Paediatric morbidity and mortality in Sierra Leone. Have things changed after the 2014/2015 Ebola outbreak? [version 1; peer review: 2 approved with reservations]

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Abstract

Background: Sierra Leone was severely affected by the 2014/2015 Ebola outbreak and is likely to have had longer term repercussions on the health system including on paediatric morbidity and mortality. We thus assessed under-five morbidity and mortality for malaria, acute respiratory Infections (ARI)/pneumonia, watery diarrhoea and measles during the post-Ebola period in Sierra Leone and compared this with the pre- and intra-Ebola periods.

Methods: This was a retrospective cross-sectional study using program data from the District Health Information system (DHIS2) and sourced from 14 districts in Sierra Leone. It included under-five children from 1,200 health facilities country-wide. Study periods included: before (June 1st, 2013-April 30th, 2014); during (June 1st, 2014-April 30th, 2015); and after Ebola (June 1st, 2016-April 30th, 2017).

Results: Malaria, ARI/pneumonia and diarrhoea consultations declined during Ebola but recovered to pre-Ebola levels in the post-Ebola period. During the post-Ebola period, there was a highly significant reduction in case-fatality for the first three morbidities compared to the pre-Ebola period (P<0.0001). Average number of measles cases increased from 48/month in the pre-Ebola period to 568/month (12-fold increase) post-Ebola. Although there was no difference in measles case-fatality between the pre- and post-Ebola periods, case-fatality post-Ebola was significantly lower than during Ebola (Relative Risk: 0.05, 95% confidence interval 0.02-0.15, P<0.0001).

Conclusions: Consultations for under-five children at health facilities in Sierra Leone recovered to pre-Ebola levels and case-fatality for common childhood illnesses declined significantly. This is a change for the better. However, the high level of reported measles cases in the post-Ebola period indicates gaps in immune status and needs focused attention.
Keywords
Sustainable Development Goals, Outbreak response, Universal Health Coverage, Operational Research, SORT IT

This article is included in the Disease Outbreaks gateway.

This article is included in the TDR gateway.

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Author roles: Sesay T: Conceptualization, Data Curation, Formal Analysis, Investigation, Methodology, Project Administration, Resources, Software, Supervision, Validation, Visualization, Writing – Original Draft Preparation, Writing – Review & Editing; Denisiuk O: Data Curation, Formal Analysis, Methodology, Software, Validation, Visualization, Writing – Original Draft Preparation, Writing – Review & Editing; Zachariah R: Conceptualization, Formal Analysis, Funding Acquisition, Methodology, Resources, Software, Supervision, Validation, Writing – Original Draft Preparation, Writing – Review & Editing

Competing interests: No competing interests were disclosed.

Grant information: The programme was funded by the Special Programme for Research and Training in Tropical Diseases hosted at the World Health Organization (TDR). The funders had no role in study design, data collection and analysis, decision to publish, or preparation of the manuscript.

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Introduction

In 2017, a cross-sectional study documented country-wide morbidity for four common childhood illnesses: malaria, acute respiratory infections (ARI)/pneumonia, watery diarrhoea and measles. There were two main findings. First, during the Ebola outbreak, health facility visits for malaria, ARI and watery diarrhoea dropped significantly nation-wide, without returning to pre-Ebola levels post-outbreak. As these morbidities have similar symptom patterns as Ebola, people may have avoided accessing formal health services to avoid being considered “an Ebola case”. Second, measles cases increased dramatically by six-fold during Ebola and the immediate post-Ebola periods. This was attributed to cessation of measles vaccination activities during the Ebola outbreak.

The outbreak was declared over in November 2015. Since then, there have been considerable investments into the health system by Government and development partners. One of the limitations of the 2017 study was that it only included the immediate six-month period after the Ebola outbreak, which might have been too early to assess health system recovery. It is now expected that the country would have fully recovered from the outbreak, but there has been no formal evaluation in this regard.

We thus conducted a similar country-wide study assessing morbidity and mortality for the same childhood illnesses using a longer post-Ebola period and compared this data with the pre- and intra-Ebola periods.

Methods

This was a retrospective analysis using routine program data from the District Health Information system (DHIS2) and sourced from all 14 districts in Sierra Leone (see Underlying data). The study setting was described in detail before. In brief, the health infrastructure is tiered into tertiary hospitals, district hospitals and Peripheral Health Units (PHUs). PHUs include Community Health Centres (CHCs), Community Health Posts (CHPs) and Maternal and Child Health Posts (MCHPs). The Ministry of Health and Sanitation provides free primary health care for children under five across 1,250 health facilities nationwide.

The study population included all children under-five years attending public health facilities nationwide. No children were excluded.

Study periods included: before (June 1st 2013-April 30th 2014); during (June 1st 2014-April 30th 2015); and after Ebola (June 1st 2016-April 30th 2017).

We exported data on health facility visits and mortality for malaria, (ARI)/pneumonia, watery diarrhoea and measles from the DHIS2 to Microsoft excel (2016) for analysis. Differences between groups were assessed using Pearson’s \( \chi^2 \) test (Chi square). Levels of significance were set at \( P \leq 0.05 \).

Discussion

This study shows that health facility consultations for malaria, ARI/Pneumonia and watery diarrhoea recovered to pre-Ebola levels and were accompanied by significant country-wide reductions in case-fatality compared to the pre-Ebola period. Despite a dramatic increase in measles cases post-Ebola, there was a significant mortality reduction, suggesting overall improvements in clinical care.

A study strength is that we included data from 1250 health facilities and for similar periods before, during and after the outbreak. A limitation is that our data did not include private health facilities resulting in possible underestimation of disease burden.

There were two key findings. First, the reductions in case fatality from malaria, ARI and watery diarrhoea could be associated with improved post-Ebola health seeking behaviour and re-establishment of community confidence in the health system. The post-Ebola recovery plan of the Government of Sierra Leone with enhanced financial, technical and training
Table 1. Morbidity and case-fatality for four under five morbidities before, during and after the Ebola outbreak in Sierra Leone.

|                      | Pre-Ebola | Ebola | Post-Ebola | P-value
<table>
<thead>
<tr>
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<tbody>
<tr>
<td></td>
<td>n</td>
<td>N</td>
<td>N</td>
<td></td>
</tr>
<tr>
<td>Malaria</td>
<td></td>
<td></td>
<td></td>
<td></td>
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<tr>
<td>Cases</td>
<td>989068</td>
<td>724881</td>
<td>1056354</td>
<td></td>
</tr>
<tr>
<td>Deaths</td>
<td>2564</td>
<td>1205</td>
<td>2112</td>
<td>&lt;0.0001</td>
</tr>
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<td>Deaths/1000</td>
<td>2.6</td>
<td>1.7</td>
<td>2.0</td>
<td></td>
</tr>
<tr>
<td>ARI/Pneumonia</td>
<td></td>
<td></td>
<td></td>
<td></td>
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<tr>
<td>Cases</td>
<td>717345</td>
<td>521860</td>
<td>735836</td>
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<td>Deaths</td>
<td>849</td>
<td>794</td>
<td>568</td>
<td>&lt;0.0001</td>
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<tr>
<td>Deaths/1000</td>
<td>1.2</td>
<td>1.5</td>
<td>0.8</td>
<td></td>
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<tr>
<td>Watery Diarrhoea</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Cases</td>
<td>200006</td>
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<td>203520</td>
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<tr>
<td>Deaths</td>
<td>361</td>
<td>150</td>
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<tr>
<td>Deaths/1000</td>
<td>1.8</td>
<td>1.2</td>
<td>1.2</td>
<td>&lt;0.0001</td>
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<tr>
<td>Measles</td>
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<td></td>
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<td></td>
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<tr>
<td>Cases</td>
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<td>962</td>
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<tr>
<td>Deaths</td>
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<td></td>
</tr>
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<td>Deaths/1000</td>
<td>1.9</td>
<td>16.6</td>
<td>1.0</td>
<td>0.5</td>
</tr>
</tbody>
</table>

1 Pre-Ebola: June 1st 2013 – April 30th 2014; Ebola: June 1st 2014 – April 30th 2015; Post-Ebola: June 1st 2016 – April 30th 2017.

2 Chi-square comparing the pre-Ebola and post-Ebola periods.
support from partners may also have contributed. Furthermore, community health worker activities including early identification and referrals of ill children were promoted which in turn would contribute to reducing mortality.

Second, increased measles cases during and after Ebola could be attributed to vaccination service cessation during Ebola in line with the recommendation to avoid invasive procedures as a way of minimizing Ebola-related occupational risks. Many children would have missed their measles vaccination, resulting in a reduction in herd immunity as well as an accumulation of unvaccinated children. Measles coverage among children under two years in 2017 (post-Ebola) stood at 80% while pre-Ebola this was at a low 78.6%. This implies that measles vaccination coverage was already below the desired level prior to Ebola and remained below desired levels after Ebola. This calls for measles vaccination campaigns at a higher age cut-off (6 months–15 years) to increase vaccination coverage to at least 95%. Laboratory confirmation is also needed to ensure that reported cases are not being mis-diagnosed since measles diagnosis is largely clinical.

In conclusion, consultations of under-five children at health facilities in Sierra Leone recovered to pre-Ebola levels and case-fatality for common childhood illnesses declined significantly. This is a change for the better. However, the high level of reported measles cases in the post-Ebola period needs focused attention.

Data availability

Source data

The Sierra Leone Health Management Information Systems, the District Health Information System 2 (DHIS2), is accessible with a Ministry of Health and Sanitation (MOHS) login through https://sl.dhis2.org/. The Directorate of Policy, Planning, and Information (DPPI) can be contacted to arrange access through Dr. Francis Smart (drfsmart@gmail.com), Director, DPPI, MOHS.

Underlying data

Repository: Dataset 1. Sesay_Tom_SORTIT2_paed_data. https://doi.org/10.17605/OSF.IO/SYP7G

This project contains the following underlying data:
- Sesay_T_casefatality_data.xlsx (case fatality data)
- Sesay_T_morbidity_data.xlsx (morbidity data)
- Sesay_T_paed_datadictionary.xlsx (data dictionary)

Data are available under the terms of the Creative Commons Zero “No rights reserved” data waiver (CC0 1.0 Public domain dedication).

Grant information

The programme was funded by the Special Programme for Research and Training in Tropical Diseases hosted at the World Health Organization (TDR).

The funders had no role in study design, data collection and analysis, decision to publish, or preparation of the manuscript.

Acknowledgement

This research was conducted through the Structured Operational Research and Training Initiative (SORT IT), a global partnership coordinated by the Special Programme for Research and Training in Tropical Diseases at the World Health Organization (WHO/TDR) and implemented with partners. The training model is based on a course developed jointly by the International Union Against Tuberculosis and Lung Disease (The Union) and Médecins sans Frontières (MSF). The specific SORT IT programme which resulted in this publication was jointly developed and implemented by: WHO/TDR, the Sierra Leone Ministry of Health and Sanitation, WHO Sierra Leone and the Centre for Operational Research, The Union, Paris, France, Alliance for Public Health, Ukraine; Institute of Tropical Medicine, Antwerp, Belgium; and Sustainable Health Systems, Freetown, Sierra Leone.

References

6. Statistics Sierra Leone (SSL) and ICF International: Sierra Leone Demographic and Health Survey 2013. Freetown, Sierra Leone and Rockville, Maryland, USA: SSL and ICF International. 2014. Reference Source
Open Peer Review

Current Peer Review Status:  

Version 1

Reviewer Report 28 October 2019

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Justin Lessler

Department of Epidemiology, Johns Hopkins Bloomberg School of Public Health, Baltimore, MD, USA

"Paediatric morbidity and mortality in Sierra Leone. Have things changed after the 2014/2015 Ebola outbreak?" provides a straight forward analysis of changes in reports of, and mortality from, four diseases before, during and after the 2014/2015 Ebola outbreak based on data from DHIS2 in Sierra Leone. The report provides simple, but important, statistics on morbidity and mortality for malaria, ARI/pneumonia, watery diarrhea, and measles. While the data and analysis is important, I have some significant concerns with the statistical analysis and interpretation of the results:

Statistics:
While the use of chi-squared test to identify that significant differences in mortality exist is appropriate, it only is being used to compare pre- and post Ebola periods and does not help with evaluation of the size of the changes. I would suggest the authors instead select a reference period and report relative rates and confidence intervals, so the data can be better understood. Further, the authors do not perform statistical tests to support some of their statements, some of which are noted below. More tests for differences in reported rates between periods and some analysis of changes in the Ebola period would better support their conclusions.

Interpretation:
The authors make several statements in the discussion and abstract that do not correspond to the analyses done. For instance, they make statements regarding trends in case reports, but perform no statistical test as to whether between period changes in reports are significant. Further, in some cases they frame speculation as a key finding of the paper, such as when they attribute changes to community confidence in the health system. Such speculations need to be clearly framed as such.

Overall, this paper could be made far stronger and more valuable with the addition of just a few additional basic statistical analyses that were more aligned with the conclusions the authors which to highlight in the discussion. This analysis could likely be done while preserving the admirable
conciseness and brevity of the paper.

Specific notes:
- Introduction, "to early to assess...": Could this not just be recovery, but also improvements, as significant investments followed the Ebola outbreak.
- Discussion, "First, the reductions in case fatality...": There is not really any evidence for this association in the paper, it should be, at the very least, more clear this is speculation.
- Discussion, "Second, increased measles cases...": If this is going to be said it needs to be better supported by the statistical analysis in the paper.
- Discussion, "In conclusion, consultations...": This was not explicitly tested in the analysis, so I am not sure it can be concluded.

Is the work clearly and accurately presented and does it cite the current literature?
Partly

Is the study design appropriate and is the work technically sound?
Partly

Are sufficient details of methods and analysis provided to allow replication by others?
Yes

If applicable, is the statistical analysis and its interpretation appropriate?
Partly

Are all the source data underlying the results available to ensure full reproducibility?
Yes

Are the conclusions drawn adequately supported by the results?
No

Competing Interests: No competing interests were disclosed.

Reviewer Expertise: Epidemiology, statistics, infectious disease dynamics

I confirm that I have read this submission and believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard, however I have significant reservations, as outlined above.

Author Response 09 Dec 2019
Tom Sesay, Ministry of Health and sanitation, Sierra Leone

Author Response to Reviewer Report for Version 1 from Justin Lessler 28 Oct 2019
Thank you for your thoughtful review of the manuscript and your suggestions. We have responded as below.

1. Reviewer comment: “Paediatric morbidity and mortality in Sierra Leone. Have things changed after the 2014/2015 Ebola outbreak?” provides a straightforward analysis of changes in reports of, and mortality from, four diseases before, during and after the 2014/2015 Ebola outbreak based on data from DHIS2 in Sierra Leone. The report provides simple, but important, statistics on morbidity and mortality for malaria, ARI/pneumonia, watery diarrhea, and measles.

Response: Thank you for this comment.

2. Reviewer comment: While the data and analysis is important, I have some significant concerns with the statistical analysis and interpretation of the results. Statistics: While the use of chi-squared test to identify that significant differences in mortality exist is appropriate, it only is being used to compare pre- and post Ebola periods and does not help with evaluation of the size of the changes. I would suggest the authors instead select a reference period and report relative rates and confidence intervals, so the data can be better understood. Further, the authors do not perform statistical tests to support some of their statements, some of which are noted below. More tests for differences in reported rates between periods and some analysis of changes in the Ebola period would better support their conclusions.

Response: Thank you for this point. We have performed t-tests for the differences in the mean monthly cases of the intra-Ebola and post-Ebola periods relative to the pre-Ebola period at a level of significance of 5%. We have also updated the figures by disease condition, which better displays the differences between the periods.

3. Reviewer comment: Interpretation: The authors make several statements in the discussion and abstract that do not correspond to the analyses done. For instance, they make statements regarding trends in case reports, but perform no statistical test as to whether between period changes in reports are significant. Further, in some cases they frame speculation as a key finding of the paper, such as when they attribute changes to community confidence in the health system. Such speculations need to be clearly framed as such.

Response: We have taken note of this comment and, accordingly, removed comments that could be considered as speculations.
4. Reviewer comment: Overall, this paper could be made far stronger and more valuable with the addition of just a few additional basic statistical analyses that were more aligned with the conclusions the authors which to highlight in the discussion. This analysis could likely be done while preserving the admirable conciseness and brevity of the paper.

Response:
We have included the basic statistical tests to test the statistical significance of differences between periods, as mentioned above.

5. Reviewer comment: Introduction, "to early to assess...": Could this not just be recovery, but also improvements, as significant investments followed the Ebola outbreak.

Response:
This comment is noted, and we have added in a phrase on possible improvements.

6. Reviewer comment: Discussion, "First, the reductions in case fatality..." There is not really any evidence for this association in the paper, it should be, at the very least, more clear this is speculation.

Response:
The reductions in case fatality were indicated by a Chi-square test of the difference between the pre-Ebola and the post-Ebola period which showed a significant reduction in case fatality in the post-Ebola period.

7. Reviewer comment: Discussion, "Second, increased measles cases...": If this is going to be said it needs to be better supported by the statistical analysis in the paper.

Response:
We have included a t-test of the average monthly measles cases, which showed significant increase in the number of measles cases between the pre-Ebola and post-Ebola periods.

8. Reviewer comment: Discussion, "In conclusion, consultations...": This was not explicitly tested in the analysis, so I am not sure it can be concluded.

Response:
We have ensured that the relevant statistical tests have been completed, with corresponding results to support the conclusions.
The authors of this paper have looked at trends in the pediatric morbidity and mortality prior to, during and after the 2014-15 Ebola outbreak in West Africa. The impact of large scale outbreaks on health systems remains a critical area of research, and the authors should be congratulated in contributing to this area of work. The comments below are intended to better contextualize the analysis and conclusions proposed by the authors.

○ The authors divide the paper into three distinct time periods, with ‘post-Ebola’ defined as beginning in June 2016 and extending through April 2017. While the end of the Ebola outbreak in Sierra Leone was declared in November 2015, there continued to be additional Ebola clusters in the country and the region into the ‘post-Ebola’ study period. The policy of mandatory Ebola testing for all deaths also continued until June 2016.

○ During the ‘post-Ebola’ time period in the paper, there was an enormous influx of funding to improve health systems at both the facility and community levels that was organized under the Presidential Recovery Plan. This funding supported the national and sub-national health systems but was also supplemented by NGO and other non-governmental health services. By the end of 2017, there were substantial reductions in the funds available for health systems and service delivery.

○ The continued specter of Ebola transmission and the rapid scale-up in health resources likely had both negative and positive impacts on health systems utilization that are difficult to quantify. The authors have documented significantly lower case fatality rates for three diseases than the pre-Ebola time period which would support the conclusion that there was regained community trust in the health services. However, those services were not the same ones available prior to the Ebola outbreak and the services are difficult to maintain with reduced external resources.

○ In order to compare the pre and post-Ebola periods, I would recommend that the analysis be extended to the immediate post-Ebola period included in this paper as well as a longer-term post-Ebola period through 2018 which would better capture the more sustainable health system in Sierra Leone rather than the one served by a short-term, high-volume influx of external funding.
Aside from the clear spikes in measles consultations, Figure 1 is difficult to interpret when displayed by time period. The differences may be more impactful if compared by disease instead.

The analysis supports the conclusion that measles surged during and post-Ebola and this is backed up by other research and program documentation. However, it does not necessarily support the recommendation to extend the cut off age for vaccination. The recommendation to conduct laboratory confirmation for measles is also not necessarily supported by the data presented. Is this because a number of the measles outbreaks were determined to be rubella which is currently not on the vaccine schedule? If yes, this should be documented by the data. What would be the impact on measles morbidity if there was increased diagnostic versus syndromic management of measles?

Is the work clearly and accurately presented and does it cite the current literature?
Partly

Is the study design appropriate and is the work technically sound?
Partly

Are sufficient details of methods and analysis provided to allow replication by others?
Yes

If applicable, is the statistical analysis and its interpretation appropriate?
Yes

Are all the source data underlying the results available to ensure full reproducibility?
Yes

Are the conclusions drawn adequately supported by the results?
Partly

**Competing Interests:** No competing interests were disclosed.

**Reviewer Expertise:** Infectious disease epidemiology

I confirm that I have read this submission and believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard, however I have significant reservations, as outlined above.

Author Response 09 Dec 2019

Tom Sesay, Ministry of Health and sanitation, Sierra Leone

Author Response to Reviewer Report for Version 1 from Sara Hersey 08 Oct 2019

Thank you for your thoughtful review of the manuscript and your insightful suggestions. We
have responded as below.

1. Reviewer comment: The authors of this paper have looked at trends in the pediatric morbidity and mortality prior to, during and after the 2014-15 Ebola outbreak in West Africa. The impact of large scale outbreaks on health systems remains a critical area of research, and the authors should be congratulated in contributing to this area of work. The comments below are intended to better contextualize the analysis and conclusions proposed by the authors.

Response:
Thank you for recognising the importance of this study and its contribution to knowledge about the impact of the 2014/15 Ebola outbreak.

2. Reviewer comment: The authors divide the paper into three distinct time periods, with ‘post-Ebola’ defined as beginning in June 2016 and extending through April 2017. While the end of the Ebola outbreak in Sierra Leone was declared in November 2015, there continued to be additional Ebola clusters in the country and the region into the ‘post-Ebola’ study period. The policy of mandatory Ebola testing for all deaths also continued until June 2016.

During the ‘post-Ebola’ time period in the paper, there was an enormous influx of funding to improve health systems at both the facility and community levels that was organized under the Presidential Recovery Plan. This funding supported the national and sub-national health systems but was also supplemented by NGO and other non-governmental health services. By the end of 2017, there were substantial reductions in the funds available for health systems and service delivery.

The continued specter of Ebola transmission and the rapid scale-up in health resources likely had both negative and positive impacts on health systems utilization that are difficult to quantify. The authors have documented significantly lower case fatality rates for three diseases than the pre-Ebola time period which would support the conclusion that there was regained community trust in the health services. However, those services were not the same ones available prior to the Ebola outbreak and the services are difficult to maintain with reduced external resources.

In order to compare the pre and post-Ebola periods, I would recommend that the analysis be extended to the immediate post-Ebola period included in this paper as well as a longer-term post-Ebola period through 2018 which would better capture the more sustainable health system in Sierra Leone rather than the one served by a short-term, high-volume influx of external funding.

Response:
Thank you for this recommendation which we do understand and accept. We do acknowledge the need for further research into the sustainability of the early post-Ebola gains in a much longer post-Ebola period. However, the objective of this study was to assess whether the investments in the health system had an impact on service delivery in the post-
Ebola period, as defined in the manuscript. In addition, the ethical approval obtained for this study was only for the periods mentioned in the paper. This, however, is an area we will be interested to investigate in the near future. We have added a recommendation to this effect in the discussion section.

3. Reviewer comment: Aside from the clear spikes in measles consultations, Figure 1 is difficult to interpret when displayed by time period. The differences may be more impactful if compared by disease instead.

Response:
We agree with your recommendation, and accordingly have completed the analysis separately by the disease conditions. We have amended the Figure 1 based on this.

4. Reviewer comment: The analysis supports the conclusion that measles surged during and post-Ebola and this is backed up by other research and program documentation. However, it does not necessarily support the recommendation to extend the cut off age for vaccination. The recommendation to conduct laboratory confirmation for measles is also not necessarily supported by the data presented. Is this because a number of the measles outbreaks were determined to be rubella which is currently not on the vaccine schedule? If yes, this should be documented by the data. What would be the impact on measles morbidity if there was increased diagnostic versus syndromic management of measles?

Response:
We do agree with this recommendation. We have removed the recommendation to extend the cut off age for vaccination and added the recommendation to increase the coverage of the second measles dose at 15 months.

Competing Interests: No competing interests were disclosed.
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