

# **Two small-data $n \ll p$ case studies: hydroxychloroquine, Marseilles and Meijel**

**High dimensional data – One World Symposium**

Thursday 27 August, 2020

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**Didier Raoult**

Gautret et al. (2020), Hydroxychloroquine and azithromycin as a treatment of COVID-19: results of an open-label non-randomized clinical trial. *International Journal of Antimicrobial Agents* 56 105949 (6pp.)

<https://rpubs.com/gill1109/raoult>



**Rob Elens, MD**

Family doctor and alternative medicine practitioner, Meijel, Netherlands

Dutch TV appearances, Dr Elens is a figurehead of a popular pro-HCQ movement

<https://rpubs.com/gill1109/elens>

## Marseilles

	good outcome	bad outcome
treatment	15	11
control	2	14

p-value = 0.004491

Gautret et al. (2020)

6 patients removed from treatment group  
for “non-compliance”

Remaining: 14, 6; 2, 14

p-value = 0.001

## Meijel

	good outcome	bad outcome
treatment	13	13
control	0	10

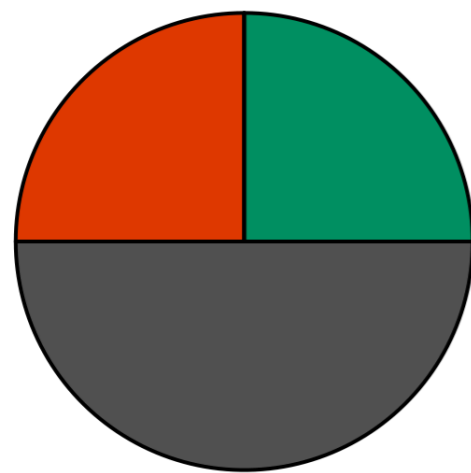
p-value = 0.005848

## Fisher exact tests

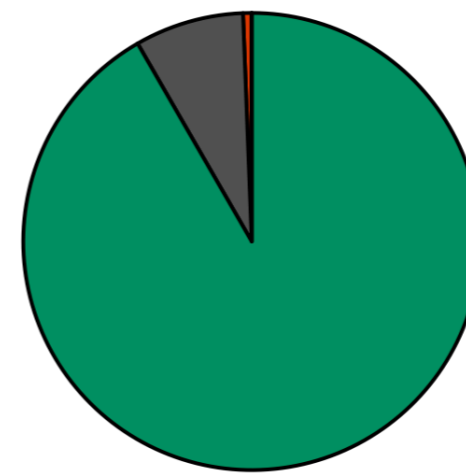
“Outcome” = Covid-19 virus absent/present after 6 days

# The Bayesian conclusion (Marseilles)

## JASP; AB test (R)



- $P(H+) = 0.250$
- $P(H-) = 0.250$
- $P(H0) = 0.500$

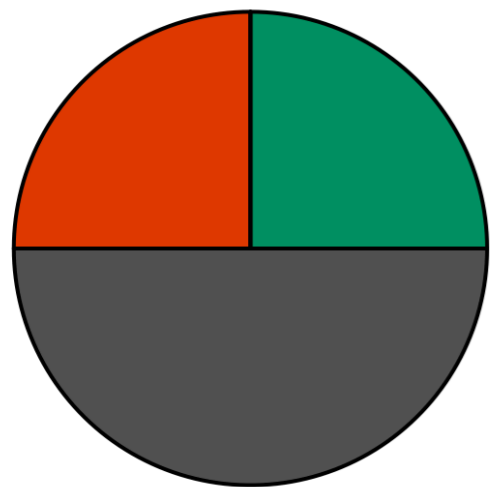


- $P(H+ | \text{data}) = 0.917$
- $P(H- | \text{data}) = 0.006$
- $P(H0 | \text{data}) = 0.077$

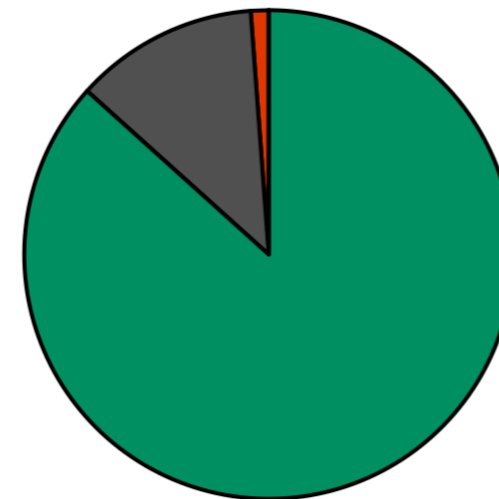
Left: prior; right: posterior  
A posteriori, still 8% chance of no difference!

# The Bayesian conclusion (Meijel)

## JASP; AB test (R)



- $P(H+) = 0.250$
- $P(H-) = 0.250$
- $P(H0) = 0.500$



- $P(H+ | \text{data}) = 0.867$
- $P(H- | \text{data}) = 0.012$
- $P(H0 | \text{data}) = 0.121$

Left: prior; right: posterior

A posteriori, still 12% chance of no difference at all!

# Confounders

**Data = two ca. 40 x 40 spreadsheets**

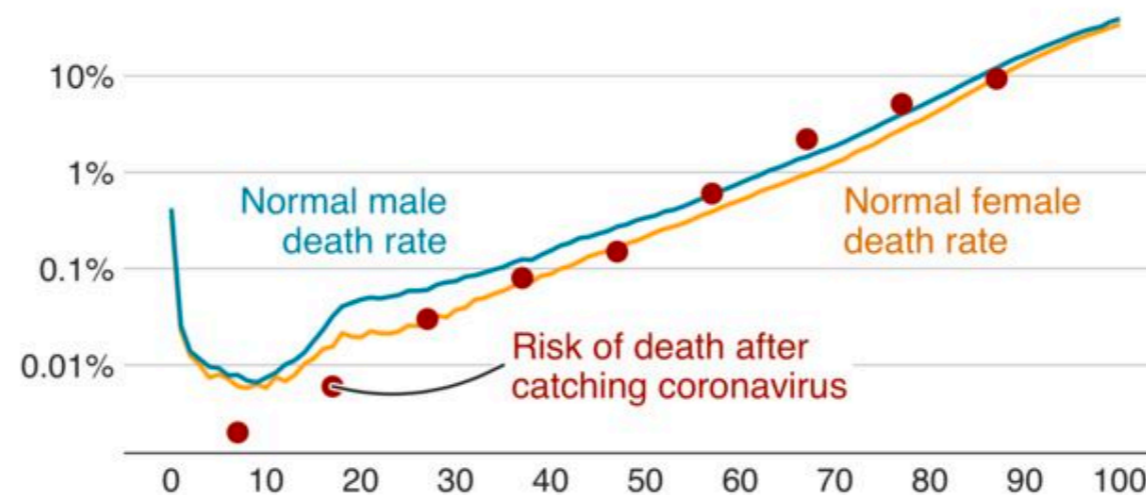
- Treatment (binary)
- Outcome (disease free after 6 days)
- Age (from 20 to 90); sex
- Comorbidities (obesity, diabetes, ... : yes/no)
- Symptoms (sense of smell gone, fever, ... : yes/no)
- Also some other numerical variables (blood pressure, ...)

# First findings

- “Common sense” and medical knowledge reduces # confounding variables to half a dozen
- Logistic regression on all 6 gives nonsense
- Lasso on all 6 gives nonsense
- Logistic regression with just one or two covariates shows realistic (significant) coefficients; effect of age (ages range from 20 to 90) exactly what we expect ... just like a life table

## Risk of dying if you get coronavirus v normal annual risk

Risk of dying each year by age (GB)



Log scale used to see differences in rates at younger ages

Source: Prof. Sir David Spiegelhalter, ONS, Imperial College London

# Present experiments

- Compute a “standardised age” from mortality statistics (risk of death this year, given age, sex, comorbidities)
- Compute a “symptoms severity index” – we can now do this using much bigger published data sets!
- Now we have just two continuous covariates (use simple spline curve) and one discrete. Results coming very soon, I hope!

Hoang Van Thuan,  
Marseilles and Vietnam



Leila  
Schneps,  
Paris

Dipro Mondal,  
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