

# Title: Infectious Agents Associated with Stillbirths and Early Neonatal (0-2 days) Deaths in Sub-Saharan Africa and South Asia: Findings from Child Health and Mortality Prevention Surveillance (CHAMPS), 2016-2019

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## Background

- In low and middle-income countries, infection is a major cause of stillbirths and neonatal deaths
- Significantly over time, the causative pathogens have changed but epidemiologic evidence is scanty

## Objective

To characterize the pathogens contributing to stillbirths and neonatal deaths occurring within the first two days of life in sub-Saharan Africa and South Asia

## Methods

- CHAMPS is a multi-country programme and the sites are –

- The objective of CHAMPS is to investigate the causes of stillbirths and children under 5 years old deaths using laboratory and clinical data including verbal autopsy
- Expert panels review the data and assign underlying, immediate and morbid causes of death (“causal chain”)
- CHAMPS uses below methods to detect pathogens –
  - Microbial Culture
  - Molecular Assay using TaqMan Array Card (TAC)

## Results

We examined 440 stillbirths and 433 neonatal deaths occurring in the first two days of life between December 2016 and December 2019 from all CHAMPS sites except India.

**Congenital infection was present in the causal chain of 12% stillbirths and 28% (0-2 days) deaths**

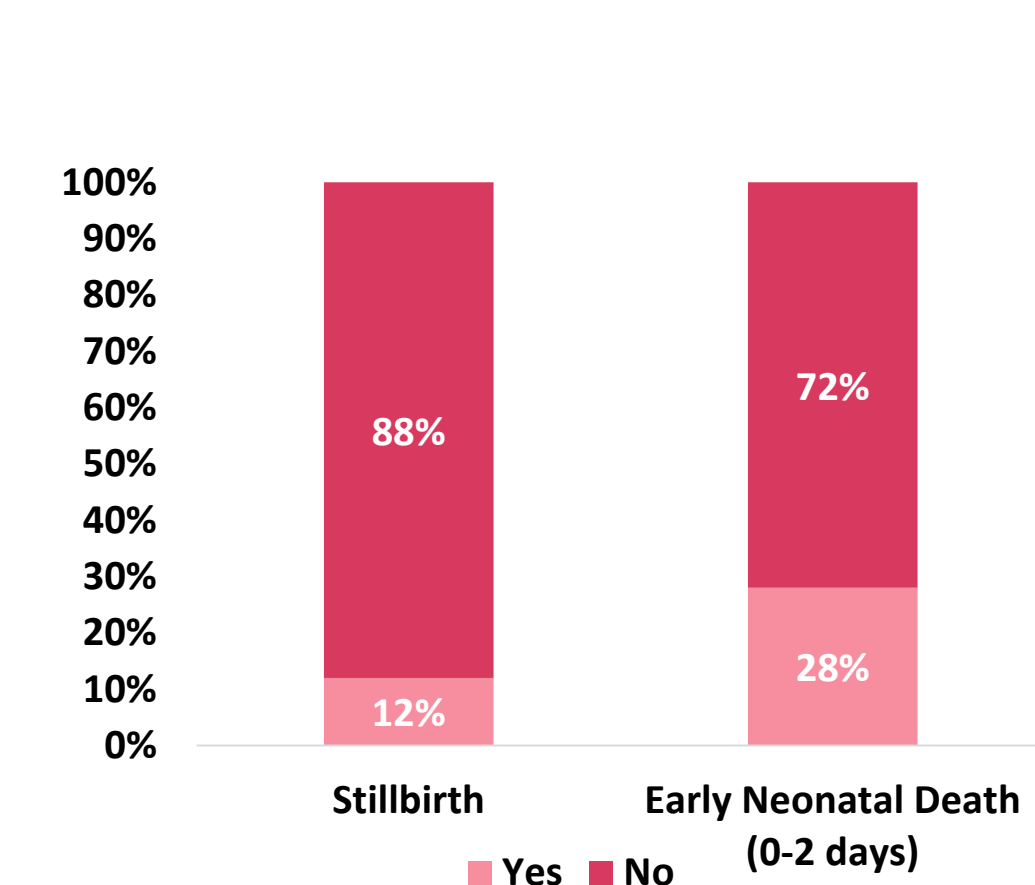


Figure 1: Infection in the causal chain of death

**Major pathogens identified among stillbirths and neonates (0-2 days) were gram-positive & gram-negative bacteria and viruses**

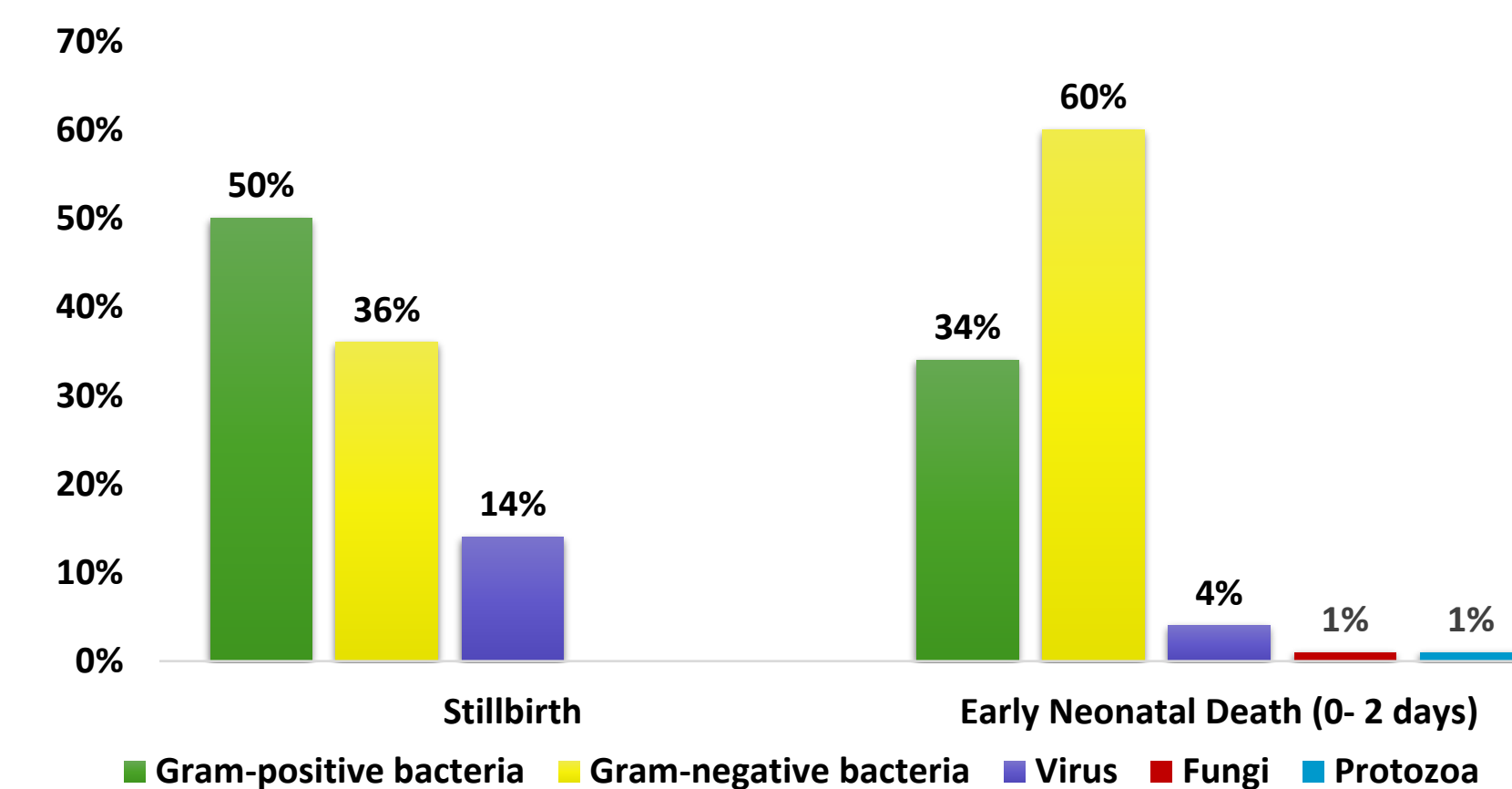


Figure 2: Type of pathogen identified

- Half of the pathogens identified (N=64) among the stillbirth were gram-positive bacteria (50%, 32/64), 36% (23/64) were gram-negative, and 14% (9/64) were viruses (Fig 2)
- We identified 140 pathogens in the causal chain for neonates; 60% (84/140) were gram-negative bacteria, 34% (47/140) gram-positive, and 4% (6/140) viruses (Fig 2)

Table 3: Etiologic agents identified in the causal chain in stillbirth, 2016-2019

Infectious agent type	Specific agents	Stillbirths
Gram-positive bacteria (N=32)	Streptococcus agalactiae	14
	Enterococcus faecalis	8
	Streptococcus spp.	6
	Streptococcus pneumoniae	3
	CoNS	1
Gram-negative bacteria (N=23)	Escherichia coli / Shigella spp.	9
	Klebsiella pneumoniae	3
	Treponema pallidum	3
	Enterobacter spp.	2
	Ureaplasma spp.	2
	Klebsiella oxytoca	1
	Haemophilus influenzae type B	1
	Pseudomonas aeruginosa	1
	Sneathia amnii	1
	Virus (N=9)	Cytomegalovirus
Varicella zoster virus		1

Table 4: Etiologic agents identified in the causal chain in early neonatal deaths (0-2 days), 2016-2019

Infectious agent type	Specific agents	Early Neonatal death (0-2 days)
Gram-negative bacteria (N=84)	Acinetobacter baumannii	17
	Acinetobacter spp.	1
	Chlamydia trachomatis	1
	Enterobacter cloacae	2
	Enterobacter spp.	1
	Escherichia coli	23
	Haemophilus influenza	2
	Klebsiella pneumoniae	31
	Morganella morganii	1
	Proteus mirabilis	1
	Pseudomonas spp.	1
	Salmonella spp.	1
	Ureaplasma spp.	2
	CoNS	1
Gram-positive bacteria (N=47)	Enterococcus faecalis	3
	Enterococcus faecium	1
	Enterococcus spp.	1
	Listeria monocytogenes	3
	Staphylococcus aureus	4
	Staphylococcus spp.	1
	Streptococcus agalactiae	22
	Streptococcus pneumoniae	4
	Streptococcus pyogenes	1
	Streptococcus spp.	6
Virus (N=6)	Cytomegalovirus	4
	HIV	2
Fungi (N=2)	Candida albicans	1
	Candida glabrata	1
Protozoa (N=1)	Toxoplasma gondii	1

## Conclusion

- The frequency of Gram-positive vs Gram-negative bacteria was divergent when comparing stillbirths and neonatal deaths
- The differences highlight the need for better characterization of cascades contributing to utero-fetal infection scenarios
- Understanding the source of infection through effective screening during pregnancy is crucial to prevent their associated morbidity and mortality



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