Sapovirus Outbreaks in Long-Term Care Facilities, Oregon and Minnesota, USA, 2002–2009

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We tested fecal samples from 93 norovirus-negative gastroenteritis outbreaks; 21 outbreaks were caused by sapovirus. Of these, 71% were caused by sapovirus genogroup IV and 66% occurred in long-term care facilities. Future investigation of gastroenteritis outbreaks should include multi-organism testing.

Viral gastroenteritis outbreaks are associated with illness and death when they occur in institutional settings, notably in long-term care facilities (LTCFs) for the elderly (1). Although most reported outbreaks in LTCFs are caused by norovirus (2), some have similar epidemiologic characteristics but are norovirus-negative after ≥2 fecal samples are tested by real-time reverse transcription PCR (RT-PCR). Epidemiologically, these norovirus-like gastroenteritis outbreaks are characterized by 24–48-hour incubation periods, if known; vomiting in ≥50% of affected persons; and 12–60-hour median illness durations (3).

Norovirus and sapovirus are separate genera of the family Caliciviridae. Sapovirus was first detected in 1977 as the cause of a gastroenteritis outbreak in a home for infants in Sapporo, Japan (4), and was thereafter reported primarily among young children with acute gastroenteritis (5). After sapovirus RT-PCR was developed (6), sapovirus outbreaks were discovered in LTCFs and other settings populated by adults (7–9). Sapovirus genogroups I, II, IV, and V (GI, GII, GIV, and GV, respectively) infect humans (10). This report describes sapovirus outbreaks in Oregon and Minnesota, USA, during 2002–2009.

The Study

The Oregon and Minnesota state public health departments investigated 2,161 gastroenteritis outbreaks reported during 2002–2009. Of these, 1,119 (52%) were caused by laboratory-confirmed norovirus (defined as ≥2 norovirus-positive fecal samples by RT-PCR); 466 (22%) were caused by bacteria, parasites, and other agents; 403 (19%) had no fecal samples to analyze; 142 (7%) were norovirus negative (defined as ≥2 norovirus-negative fecal samples by RT-PCR) and, when tested, were negative for enteric bacterial pathogens; and 31 (<1%) had a single norovirus-negative stool sample. Outbreak-related fecal samples were archived when any specimen remained after analysis, creating a convenience sample of feces for this and other studies.

The Minnesota Public Health Laboratory tested feces from 93 (66%) of the 142 norovirus-negative outbreaks with RT-PCRs for astrovirus, adenovirus, rotavirus, norovirus, and sapovirus (6). Sapoviruses were genotyped by sequence analysis of the capsid gene (11).

Defining a sapovirus outbreak in this study as ≥1 sapovirus-positive fecal sample, 21 (23%) of the 93 norovirus-negative outbreaks were found to be caused by sapovirus. Adenovirus or norovirus were also identified in 4 (19%) of the 21 sapovirus outbreaks (Table 1). The unexpected norovirus finding is likely due to slight variations in testing methods between state public health laboratories and viral loads nearing the detection level of the RT-PCR.

Of 21 sapovirus outbreaks, LTCFs accounted for 12 (66%); grade schools for 2 (10%); and a prison, a large psychiatric hospital, a cruise ship, a restaurant, and a bed and breakfast for 5 (24%). During 2007, 10 outbreaks (48%) occurred; 14 outbreaks (67%) occurred during the colder months (November–March) of each observed year. Person-to-person transmission accounted for 18 (86%) of 21 outbreaks. On the basis of the outbreak setting, foodborne transmission was suspected, but not confirmed, in 3 (14%) of 21 sapovirus outbreaks; food items were not implicated. Outbreaks involved 5–44 persons (median 34 persons) per outbreak and lasted 1–28 days (median 15 days) (Table 2).

Clinical data were available for 141–269 patients from 14 sapovirus outbreaks in which neither adenovirus nor norovirus were identified. Of 141 patients, 32 (23%) had fevers. Of 269 patients, 132 (49%) had vomiting, and 238 (88%) had diarrhea (Table 2). In Oregon, 1 person with sapovirus was hospitalized and 1 died; no hospitalizations or deaths occurred in Minnesota among persons with sapovirus. Symptoms lasted 24–105 hours (median 48 hours) (data not shown).

Four (19%) of 21 sapovirus outbreaks were caused by sapovirus GI, 1 (5%) by sapovirus GII, 15 (71%) by...
### Table 1. Microbiology of 21 sapovirus outbreaks, Oregon and Minnesota, USA, 2003–2009*  

| State | Outbreak no. | Fecal samples, no. | Genotype Results |
|-------|--------------|--------------------|------------------|
| MN    | 2002–438     | 1                  | IV Sapovirus only |
| MN    | 2002–439     | 3                  | IV Sapovirus only |
| MN    | 2003–644     | 1                  | II Sapovirus only |
| OR    | 2004–066     | 1                  | V Sapovirus only  |
| MN    | 2006–924     | 2                  | IV Sapovirus only |
| OR    | 2007–001     | 3                  | IV Sapovirus only |
| OR    | 2007–013     | 3                  | IV Sapovirus only |
| OR    | 2007–023     | 3                  | IV Sapovirus only |
| OR    | 2007–028     | 4                  | IV Sapovirus only |
| OR    | 2007–039     | 4                  | IV Sapovirus only |
| OR    | 2007–046     | 4                  | IV Sapovirus only |
| OR    | 2007–086     | 4                  | IV Sapovirus only |
| OR    | 2007–091     | 4                  | IV Sapovirus only |
| OR    | 2007–221     | 1§                 | I Sapovirus only  |
| OR    | 2007–228     | 1                  | I Sapovirus only  |
| OR    | 2008–109     | 1                  | I Sapovirus only  |
| OR    | 2008–128     | 3                  | I Sapovirus only  |
| MN    | 2008–1308    | 3                  | I Sapovirus only  |
| MN    | 2008–1327    | 3                  | IV Sapovirus only |
| OR    | 2009–146     | 3                  | IV Sapovirus only |
| OR    | 2009–167     | 2                  | IV Sapovirus only |

*MN, Minnesota; OR, Oregon; G, genogroup.  
†One norovirus GI–positive sample.  
‡One adenovirus-positive sample.  
§Norovirus GII co-infection.  
¶Two adenovirus-positive samples.

### Table 2. Descriptive epidemiology of 21 sapovirus outbreaks, Oregon and Minnesota, USA 2002–2009*  

| Infection and state | Outbreak no. | Setting | Transmission | Outbreak features | Symptoms, no. patients |
|---------------------|--------------|---------|--------------|-------------------|------------------------|
| Sapovirus only      | MN 2002–438 | Grade school | Person-to-person | 2002 Apr 11 15 NA NA NA |
|                     | MN 2002–439 | Long-term care | Person-to-person | 2002 Apr 1 34 NA NA NA |
|                     | MN 2003–644 | Grade school | Person-to-person | 2003 Dec 8 17 NA NA NA |
|                     | OR 2004–066 | Long-term care | Person-to-person | 2003 Mar 17 44 23 44 8 |
|                     | MN 2006–924 | Long-term care | Person-to-person | 2006 Feb 13 24 9 24 11 |
|                     | OR 2007–013 | Long-term care | Person-to-person | 2007 Jan 15 12 7 9 NA |
|                     | OR 2007–023 | Long-term care | Person-to-person | 2007 Jan 7 35 16 33 3 |
|                     | OR 2007–028 | Long-term care | Person-to-person | 2007 Jan 9 12 7 5 6 |
|                     | OR 2007–039 | Long-term care | Person-to-person | 2007 Jan 22 14 8 12 2 |
|                     | OR 2007–046 | Long-term care | Person-to-person | 2007 Jan 5 15 11 15 0 |
|                     | OR 2007–086 | Long-term care | Person-to-person | 2007 Feb 13 8 6 7 NA |
|                     | OR 2007–221 | Long-term care | Person-to-person | 2007 Nov 10 34 14 27 NA |
|                     | OR 2008–109 | Long-term care | Person-to-person | 2008 Aug 1 5 3 5 NA |
|                     | MN 2008–1308 | Cruise ship | Foodborne suspected | 2008 Aug 1 5 3 5 NA |
|                     | MN 2008–1327 | Bed and breakfast | Foodborne suspected | 2008 Nov 3 7 2 7 2 |
|                     | OR 2009–146 | Psychiatric hospital | Person-to-person | 2009 Jul 9 13 9 11 NA |
|                     | OR 2009–167 | Long-term care | Person-to-person | 2009 Aug 11 22 7 18 NA |
| Sapovirus and norovirus | OR 2007–001 | Prison | Person-to-person | 2006 Dec 23 154 70 119 1 |
|                     | OR 2007–221 | Long-term care | Person-to-person | 2007 Nov 16 34 8 29 NA |
| Sapovirus and adenovirus | OR 2007–081 | Long-term care | Person-to-person | 2007 Feb 13 25 15 25 NA |
|                     | OR 2008–128 | Restaurant | Foodborne suspected | 2008 Apr 4 26 10 25 NA |

*MN, Minnesota; OR, Oregon; NA, data were not collected or could not be analyzed.  
†Median no. days: 15 (range 1–28 days).  
‡Laboratory-confirmed and epidemiologically linked cases with systematically collected symptoms; these are not complete case counts. Median no. cases: 34 (range 5–44 cases).  
§Of 269 patients, vomiting was reported for 132 (49%) and diarrhea for 238 (88%).  
¶Of 141 patients, fever was reported for 32 (23%).
Sapovirus Outbreaks

Sapovirus GIV, and 1 (5%) by sapovirus GV (Table 1). The genogroup-specific differences between outbreak settings and between the proportions of vomiting, diarrhea, and fever were not statistically significant. Seventy-three percent of sapovirus GIV outbreaks occurred in 2007. A representative sequence from each outbreak was placed in the phylogenetic tree (Figure). Of 14 sapovirus outbreaks with ≥2 sapovirus-positive samples, sequences from 12 were identical within the outbreaks, and 2 had ≈2 different sequences (Figure).

Conclusions
In this study, the high (66%) proportion of sapovirus outbreaks in LTCFs among 21 outbreaks of previously unknown etiologies is likely to be an artifact of legally mandated outbreak reporting by health care facilities rather than the true distribution of sapovirus outbreaks in Oregon and Minnesota. Still, elderly residents of LTCFs are especially vulnerable to rapid transmission of viral enteric pathogens and serious complications from infection with these agents (12), and therefore merit the attention of public health.

Our data, together with a recent study in Canada (7), demonstrate that sapovirus has been circulating among the institutionalized elderly since at least 2002 and that sapovirus outbreaks increased in 2007 as part of a worldwide surge in gastroenteritis outbreaks (2,7,9). Before these retrospective studies, sapovirus infections among adults ≥65 years old had been reported as single cases at a low (3%) rate in 2002 (13) and as nosocomial outbreaks in 2010 and 2005 (8,14). In 2010, Svraka et al. reported an age distribution shift from younger to older persons (9).

Sapovirus outbreaks occurred in the same settings and had the same seasonal distribution as norovirus outbreaks (2,15). Our study adds clinical details to information provided by studies in Canada and Europe (7,9). The clinical profile of sapovirus outbreaks in this study (49% vomiting, 88% diarrhea, and 23% fever, plus a median duration of 48 hours) approximates the criteria of Kaplan et al. (3), which are still used to evaluate norovirus outbreaks when laboratory resources are limited. We found, however, that sapovirus and norovirus outbreaks are clinically and epidemiologically similar enough to be indistinguishable without laboratory testing.

This study has at least 3 limitations. First, testing a convenience sample of fecal specimens from norovirus-negative outbreaks might have introduced selection bias, the impact of which is uncertain. Second, because outbreak reporting from institutions other than LTCFs is not legally mandated, outbreaks in these settings are underreported. Third, feces from norovirus-positive outbreaks were not assayed for sapovirus. Previously undetected norovirus GI and GII discovered among 21 sapovirus outbreaks indicates that outbreaks might have had >1 etiology. It is therefore likely that the number of sapovirus outbreaks was underestimated.

In summary, gastroenteritis outbreaks in LTCFs should be investigated by public health departments in conjunction with GenBank sequences of well-characterized genogroups were aligned with outbreak sequences, and a phylogenetic tree was created by the neighbor-joining method by using BioNumerics (Applied Maths, Austin, TX, USA). Genotypes were assigned on the basis of >95% similarity to reference strains. Outbreak strain sequences were deposited in GenBank under accession nos. HM800902–HM800920.

![Figure. Phylogenetic tree of sapovirus sequences from outbreaks of acute gastroenteritis reported to state public health departments in Oregon and Minnesota, 2002–2009, on the basis of partial capsid nucleotide sequences. Reference strains [GenBank accession numbers] include Sapporo/1982/JP [U65427], Parkville/1994/US [U73124], Stockholm318/1997/SE [AF194152], Chiba000496/2000/JP [AJ606693], Ehime2K-814/2000/JP [AJ606698], London/1992/UK [U95645], Mex340/1990/MX [AF435812], cruise ship/2000/US [AY289804], PEC-Cowden/1980/US [AF182760], Hou7-1181/1990/US [AF435814], and Argentina39/AR [AY289803]. Boldface indicates state-assigned outbreak identification numbers. Scale bar represents percent genetic similarity between sequence types. Genogroups are indicated on the right. For genogrouping, GenBank sequences of well-characterized genogroups were aligned with outbreak sequences, and a phylogenetic tree was created by the neighbor-joining method by using BioNumerics (Applied Maths, Austin, TX, USA). Genotypes were assigned on the basis of >95% similarity to reference strains. Outbreak strain sequences were deposited in GenBank under accession nos. HM800902–HM800920.]

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with testing of fecal specimens. Public health laboratories should archive fecal samples from all gastroenteritis outbreaks until a cause is established. As in this study, testing with assays for sapovirus, astrovirus, adenovirus, and rotavirus, should be conducted when standard methods for norovirus and enteric bacterial pathogens fail to identify a causative agent.

In keeping with recent recommendations, at minimum, adding sapovirus to routine diagnostics of infections that occur in any setting and by any mode of transmission will establish etiologies of some norovirus-negative outbreaks and help define the disease impact and clinical characteristics of sapovirus infections (9,10,13). These data can in turn be used to develop and evaluate sapovirus disease management guidelines and sapovirus outbreak prevention and control measures.

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