

A mechanistic model and web-based tool for estimating the potential impact of water, sanitation, and hygiene interventions, accounting for contextual and intervention factors

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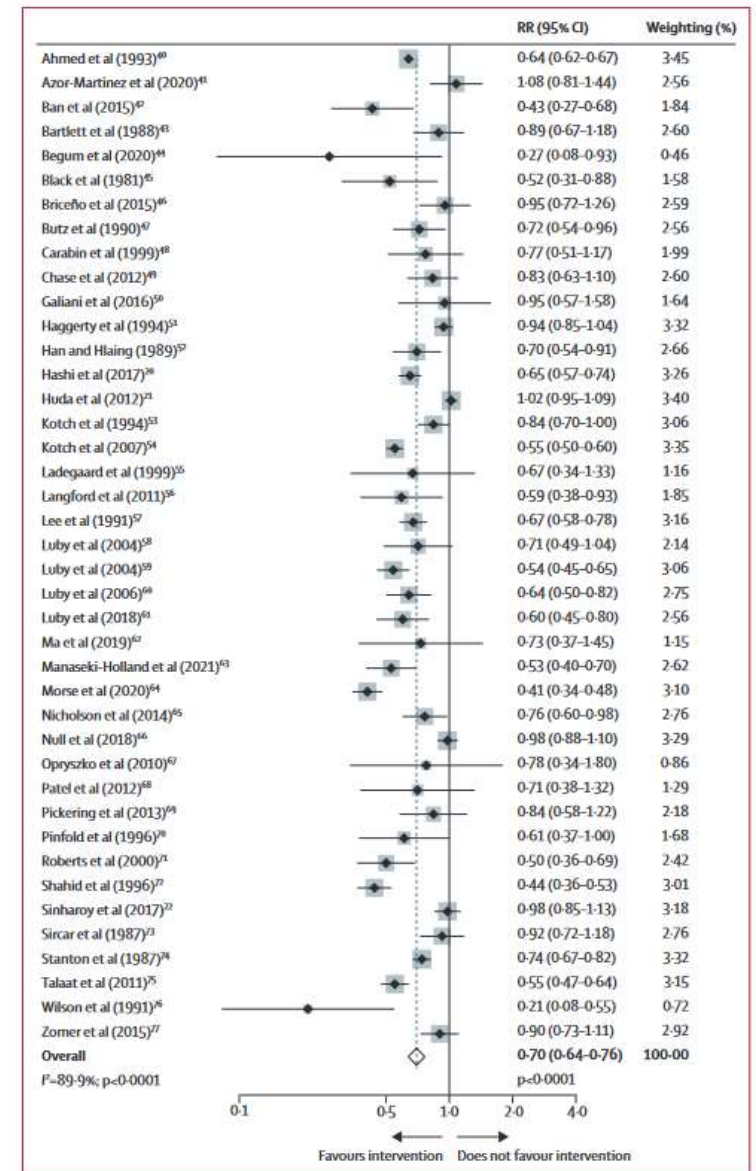
EMORY

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PUBLIC
HEALTH

BILL & MELINDA
GATES *foundation*

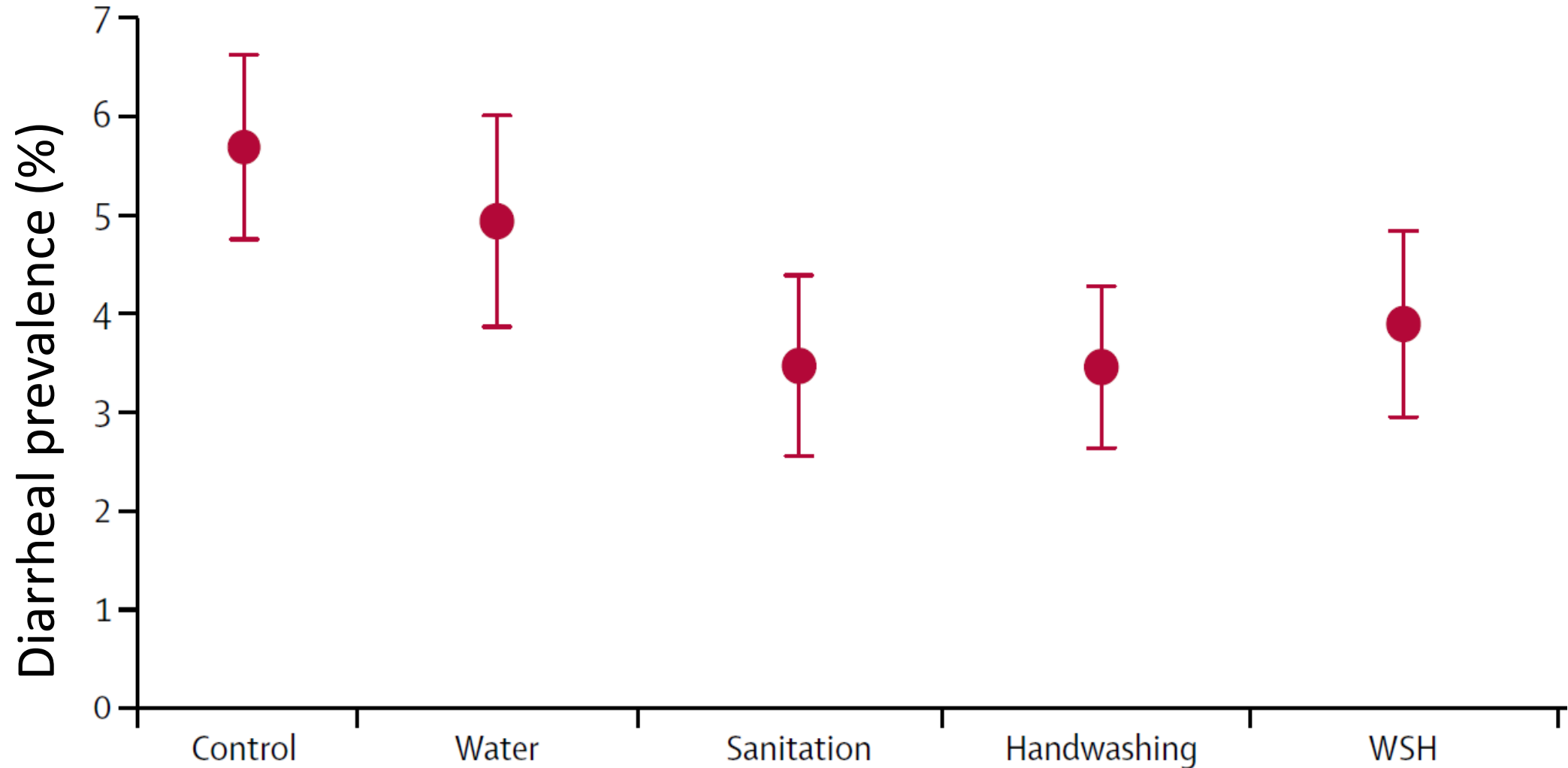
Water, sanitation, and hygiene (WASH) interventions have the potential to reduce the burden of diarrheal disease...

... but there has been substantial heterogeneity and multiple large trials with modest-to-null results.

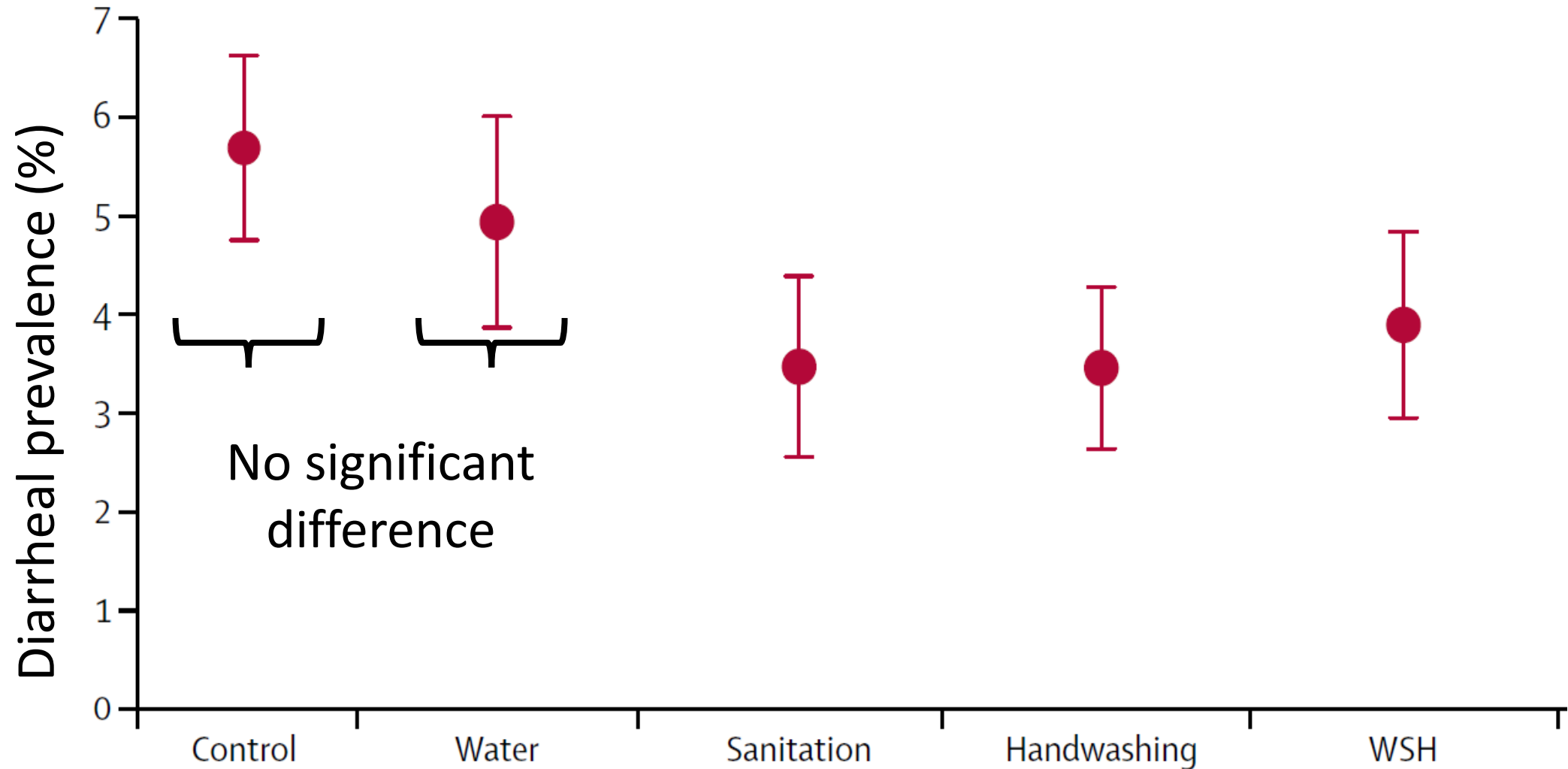


Wolf et al., 2022. *The Lancet*.
Meta-analysis of the effectiveness of hygiene interventions

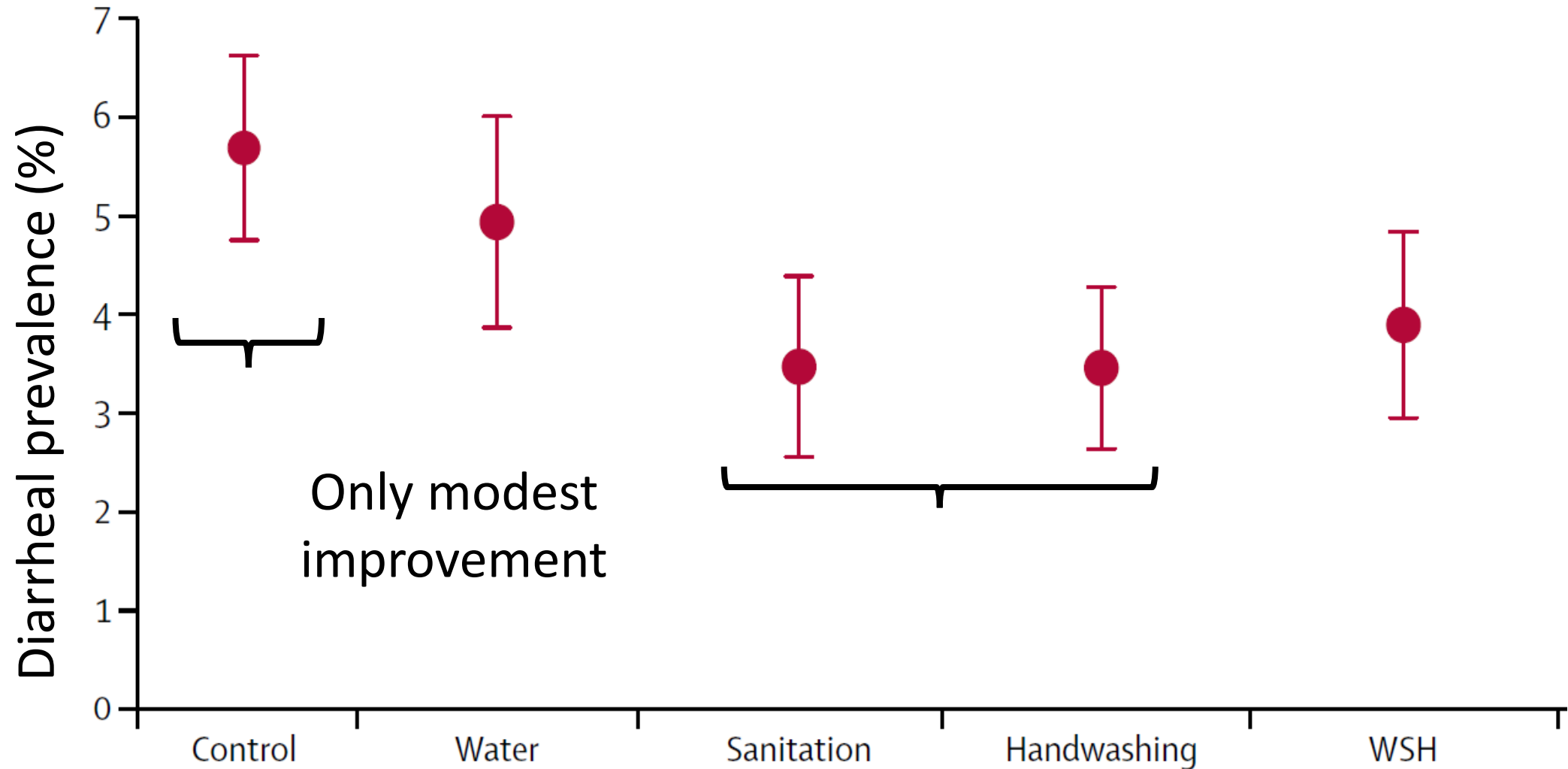
For example, the WASH Benefits Bangladesh trial found...



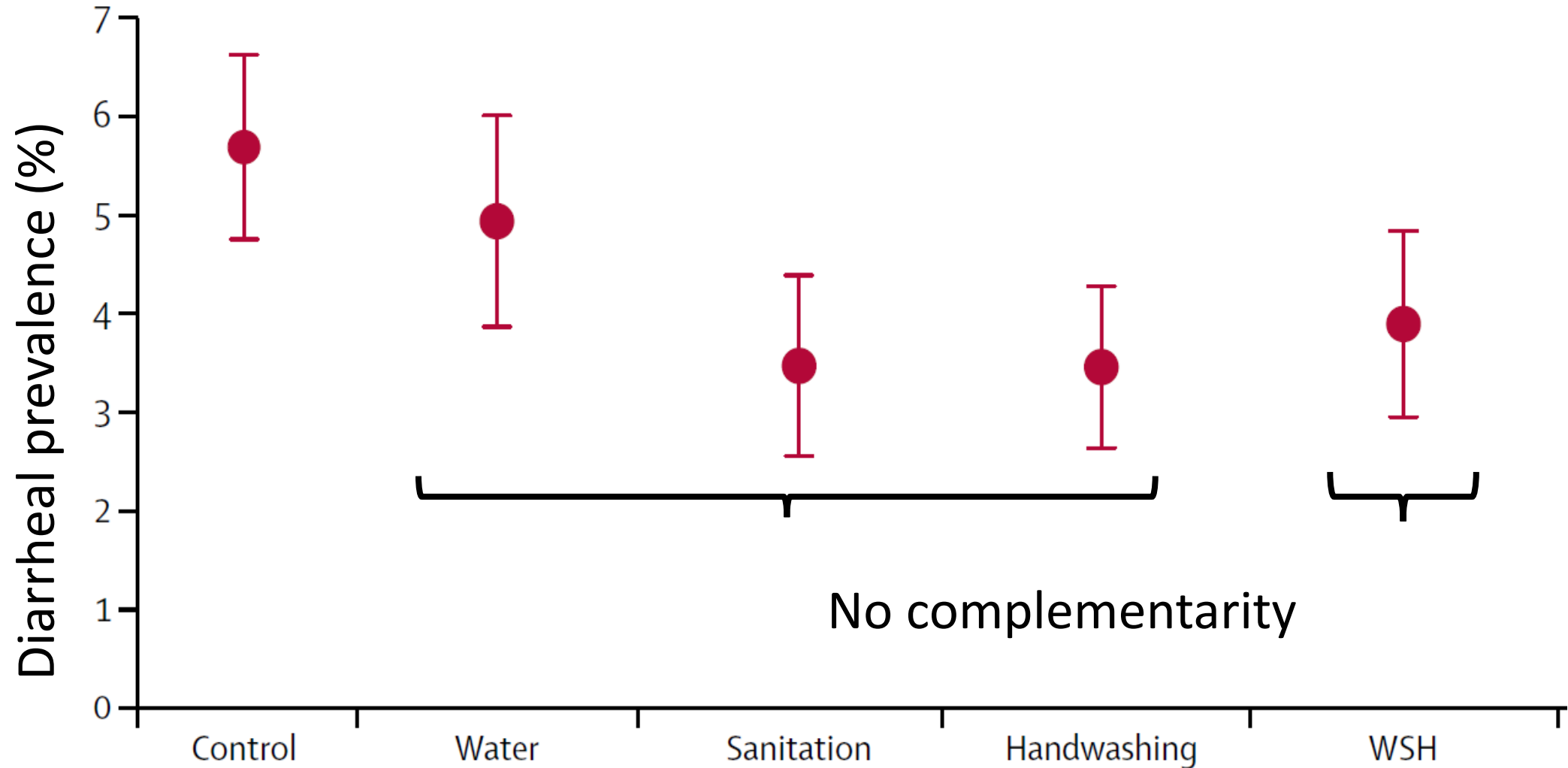
For example, the WASH Benefits Bangladesh trial found...



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Which factors contribute to heterogeneity in the effectiveness of interventions in WASH trials?

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Efficacy

Do the interventions work?

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Compliance

Are people receiving and using the interventions?

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Efficacy

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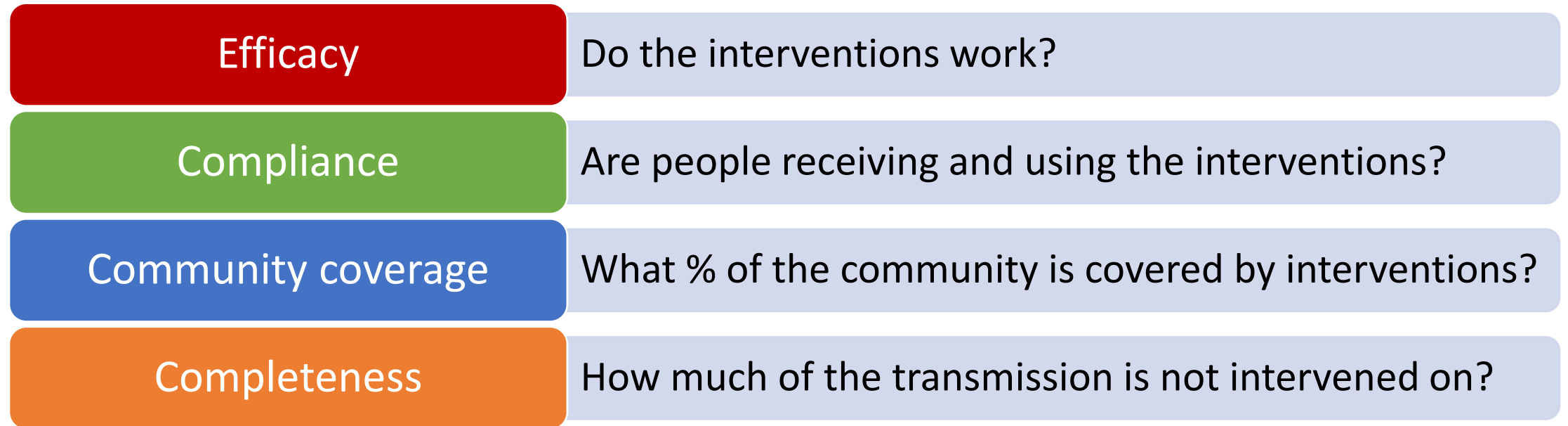
Compliance

Are people receiving and using the interventions?

Community coverage

What % of the community is covered by interventions?

Which factors contribute to heterogeneity in the effectiveness of interventions in WASH trials?



Magnitude of transmission pathways



Which factors contribute to heterogeneity in the effectiveness of interventions in WASH trials?

Efficacy

Do the interventions work?

Compliance

Are people receiving and using the interventions?

Community coverage

What % of the community is covered by interventions?

Completeness

How much of the transmission is not intervened on?

Baseline WASH conditions

Did participants already have WASH infrastructure?

Which factors contribute to heterogeneity in the effectiveness of interventions in WASH trials?

Efficacy

Do the interventions work?

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Are people receiving and using the interventions?

Community coverage

What % of the community is covered by interventions?

Completeness

How much of the transmission is not intervened on?

Baseline WASH conditions

Did participants already have WASH infrastructure?

Baseline disease conditions

Does the effectiveness depend on the disease burden?

Which factors contribute to heterogeneity in the effectiveness of interventions in WASH trials?

Efficacy

Compliance

Community coverage

Completeness

Baseline WASH conditions

Baseline disease conditions

Intervention factors are modifiable by the investigators

Contextual factors depend on the location

Many of these factors could be driven by differences in pathogen distributions

- Different pathogens use different transmission pathways to different degrees, and interventions have different efficacies

Many of these factors could be driven by differences in pathogen distributions

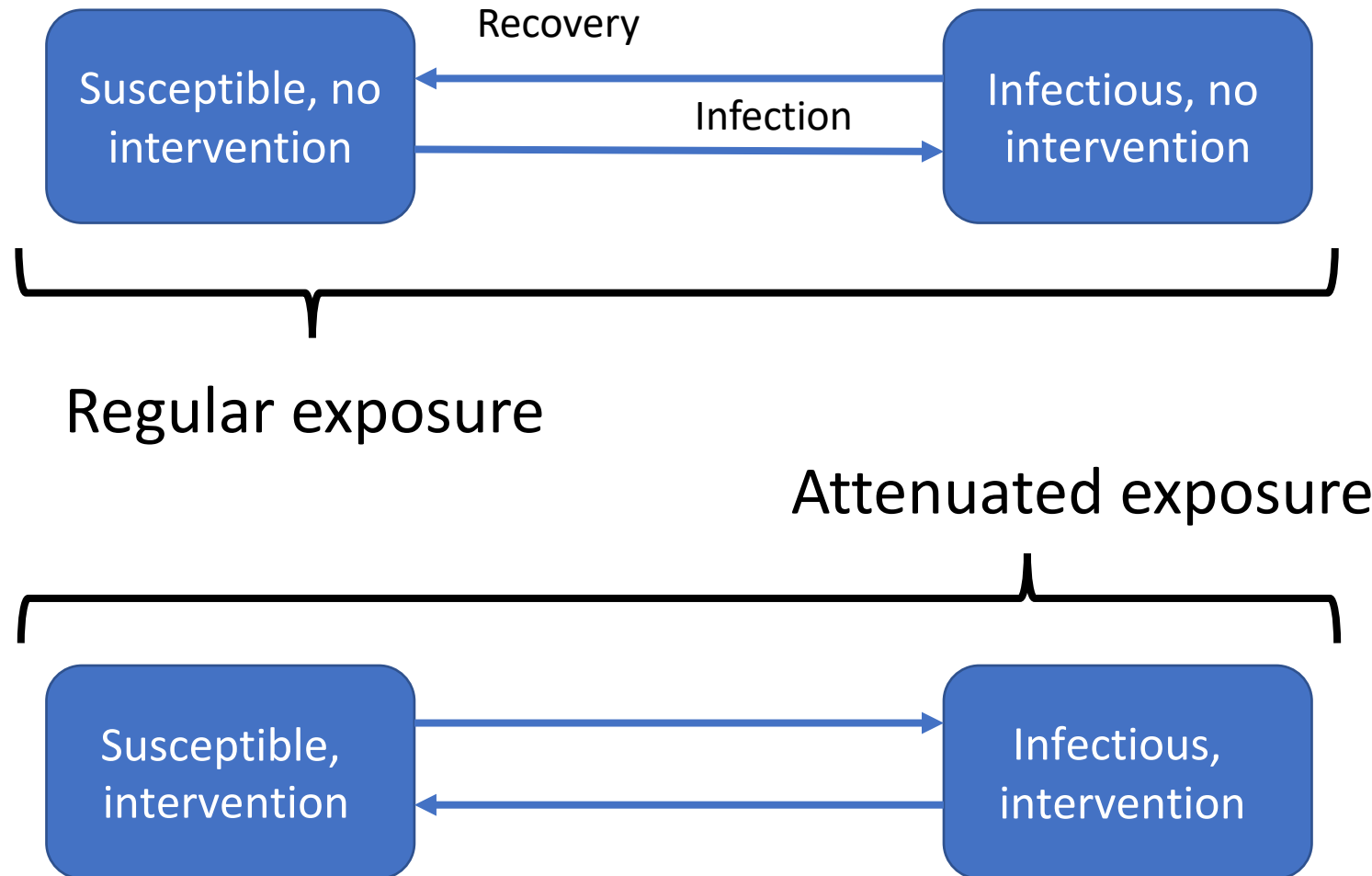
- Different pathogens use different transmission pathways to different degrees, and interventions have different efficacies

	Basse, The Gambia	Mirzapur, Bangladesh	Karachi (Bin Qasim Town), Pakistan
0-11 months			
Number of case-control pairs	258	293	284
Rotavirus	23.0% (18.8-29.1)	23.4% (18.9-28.7)	27.0% (22.7-33.5)
Adenovirus 40/41	9.6% (4.9-15.9)	8.3% (4.7-13.9)	7.2% (0.9-10.6)
<i>Cryptosporidium</i> spp	11.5% (4.6-16)	1.2% (0.1-2.7)	10.4% (5.1-16.1)
<i>Shigella</i> spp or EIEC	7.5% (3.9-12.9)	15.8% (12-20.4)	13.8% (9.7-19.7)
<i>C jejuni</i> or <i>C coli</i>	*	12.3% (7.6-19.8)	7.0% (0-17.1)
ST-EPEC	5.5% (1.8-9.7)	2.0% (0.5-3.9)	12.2% (8.9-17.6)
Norovirus GII	4.4% (1.8-11.1)	1.8% (0.2-5.3)	2.7% (0-6.2)
tEPEC	2.7% (0-5.7)	0.3% (0-2.6)	2.1% (0-5.1)
Sapovirus	1.0% (0-5.4)	1.1% (0-3.4)	4.4% (0.2-11.1)
Astrovirus	0.7% (0-4.4)	1.4% (0-3.9)	3.3% (0.1-7.6)
<i>V cholerae</i>	*	0.6% (0-1.7)	5.6% (2.3-8.8)

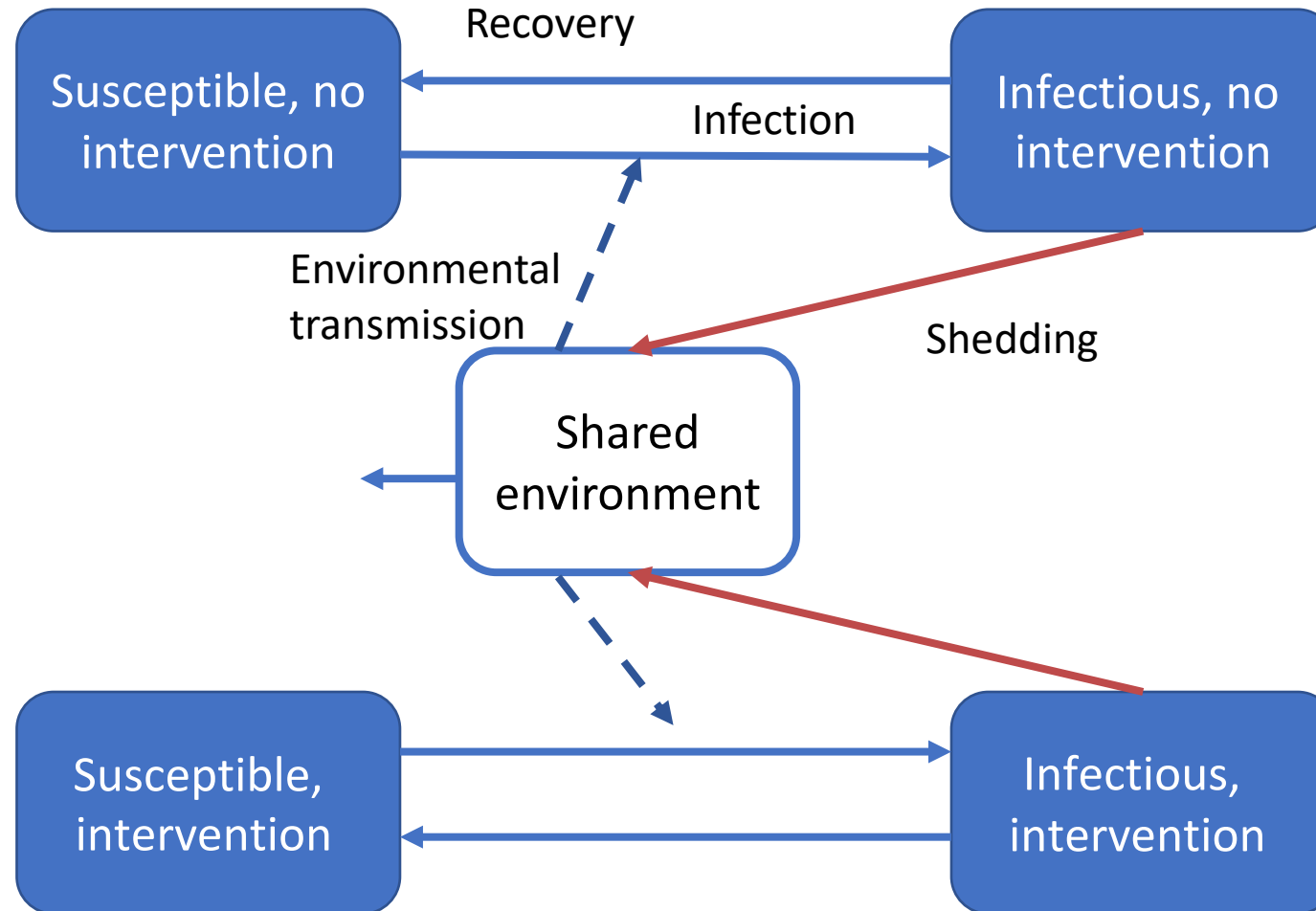
The Global Enteric Multicenter Study found substantial differences in the distribution of pathogens at different sites.

Mechanistic transmission modeling can aid in predicting and generalizing WASH trial outcomes.

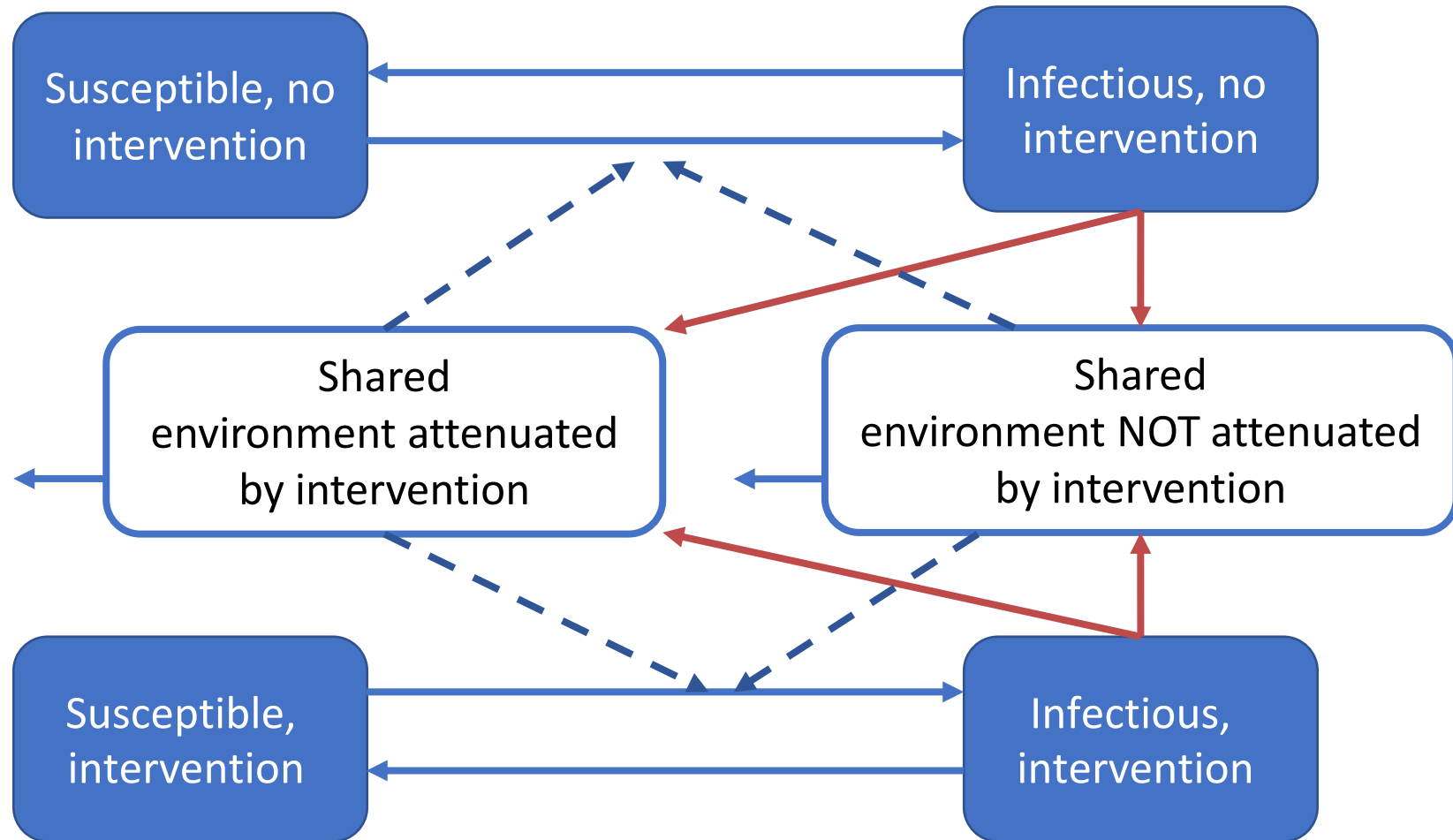
Our transmission model accounts for people using and not using interventions...



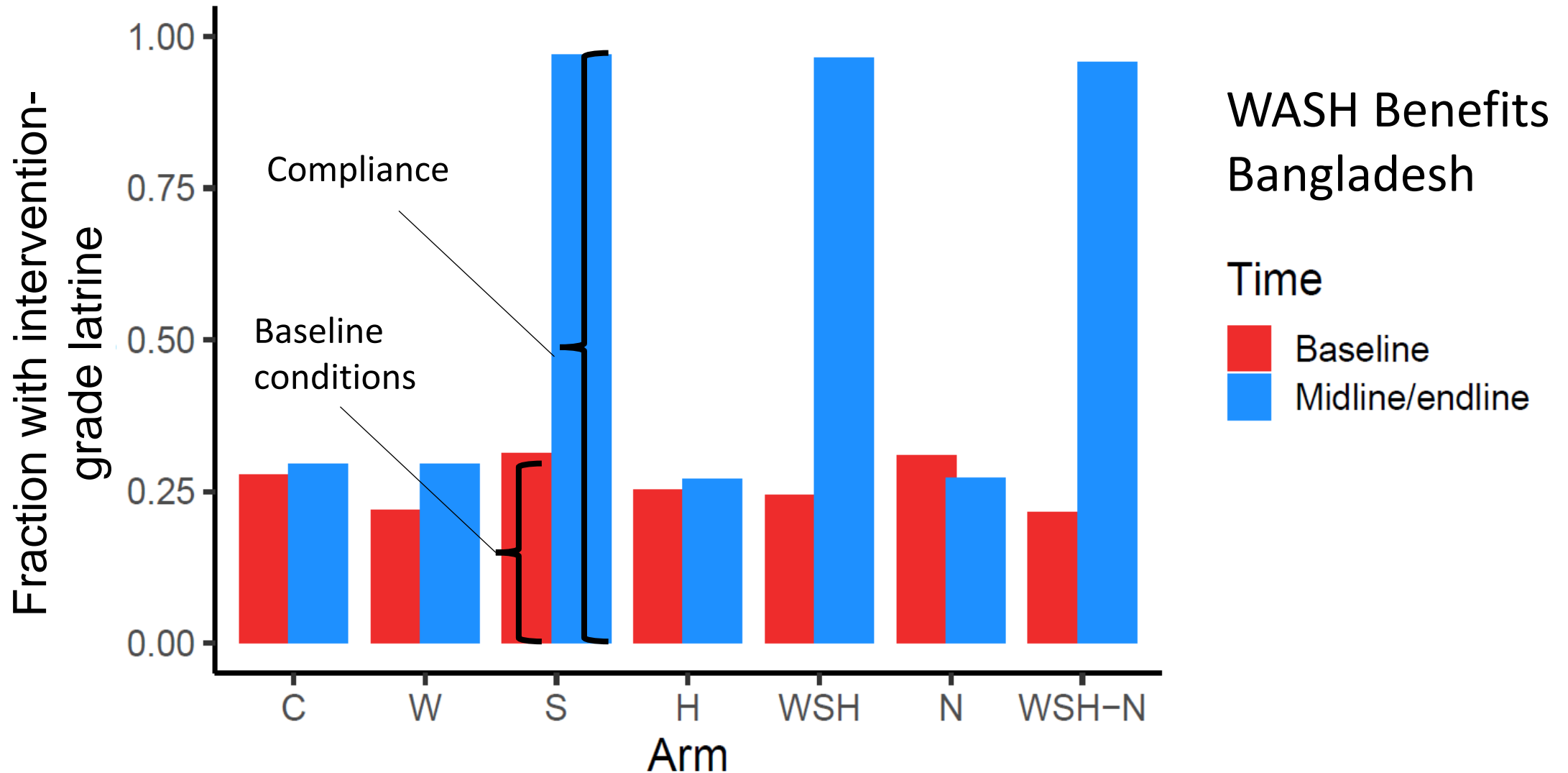
... a shared environment ...



... and multiple transmission pathways.



We also account for intervention compliance and baseline WASH conditions.

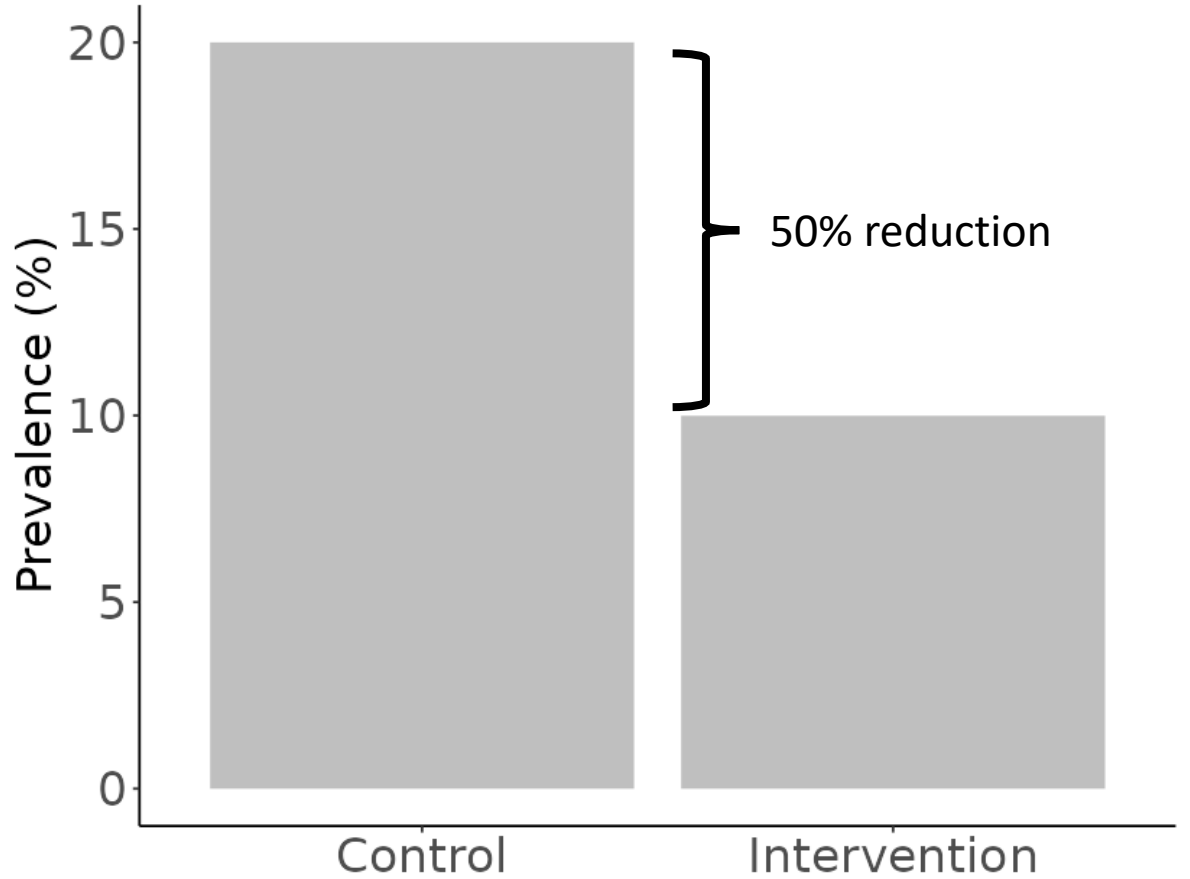
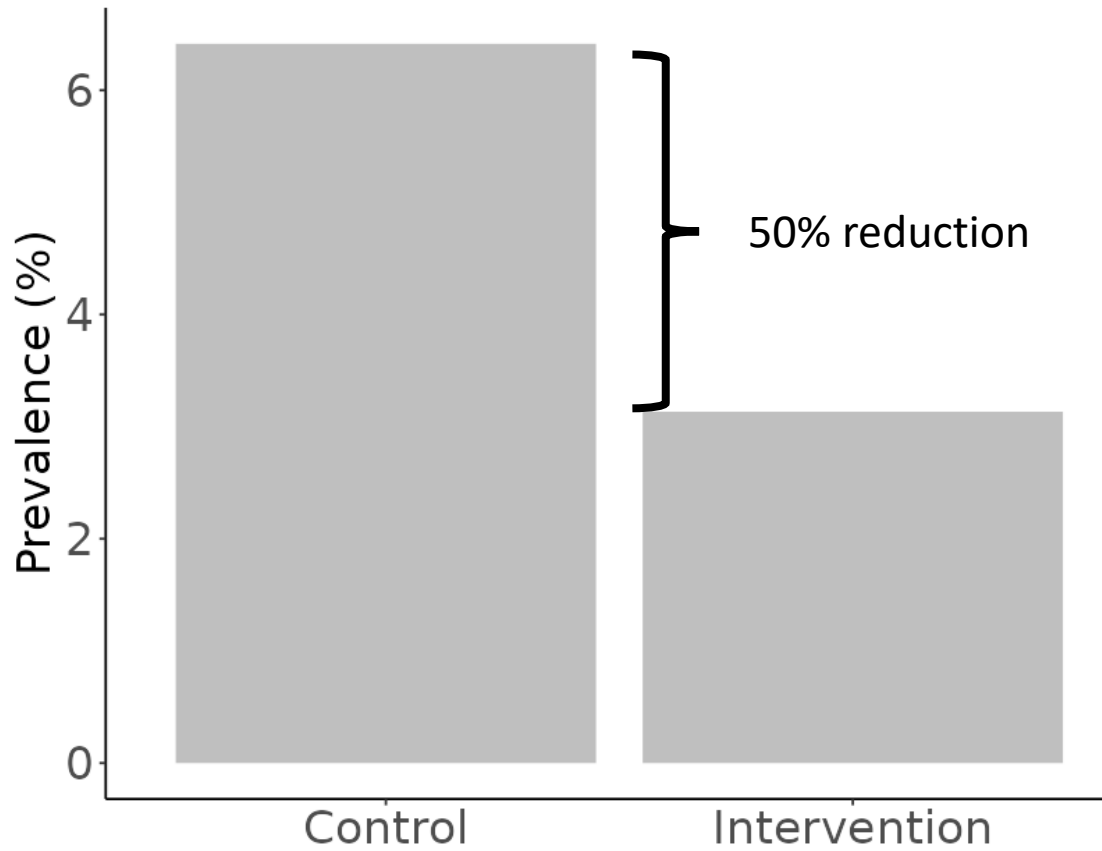


We illustrated the sensitivity of intervention effectiveness to intervention and contextual parameters in two scenarios.



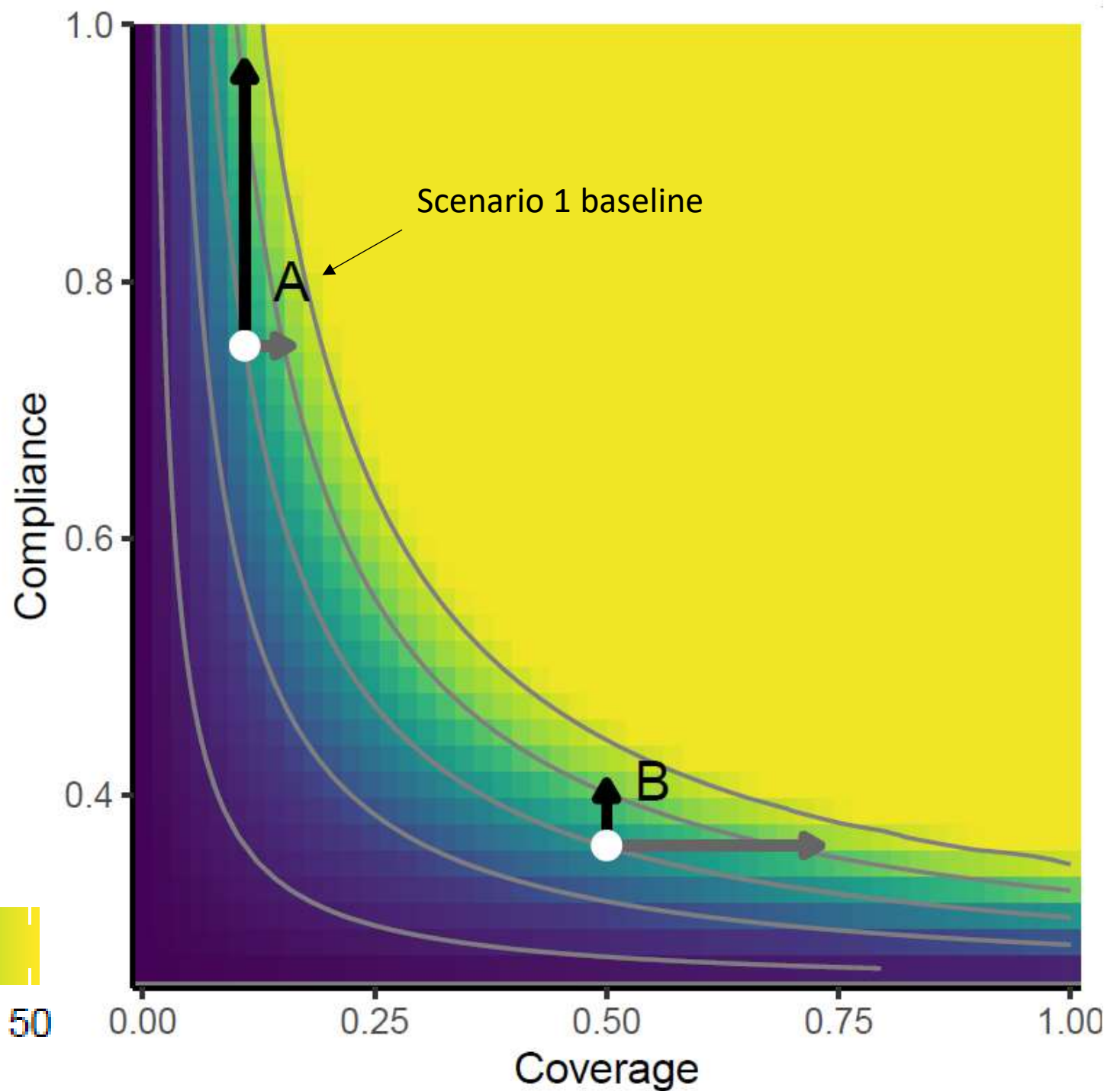
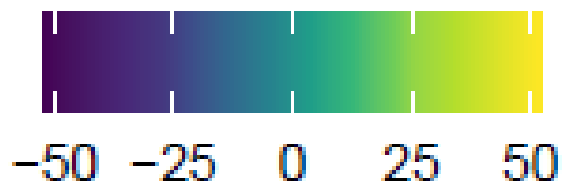
Scenario 1	Scenario 2
Lower (75%)	Higher (83%)
Higher (75%)	Lower (50%)
Lower (11%)	Higher (75%)
Higher (75%)	Lower (35%)
Higher (25%)	Lower (0%)
Lower (6.4%)	Higher (20.0%)

These scenarios were chosen to have 50% effectiveness.

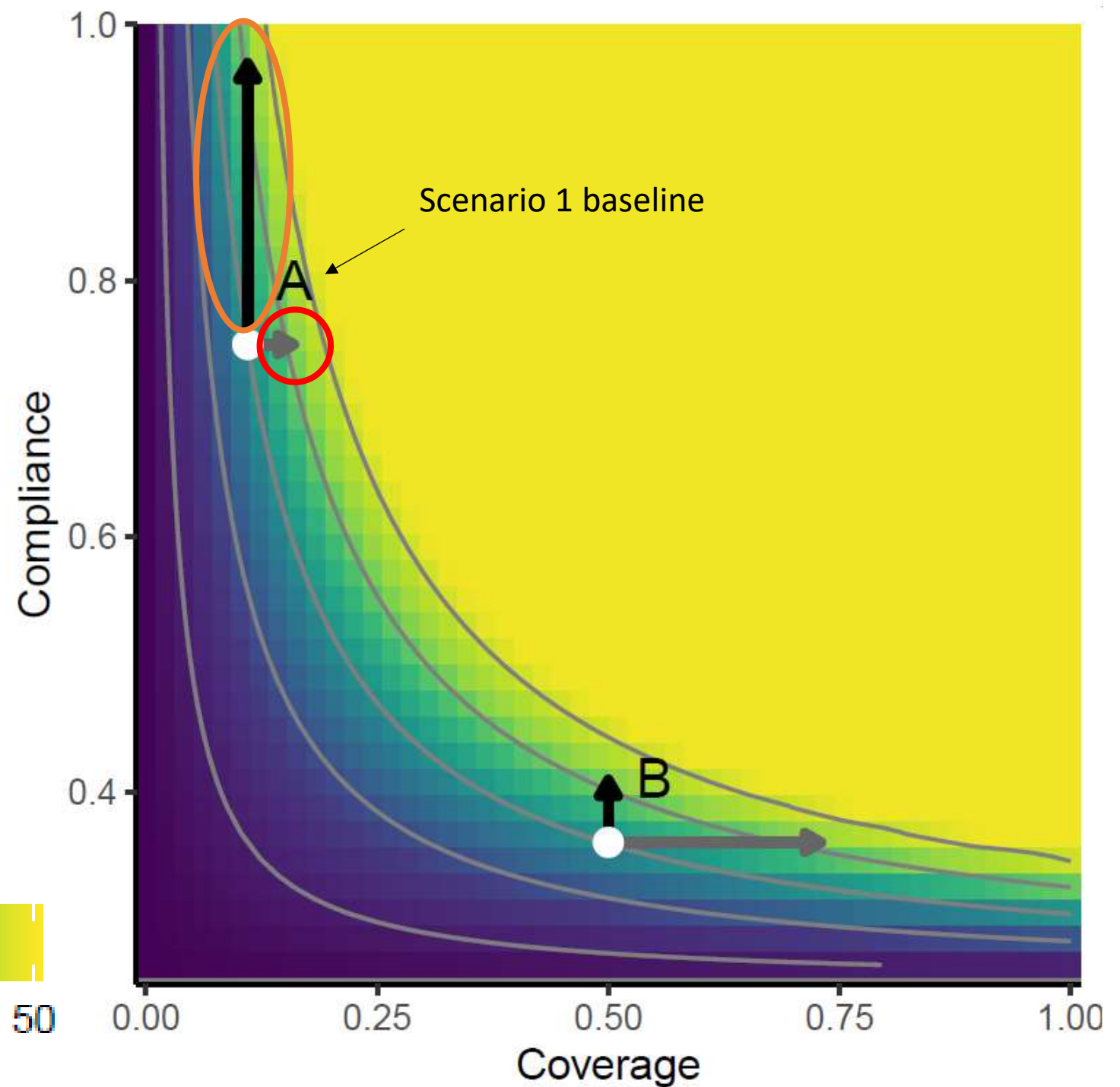
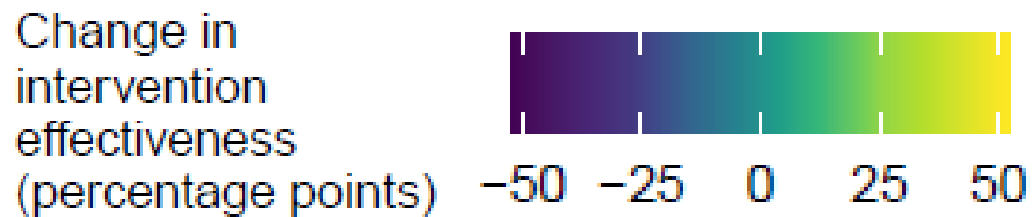


Sensitivity to one parameter depends on the other parameters

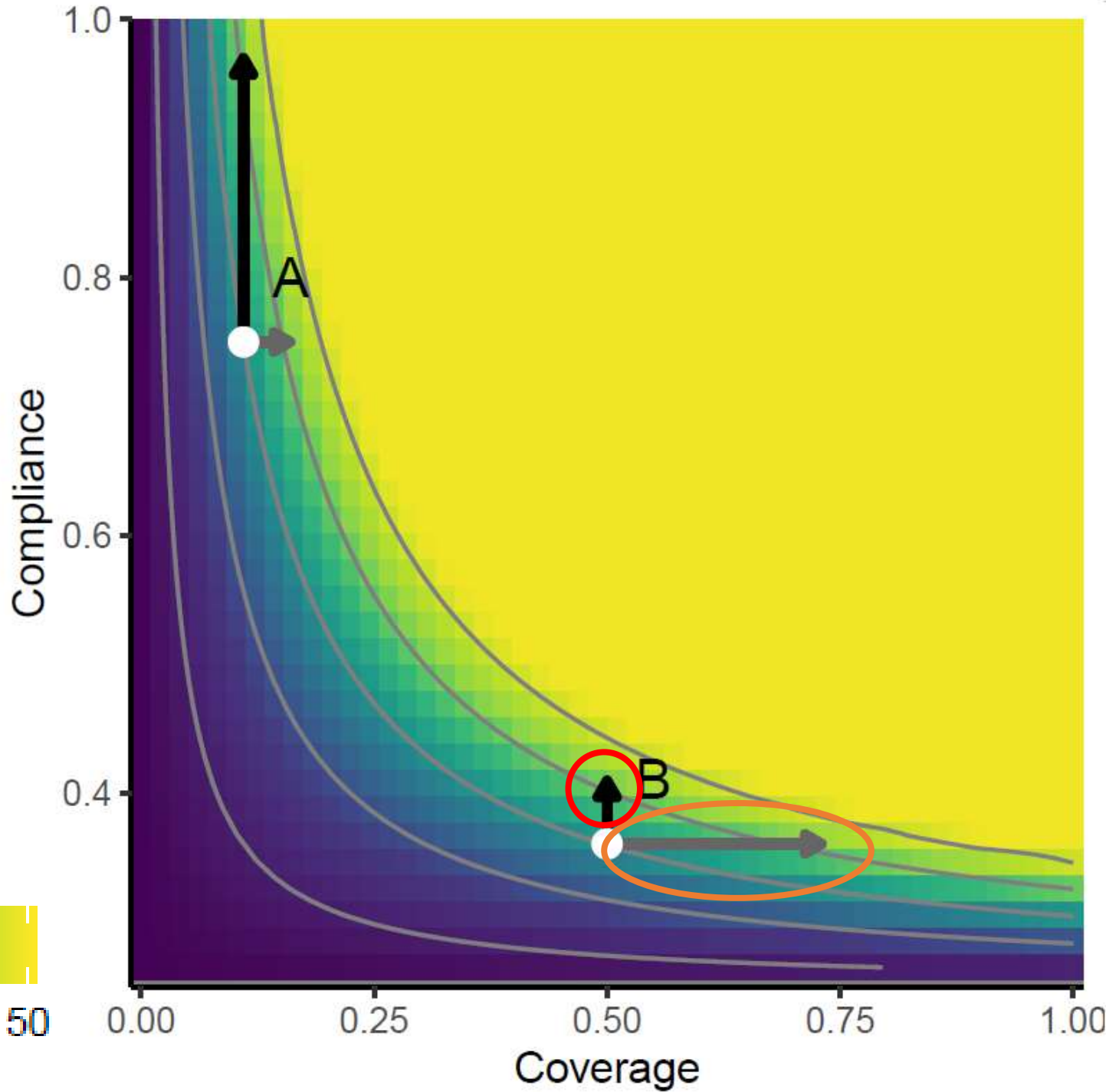
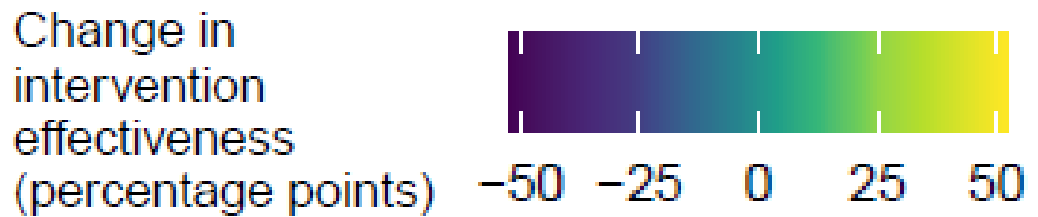
Change in intervention effectiveness (percentage points)



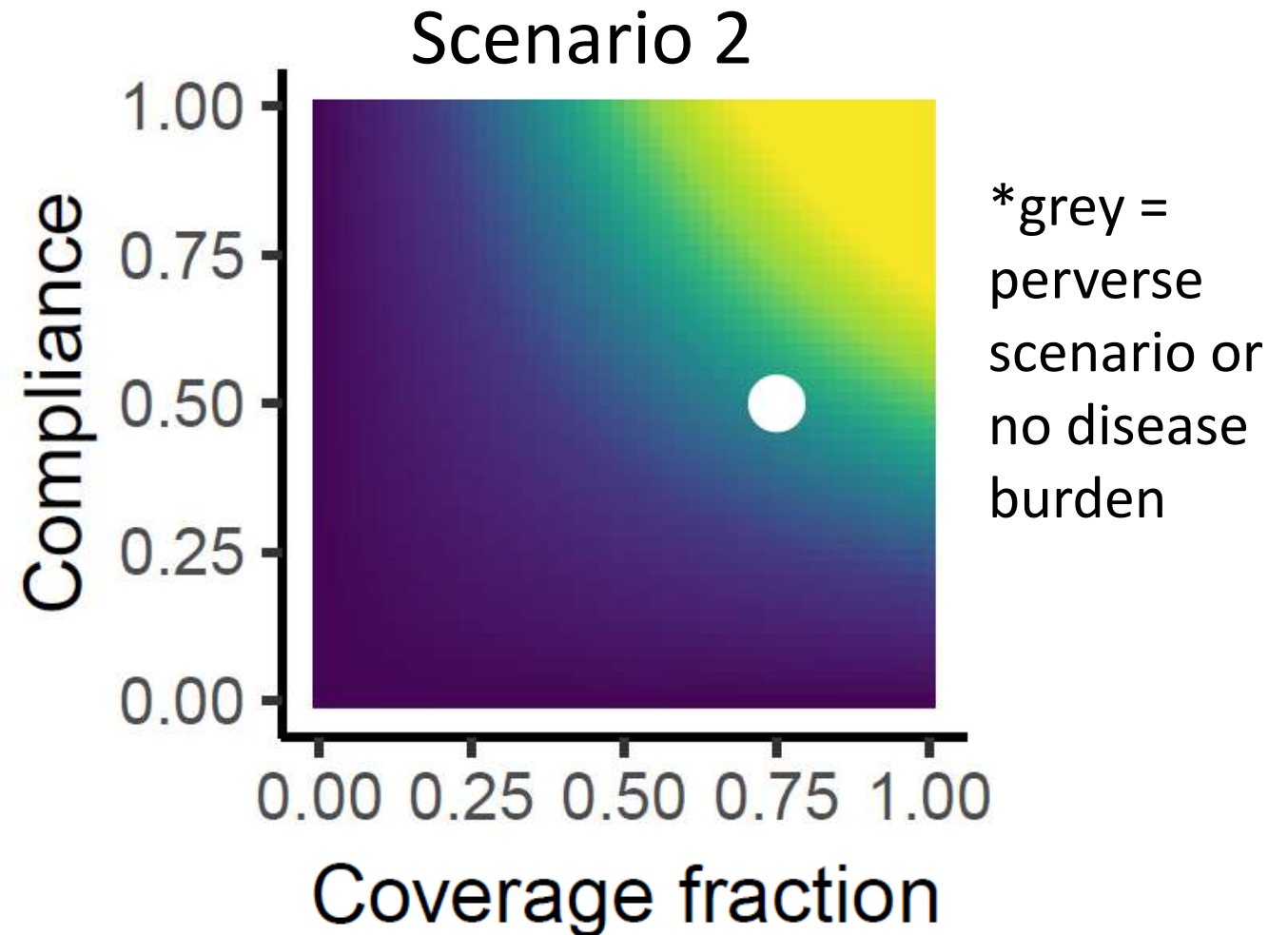
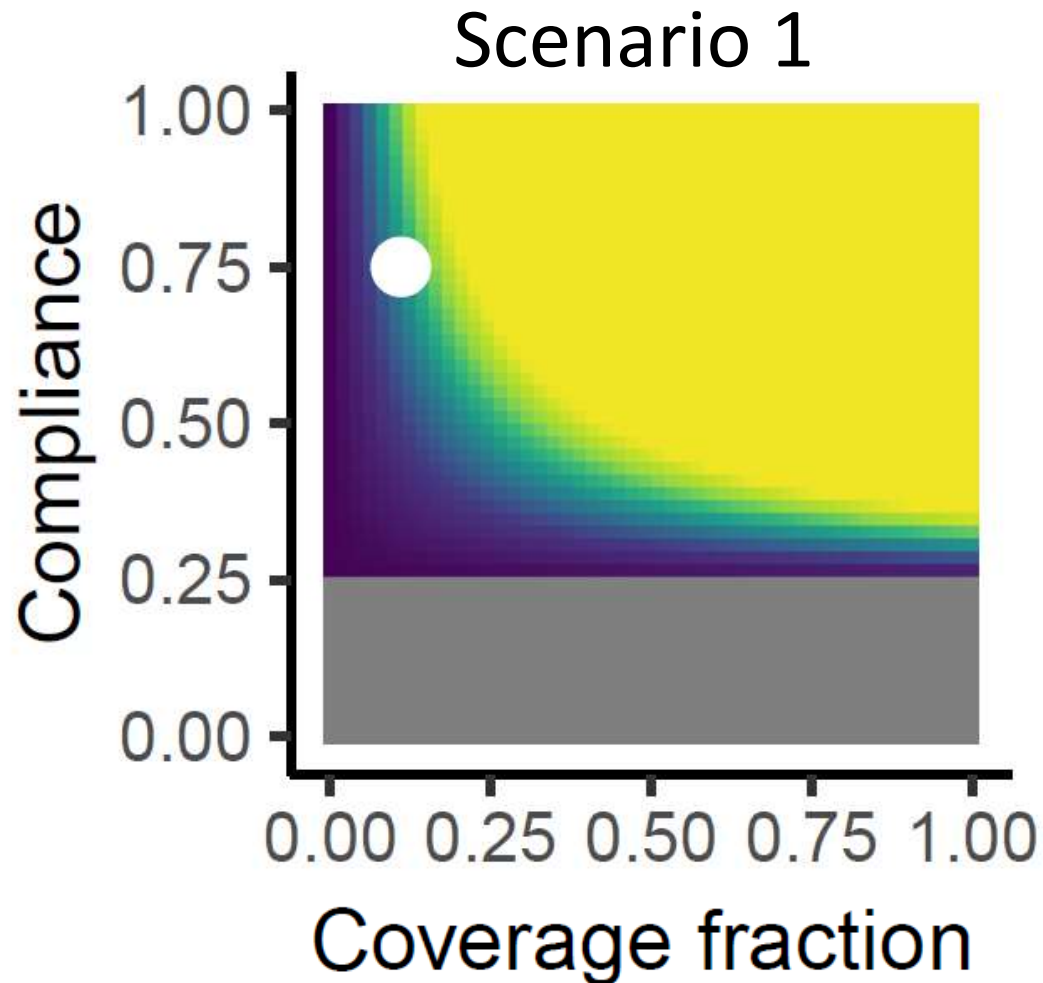
At the Scenario 1 baseline (A), effectiveness is **highly sensitive** to coverage and **less sensitive** to compliance



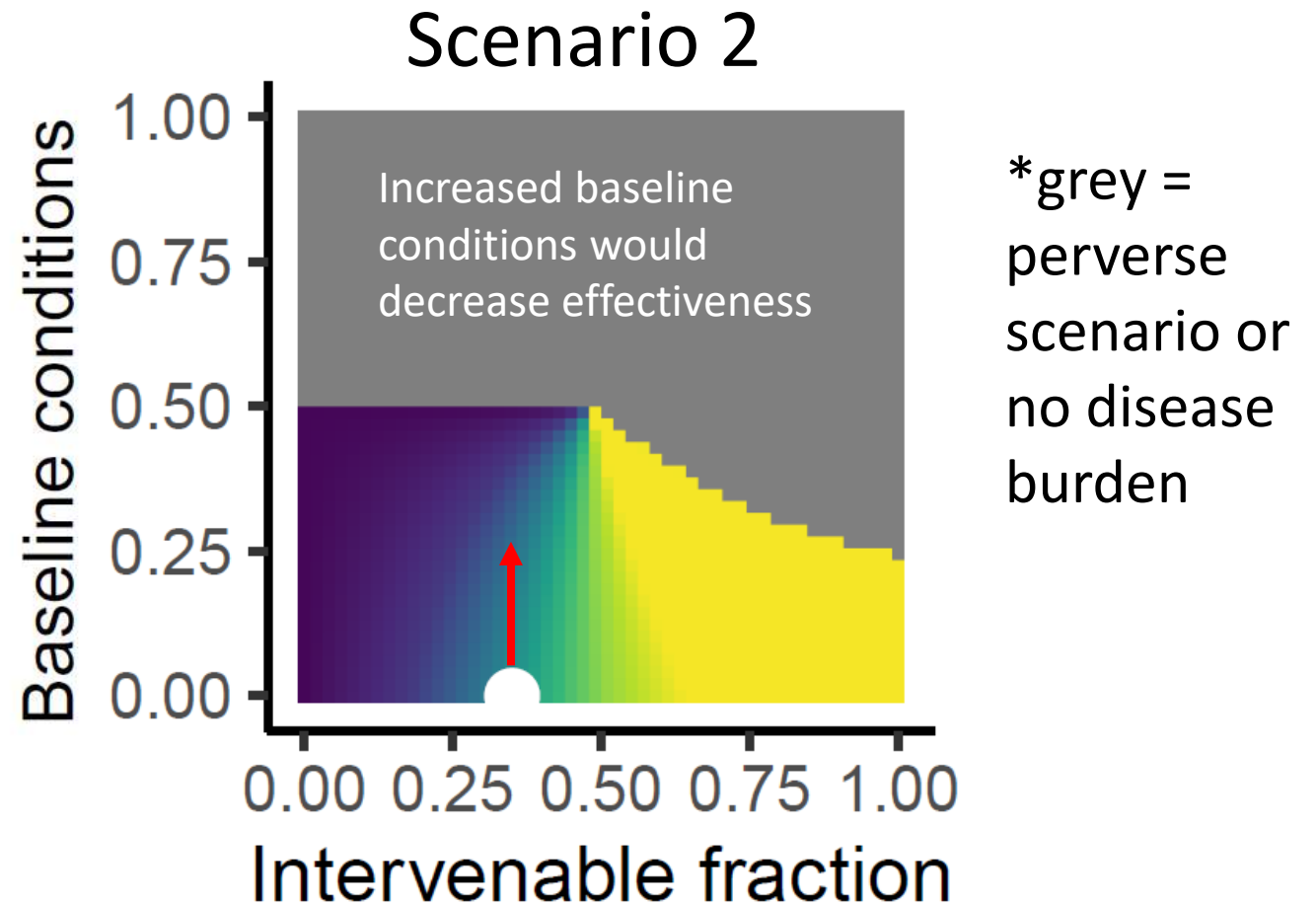
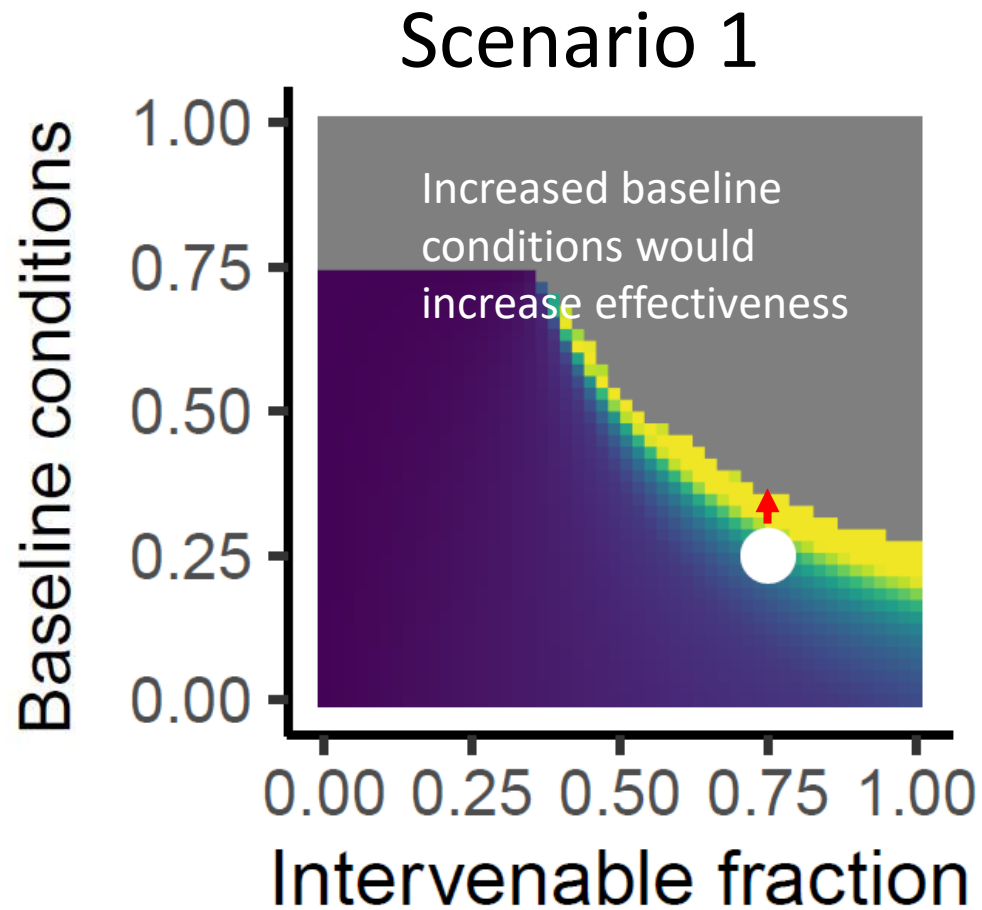
But at another point with 50% effectiveness (B), effectiveness is highly sensitive to compliance and less sensitive to coverage.



And these sensitivities further depend on the other intervention and contextual factors.



Indeed, in some cases, a combination of factors could increase effectiveness in one scenario but *decrease* it in another.



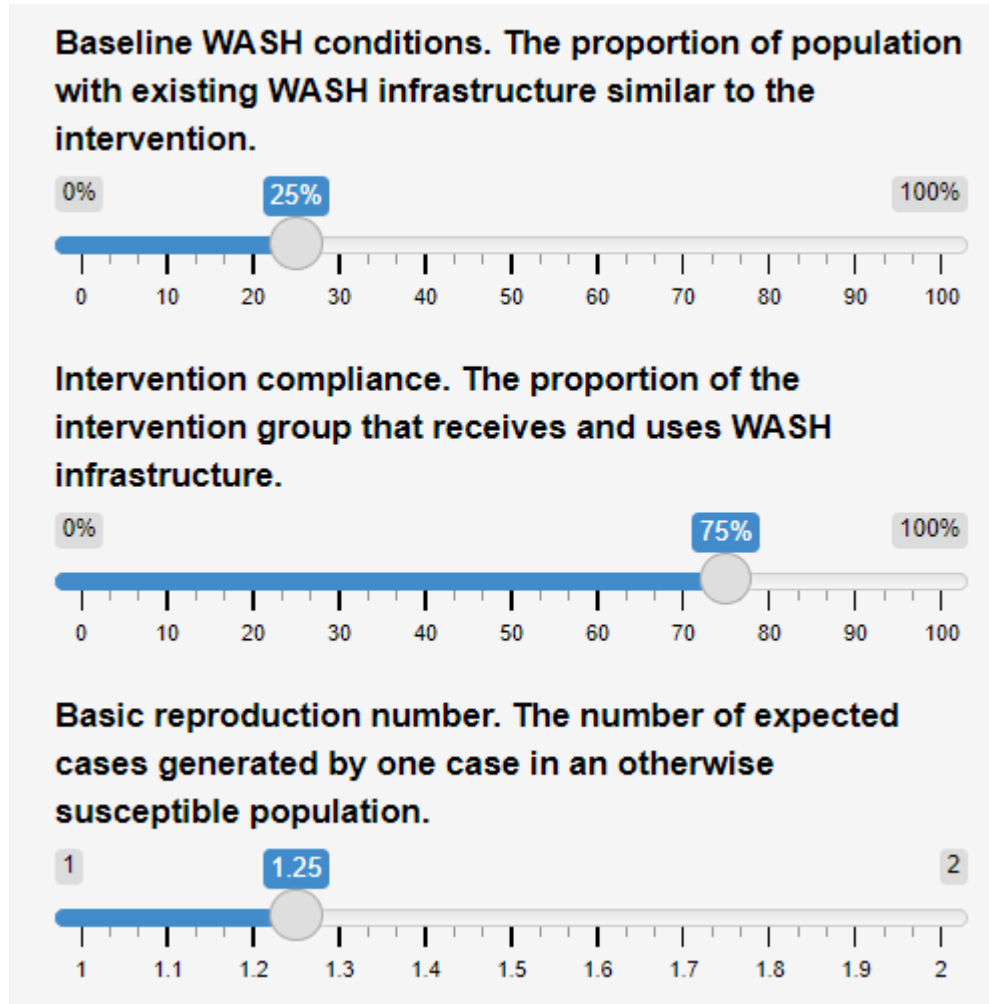
Thus, it is important
project an intervention's
effectiveness in a specific
context.

We have created a **public-
use tool** to facilitate
intervention planning for
local contexts.



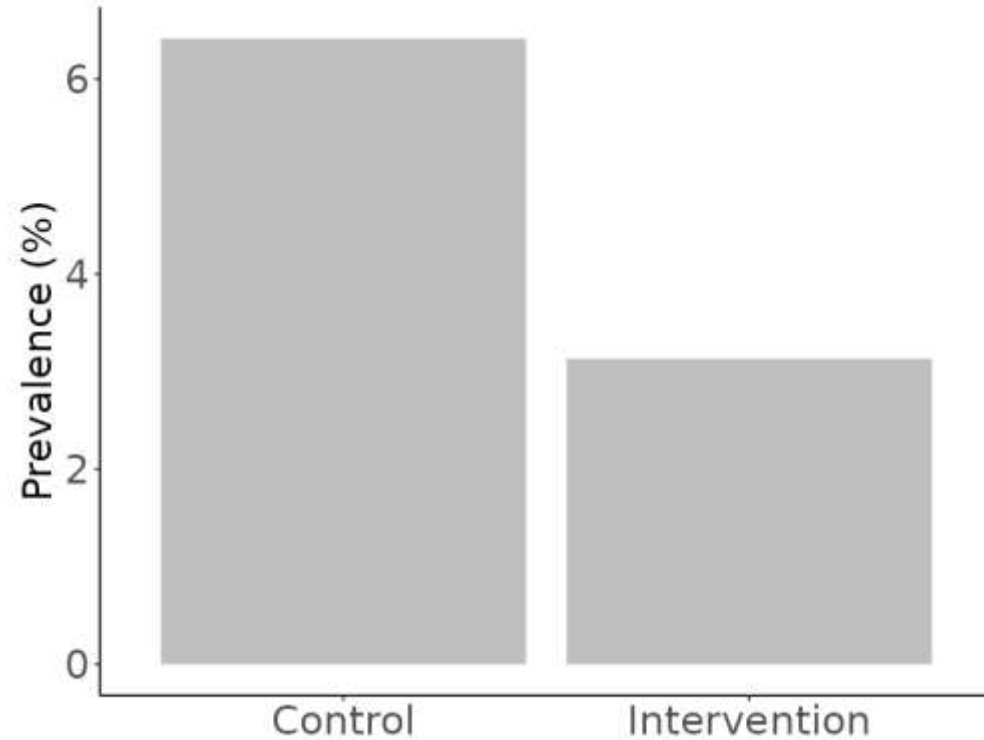
https://umich-biostatistics.shinyapps.io/sise_rct/

Set your intervention and contextual factors



https://umich-biostatistics.shinyapps.io/sise_rct/

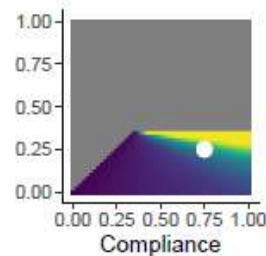
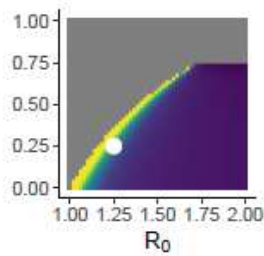
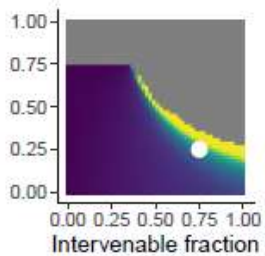
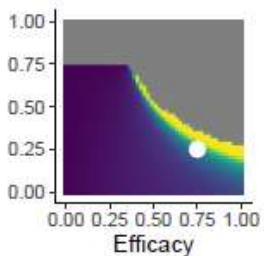
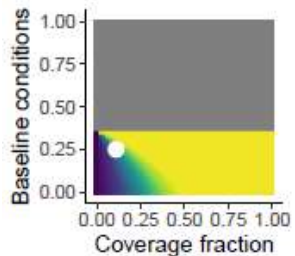
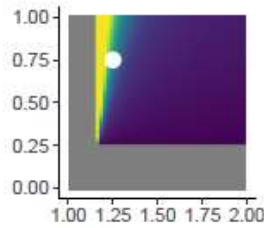
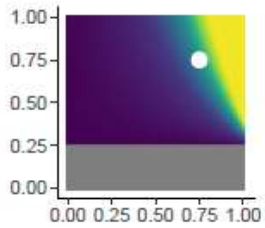
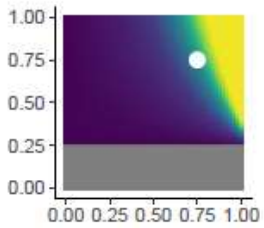
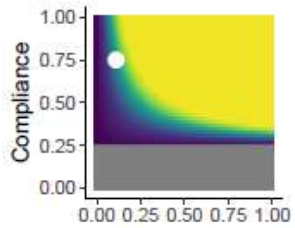
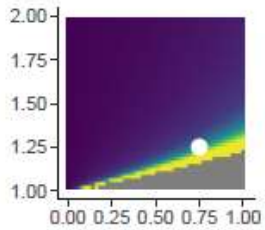
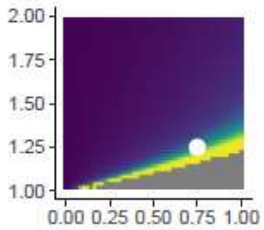
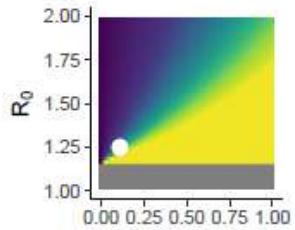
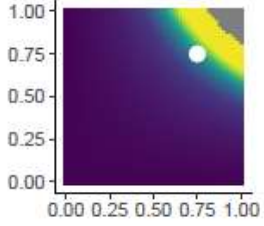
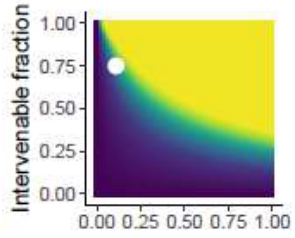
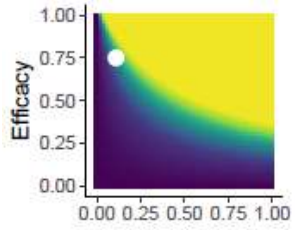
Estimate the baseline prevalence and intervention effectiveness



https://umich-biostatistics.shinyapps.io/sise_rct/

Prevalence in control arm	Prevalence in intervention arm	Non-intervenable prevalence	Intervention effectiveness
6.4	3.1	0	51.2%

See how sensitive the intervention is to each intervention and contextual factor.



https://umich-biostatistics.shinyapps.io/sise_rct/

Conclusions

- Not surprisingly, the sensitivity of intervention effectiveness to an intervention or contextual factor is highly sensitive to the other factors.
- These factors likely underly the heterogeneity in trial outcomes.
- Local contextual factors must be accounted for when developing priorities for interventions.
- Mechanistic modeling can both aid in this intervention planning and help to estimate how the results of an intervention in one location will translate to another location.

Acknowledgments

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- Ayse Ercumen
- Stephen Luby
- Amy Pickering
- Claire Null
- Mahbubur Rahman



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Questions?

Shiny app tool



https://umich-biostatistics.shinyapps.io/sise_rct/

Preprint paper



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