old with ≥3 episodes of vomiting/diarrhea in 24 hours and <7 days of symptoms were recruited in 2 pediatric emergency departments, a public health clinic and via Health Link, a provincial nurse advice phone line. Rectal swabs and stool samples were collected and tested using the Luminex xTAG GPP, an in-house 5-virus RT-qPCR panel and enteric bacterial culture. Vomiting and diarrhea data were collected at enrollment (day 0) and at day 14.

**Results.** Between Dec 9, 2014 and Apr 14, 2016, 2,184 children were enrolled and tested: 784 (36%) presented with isolated vomiting, 250 (11%) with isolated diarrhea (ID), 1,138 (52%) with both vomiting and diarrhea (V&D), 12 had missing data. The detection of enteropathogens was 56% when presenting with isolated vomiting, 55% with ID and 83% with V&D. Of the 784 children with isolated vomiting, 54% (n = 424) had one or more viruses: the most common was norovirus (NoV) (n = 244, 50%), followed by adenovirus (Adv) (91, 19%), rotavirus (Rota) (57, 12%), sapovirus (84, 17%) and astrovirus (10, 2%). Fifty-eight cases had >1 virus; co-infection with NoV and Adv was the most common (n = 23). Ten of these 424 patients also had enteric bacteria (2 *Aeromonas*, 2 ETEC, 2 *Salmonella*, 2 *Yersinia*, 1 *Campylobacter*, 1 *E. coli* O157) and 8/9 (89%) of these patients reported development of diarrhea at day 14. In comparison, 212/383 (55%) of patients with virus only reported diarrhea at follow up. Enteric bacteria with no virus was detected in 11 patients (3 *Aeromonas*, 3 *Salmonella*, 3 STEC, 1 *Campylobacter*, 1 *E. coli* O157) and 3/10 of these patients reported diarrhea.

**Conclusion.** Over 50% of AGE presented with isolated vomiting had enteric virus identified in stool or rectal swabs, representing a significant pathogen-based disease burden not previously included in healthcare planning (e.g., Rota vaccine). NoV was the predominant agent followed by Adv and Rota. Finding enteric bacteria in these cases is novel and requires further study.

**Disclosures.** All authors: No reported disclosures.

### 1181. Enteropathogen Identification by Multiplex PCR in Guatemalan Children with Acute, Non-bloody Diarrhea

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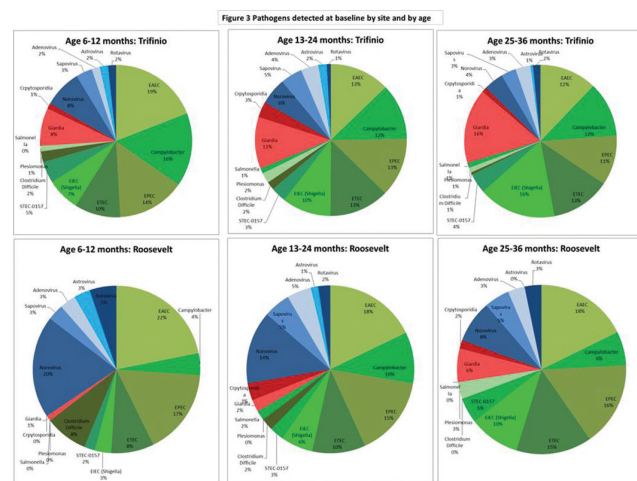
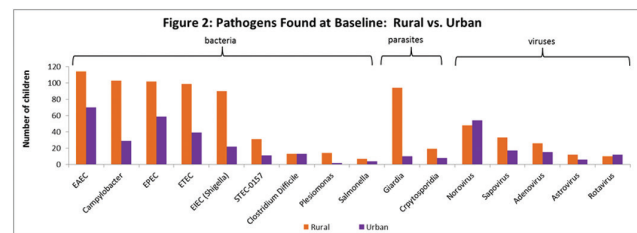
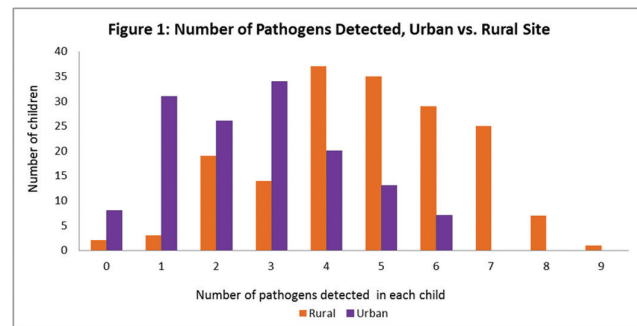
**Background.** Diarrhea is a leading cause of morbidity and mortality in children in low and middle income countries (LMICs). Assessing diarrhea etiology in LMICs is of great importance in order to better develop both therapeutic and public health strategies, but is hampered by the complexity of potential diarrheal pathogens, and diverse methodology needed for pathogen identification

**Methods.** Subjects 6 to 35 months old with acute, moderate severity, non-bloody diarrhea were enrolled in a diarrheal treatment trial, conducted at one rural (N = 172) and two urban sites (N = 144) in Guatemala. Diarrheal pathogens were determined in stool by multiplex PCR (FilmArray GI<sup>®</sup> Biofire) which allows simultaneous identification of 23 bacterial, viral, parasitic pathogens. Descriptive statistics on demographics, pathogen load, and differences in pathogen occurrence by site were performed; differences were assessed with t-test and chi<sup>2</sup> test

**Results.** Nearly all (96.8%) subjects had pathogens identified, and most had multiple potential pathogens identified (mean pathogen count: 2.7 urban and 4.8 rural; P < 0.001 (Figure 1). Notable pathogen differences were observed between rural and urban populations. Bacteria (particularly *E. coli* pathotypes and *Campylobacter*) and protozoa (particularly giardia) were more common in the rural population (Figure 2). Viral pathogens were either similar or more common (norovirus; P = 0.04) in the urban population; rotavirus was uncommon in both sites (10 rural and 12 urban cases). A similar pattern of pathogen evolution with patient age was noted in both settings, with a decrease in the relative number of viral and increase in parasitic pathogens (Figure 3). Important demographic and socioeconomic differences between rural and urban were noted: rural subjects had poorer nutritional status, underdeveloped water and sanitation facilities and more domestic animal exposure

**Conclusion.** Acute diarrheal episodes in Guatemalan children were associated with a complex spectrum of pathogens when determined by multiplex PCR, with distinct patterns in rural and urban populations. Future studies to precisely determine

diarrheal etiologies in LMICs will need to incorporate controls to sort causative organisms from those colonizing the intestine.



**Disclosures.** All authors: No reported disclosures.

### 1182. Appropriateness of a Rapid Multiplex Gastrointestinal Panel in the Investigation of Suspected Infectious Diarrhea After Implementation at an Academic Medical Center

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**Background.** The BioFire FilmArray<sup>™</sup> Gastrointestinal (GI) Panel is a 1 hour multiplex real-time PCR test that can detect the presence of 22 GI pathogens (viral, bacterial, and parasitic) known to cause infectious diarrhea. Our tertiary-care academic medical center implemented the GI Panel for all cases of suspected infectious diarrhea replacing the previous conventional testing once utilized to detect GI pathogens. Since its implementation we have not had any criteria for ordering this test to aid healthcare providers.

**Methods.** The aim of this IRB approved, retrospective investigation was to determine the appropriateness of ordering the GI panel at our academic institution. Cases were randomly selected, stratified by age group and result (specific pathogens or negative result) from May 2015 through April 2016 in the post-implementation period (n = 400 of 1117 total tests). We developed appropriateness criteria for ordering the GI panel which included: passage of at least 3 unformed stools in 24 hours plus one

or more enteric symptom (nausea, vomiting, abdominal pain/cramps, tenesmus, fecal urgency, moderate to severe flatulence), and one of the following: grossly bloody diarrhea (dysenteric), persistent diarrhea (14 – 30 days), worsening or relapsing diarrhea, fever  $\geq 101$  F°, severe diarrhea > 10 bouts in 24hrs, immunosuppression, pregnancy, food handler, infant < 1 year and their care takers, age  $\geq 65$  years old, concern for disseminated GI infection, with no previous GI panel testing in the past 30 days.

**Results.** Overall appropriateness of GI panel testing based off our generated criteria was 36% ( $n = 144/400$ ). This included all tests ordered in the outpatient clinics, emergency department, inpatient medical/surgical wards, and intensive care units.

**Conclusion.** Currently there is not a well-established standard criteria for ordering the GI panel for investigating suspected infectious diarrhea. After implementation at our academic tertiary-care medical center the GI panel was used inappropriately in most cases without a criteria for ordering in place to aid clinicians. Educating healthcare providers about appropriate testing indications is being performed. Further studies are needed to assess if our generated criteria will lead to decreased costs and unnecessary testing.

**Disclosures.** All authors: No reported disclosures.

### 1183. Clinical Predictors of *Shigella* and *Campylobacter* Infection in Children in the United States

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**Background.** Infectious gastroenteritis is a major cause of morbidity and mortality among children worldwide. While most episodes are self-limiting, for select pathogens such as *Shigella* and *Campylobacter*, etiological diagnosis may allow effective antimicrobial therapy and aid public health interventions. Unfortunately, clinical predictors of such pathogens are not well established and are based on small studies using bacterial culture for identification.

**Methods.** We used prospectively collected data from a multi-center study of pediatric gastroenteritis employing multi-pathogen molecular diagnostics to determine clinical predictors associated with 1) *Shigella* and 2) *Shigella* or *Campylobacter* infection. We used machine learning algorithms for clinical predictor identification, then performed logistic regression on features extracted plus pre-selected variables of interest.

**Results.** Of 993 children enrolled with acute diarrhea, we detected *Shigella spp.* in 56 (5.6%) and *Campylobacter spp.* in 24 (2.4%). Compared with children who had neither pathogen detected (of whom, >70% had  $\geq 1$  potential pathogen identified), bloody diarrhea (odds ratio 4.0), headache (OR 2.2), fever (OR 7.1), summer (OR 3.3), and sick contact with GI illness (OR 2.2), were positively associated with *Shigella*, and out-of-state travel (OR 0.3) and vomiting and/or nausea (OR 0.4) were negatively associated (Table). For *Shigella* or *Campylobacter*, predictors were similar but season was no longer significantly associated with infection.

**Conclusion.** These results can create prediction models and assist clinicians with identifying patients who would benefit from diagnostic testing and earlier antibiotic treatment. This may curtail unnecessary antibiotic use, and help to direct and target appropriate use of stool diagnostics.

Feature	Shigella or Campylobacter not detected (n=913)	Shigella detected (n=56)		Shigella or Campylobacter detected (n=80)	
	N (%)	N (%)	OR (95% CI)	N (%)	OR (95% CI)
Bloody diarrhea	91(10.0)	15(26.8)	<b>4.0 (1.9-8.5)</b>	23(28.8)	<b>3.8 (2.1-6.9)</b>
Fever	458(50.1)	47(83.9)	<b>7.1 (2.8-18.2)</b>	62(77.5)	<b>5.1 (2.4-10.5)</b>
Headache	236(25.9)	24(42.9)	<b>2.2 (1.2-4.2)</b>	35(43.8)	<b>2.4 (1.4-4.0)</b>
Vomiting/nausea	726(79.5)	34(60.7)	<b>0.4 (0.2-1.0)</b>	49(61.3)	<b>0.5 (0.3-1.1)</b>
Sick Contact with GI illness	206(22.6)	17(30.4)	<b>2.2 (1.1-4.3)</b>	25(31.3)	<b>2.1 (1.2-3.6)</b>
Out of state travel	131(14.4)	4(7.1)	<b>0.3 (0.1-0.95)</b>	5(6.3)	<b>0.3 (0.1-0.75)</b>
Summer season	254(27.8)	34(60.7)	<b>3.3 (1.7-6.5)</b>	35(43.8)	<b>1.6 (0.9-2.7)</b>

**Disclosures.** A. Leber, BioFire Diagnostics: Research Contractor and Scientific Advisor, Research support, Speaker honorarium and Travel expenses; J. Daly, Biofire: Grant Investigator, Grant recipient; R. Selvarangan, BioFire Diagnostics: Board Member and Investigator, Consulting fee and Research grant; Luminex Diagnostics: Investigator,

Research grant; J. Dien Bard, BioFire: Consultant and Investigator, Research grant and Speaker honorarium; K. Holmberg, BioFire Diagnostics: Employee, Salary; K. Bourzac, BioFire Diagnostics: Employee, Salary; K. C. Chapin, BioFire Diagnostics: Investigator, Research support; A. Pavia, BioFire Diagnostics: Grant Investigator, Research grant

### 1184. High Genetic Variability of Norovirus Leads to Diagnostic Test Challenges

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**Background.** It is important to understand the diagnostic accuracy of syndromic multiplex panels such as the Luminex xTAG<sup>®</sup> Gastrointestinal Pathogen Panel (GPP) as they are increasingly employed as routine diagnostic tests in laboratories worldwide. Recent evaluations in our laboratory identified lower detection rates of norovirus genogroup II (NoV GII) using the GPP as compared with our lab-developed RT-qPCR Gastroenteritis Virus Panel (GVP). This study is to characterize the NoV strains in samples with discordant NoV GII test results between GPP and GVP and determine the sensitivity of the two assays for specific NoV GII genotypes.

**Methods.** We genotyped all NoV GII strains with discordant test result in stool samples or rectal swabs collected prospectively from a cohort of children with acute gastroenteritis between December 2014 and July 2016. The sensitivity of GVP and GPP for NoV GII were compared by analyzing GVP threshold cycle (Ct) and using ten-fold serial dilutions of positive samples of various NoV GII genotypes.

**Results.** All discordant samples (11%; 63/607) tested positive for NoV GII by GVP but negative by GPP. Thirty-five percent (22/63) were successfully genotyped; 64% (14/22) of those were NoV GII genotype 2 (GII.2). The median Ct value of concordant positive was lower than those with discordant results (19.8 vs. 33.7 respectively;  $P < 0.0001$ ). GVP was 10-fold and at least 10,000-fold more sensitive than GPP in detecting NoV GII.3 and GII.2, respectively, but has similar sensitivity for NoV GII.4. The GII.2 variants with discordant test results differed genetically from the concordant GII.2 variants.

**Conclusion.** GPP has suboptimal sensitivity to detect NoV GII.2 and its use may lead to an underestimation of NoV disease burden with some cases not being detected.

**Disclosures.** All authors: No reported disclosures.

### 1185. Risk Factors and Clinical Outcomes of Cancer Patients with *Clostridium difficile* Associated Diarrhea Co-infected with a Second Enteropathogen

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**Background.** Cancer patients are at an increased risk for *C. difficile* infection (CDI) which is often identified along with other enteropathogens. The impact of co-infections on outcomes has not been established in this population. We compared the risk factors and clinical characteristics of patients with CDI mono-infection (CDIM) and patients coinfected with bacterial (CDIB) or viral (CDIV) enteropathogens.

**Methods.** Adult patients presenting with primary or recurrent CDI ( $n = 88$ ) identified on a two-step GI multiplex assay (Biofire) followed by toxin A/B EIA, were classified into CDIM ( $n = 66$ ), CDIB ( $n = 12$ ), and CDIV ( $n = 10$ ) groups. Demographic and clinical data were collected and risk factors and outcomes compared by Fisher's exact test, ANOVA, and the Kruskal-Wallis test. CDI severity was determined using Zar's criteria, presence of bacteremia, and ICU stay.

**Results.** During the study period, 2,017 diarrheal samples were submitted to the microbiology laboratory. An enteric pathogen was identified in 311 (15%) patients. CDI was identified in 88 cases of which 22 (25%) had a second pathogen. CDIM was found in 66 (21%), CDIB in 12 (4%), and CDIV in 10 (3%) subjects. The most common co-pathogens identified were diarrheagenic *E. coli* in the CDIB group (9/12, 75%) and norovirus in the CDIV group (8/10, 80%). Groups were similar in terms of demographics, number of recurrences, health care acquisition, co-morbidities, disease severity, serum creatinine at presentation, presence of toxin by EIA, and mortality. Patients with CDIM were more likely to have a recent hospitalization than the CDIB group (44/66 67% vs. 3/12 25%,  $P = 0.01$ ). Clinical symptoms at presentation were similar for the three groups except for nausea which was more common in the CDIV group when compared with CDIM (8/10, 80% vs. 25/66, 38%;  $P = 0.02$ ). The use of proton pump inhibitors was similar in the three groups. There was however, a higher proportion of patients taking GABA-like drugs within 90 days among the CDIB patients (10/12, 83%) than the group with CDIM (26/66, 40%)  $P = 0.01$ .