Evaluation of Targeted versus Universal Newborn Screening for Congenital Cytomegalovirus (cCMV)



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Background

Congenital cytomegalovirus (cCMV) infection in infants is extremely common, impacting ~1 in 200 live births. Hearing loss resulting from cCMV infection is estimated to cause 15-25% of cases of sensorineural hearing loss in children. It is the only preventable risk factor for permanent hearing loss. Despite the high prevalence of cCMV infection, newborn screening for cCMV is not common. Two approaches for cCMV screening are: 1) targeted screening of infants who do not pass their newborn hearing screening (NBHS); and 2) universal screening of all infants at birth

The aim of this project is to evaluate the efficacy of a targeted versus universal cCMV screening protocol.

Methods

A review of literature was conducted along with a review of websites of state programs in the United States which implement cCMV screening. Articles were selected from credible search engines including Google Scholar, PubMed, American-Speech-Language-Hearing-Association journals, and Purdue Libraries. Key words used were: congenital cytomegalovirus (CMV) screening, permanent hearing loss, targeted screening, universal screening, early intervention

Review of Literature Targeted cCMV Screening Approach

Main findings References

- A targeted cCMV screening program is feasible.
 - Beswick et al., 2019
 - Diener et al., 2017
 - Kadambari et al., 2015
 - Martin et al., 2021
 Fouder et al., 2017
 - Fowler et al., 2017
- Targeted cCMV screening identifies cCMV-positive infants and is cost-effective.
- Beswick et al., 2019Kadambari et al., 2015
- Targeted cCMV screenings can be performed non-invasively within the time frame to begin antiviral treatment if appropriate.
 - Diener et al., 2017
 - Kadambari et al., 2015Kimberlin et al., 2003
- Targeted screening misses cCMV-positive infants who are asymptomatic but develop later onset hearing loss. A universal screening approach is

preferable.

- Martin et al., 2021Fowler et al., 2017
- Lanzieri et al., 2017

with targeted screening but identified with universal screening. Universal screening can be

- administered at the same time as NBHS. For cCMV-positive infants with SNHL at birth, the etiology of hearing loss could be identified.
- ~4-30% of cases of cCMV-related SNHL are progressive. Identifying all positive infants via universal screening would allow for close audiometric

monitoring.

References

Universal cCMV Screening Approach

Main findings ~85% of cCMV-positive infants are asymptomatic at birth and would be missed with targeted screening but identified with universal screening.

~33% of cCMV-positive

asymptomatic at birth

They would be missed

develop progressive SNHL.

children who are

- Boppana, Ross & Fowler, 2013
- Fowler et al., 2017
- Chen, Zhong & Gu, 2020
- Goderis et al., 2016
- Goderis et al., 2014Vos et al., 2021
- Goderis et al., 2014
- Goderis et al., 2016
- Lanzieri et al., 2017

• Chen, Zhong & Gu, 2020

- Martin et al, 2021
- Fowler et al., 2017
- Berrettini, Ravecca, Sellari-Franceschini, Matteucci, Siciliano & Ursino, 1999
- Martin et al., 2021

State Protocols

Aside from screening, some states have legislative mandates requiring education about cCMV to the public and/or healthcare professionals following these protocols:

- cCMV education plus cCMV screening
- Education about cCMV only
- Utah is the only state with targeted cCMV screening and mandated education accompanied by annual funding.

Results and Clinical Decision

The literature review suggests that universal cCMV screening has greater overall efficacy than targeted screening.

- 1. cCMV infections are asymptomatic in most cases, resulting in the exclusion of majority of cCMV-positive infants when using targeted screening.
- 2. Late onset SNHL occurs in 1 out of 3 asymptomatic cCMV-positive infants, who would be missed if using targeted screening.
- 3. Barriers for implementation of universal screening are state legislation and funding limitations.

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- •Contact: Grace Haines-Gallagher (317)3134884
- References available on request.

Comparisons of Targeted vs. Universal cCMV Screening Approach

Targeted

- Screening only warranted upon failure of initial and/or subsequent NBHS
- More cost-effective because only infants who fail NBHS are screened for cCMV
- Screening may fall outside 21 day window for determining if CMV is congenital or acquired

Both

- Part of newborn screening tests
- Easy collection and analysis of samples
- Identification of hearing loss etiology
- Close monitoring of cCMV positive infants

Universal

- Screening warranted for all infants regardless of hearing status
- Administered
 simultaneously with NBHS
- Allows determination of cCMV status in asymptomatic positive infants
- Enables vigilant audiometric monitoring for progressive SNHL in positive infants born without congenital HL
- Requires more resources for screening all infants

CCM V SCREENING IN THE UNITED STATES BY STATE

