# The early use of Antibiotics for At-risk children with InfluEnza in Primary Care (the ARCHIE programme)

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#### **Disclosure of interests**

**Full disclosure of interests:** Completed ICMJE forms for all authors, including all related interests, are available in the toolkit on the NIHR Journals Library report publication page at https://doi.org/10.3310/WDFR7331.

Primary conflicts of interest: Kay Wang reports grants from the National Institute for Health and Care Research (NIHR) Academic Clinical Lectureship during the conduct of the study. Rafael Perera reports grants from the British Heart Foundation (PG/17/49/33099), NIHR School of Primary Care Research (Wittenberg\_FR16), NIHR Oxford Biomedical Research Centre (IS-BRC-1215-20008) and NIHR Community Healthcare MedTech and In Vitro Diagnostics Co-operative (MIC-2016-018) during the conduct of the study. Rafael Perera was a member of the HTA commissioning committee (2008–12). Anthony Harnden reports that in his role as Deputy Chairman of the Joint Committee for Vaccination and Immunisation, member since 2006, deputy chair from 2015, he was involved in the decision to recommend the use of the live attenuated influenza vaccine in school-age children. Malcolm G Semple reports that during the planning stages of ARCHIE he held grants from NIHR (Bronchiolitis Endotracheal Surfactant Study, 15/21/01), Medical Research Council UK and Health Protection Research Unit in Emerging & Zoonotic Infections, University of Liverpool. He is currently an independent external and non-remunerated member of Pfizer's External Data Monitoring Committee for their mRNA vaccine programme(s), from August 2022 continues, and chair of the Infectious Disease Scientific Advisory Board for Integrum Scientific LLC (Greensboro, NC, USA), from November 2018 continues, of which he is also a minority owner. He was a non-remunerated independent member of the HMG UK Scientific Advisory Group for Emergencies (SAGE), COVID-19 Response, from March 2020 to March 2022, and the HMG UK New Emerging Respiratory Virus Threats Advisory Group (NERVTAG), from 2014 continues.

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## Plain language summary

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or most children flu or a flu-like illness is a mild and short illness but is considered to be a factor for secondary complications, including bacterial infections. Children who have an underlying health condition (e.g. diabetes or Down syndrome) are particularly vulnerable to developing secondary complications. Flu and flu-like illness in these children place a great burden on families and health-care systems.

Our research explored the literature on children, flu and flu-like illness. We spoke to general practitioners who treat children and families of children who have experienced a bout of flu/flu-like illness. We conducted a clinical trial to see if giving children an antibiotic [co-amoxiclav (Augmentin®, GlaxoSmithKline UK)] early on in their illness would help them feel better more quickly and prevent complications. We conducted a substudy to take follow-up throat swabs at three times over a year to check for any signs of antibiotic resistance from co-amoxiclav treatment. We explored the cost-effectiveness of treatment with co-amoxiclav and if it was possible to identify which children would benefit most from it.

Our findings from the literature were that premature birth was an unidentified risk factor for flu/flu-like illness complications. We found that general practitioners are influenced by a variety of factors when making clinical decisions about individual children but these do not include if the child may have flu. Parents reported not being prepared for how quickly their child got worse when they had the flu and were positive about the use of antibiotics. The clinical trial indicated that there was no evidence of difference between children given co-amoxiclav and those given a placebo, or that co-amoxiclav helped children who visited a hospital outpatient service more than children who visited their general practitioner. The swab substudy did not show any evidence that co-amoxiclav led to children developing long lasting resistance to antibiotics.

We did not find any evidence that early treatment with co-amoxiclav makes it less likely that a child will be seen again by a doctor or nurse owing to their flu/flu-like illness getting worse. Our findings suggested that treatment with co-amoxiclav might lead to lower overall health-care costs, but a bigger trial would be needed to confirm this as there was no clear evidence of difference. It was not possible to identify if any specific group of children would benefit more from co-amoxiclav treatment early on in their illness. We do not recommend that children with an underlying health condition be given an antibiotic when they have a flu-like illness.

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