

What have we learned from
Covid-19 – the good and the
bad and the TB

Diagnostics

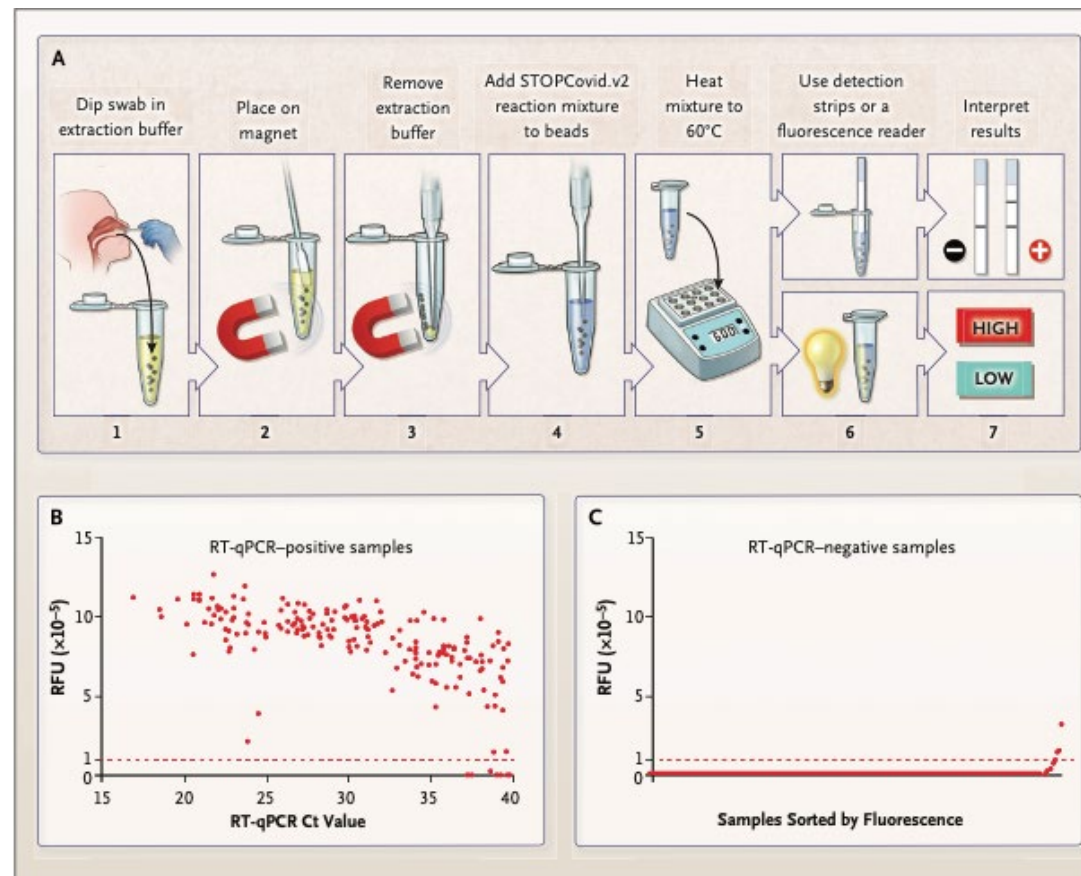
- The good
 - Widespread implementation of PCR-based testing
 - Some innovative new technologies
- The bad
 - No point-of-care PCR test
 - Rapid antigen-based testing not widely deployed

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CORRESPONDENCE



Detection of SARS-CoV-2 with SHERLOCK One-Pot Testing



Diagnostics – the lessons for TB

- PCR widely established and available
- Difficult to reduce new technology to practice

Therapeutics

- The good
 - After a slow start, many clinical trials
 - Some success with glucocorticoids and other anti-inflammatories
- The bad
 - No truly new agents
 - No dramatic successes
 - Multiple simultaneous trials with low power to detect differences
 - No real proof-of-concept for antivirals

Row	Saved	Status	Study Title	Conditions	Interventions	Locations
1	<input type="checkbox"/>	Not yet recruiting	Effectiveness of Hydroxychloroquine in Covid-19 Patients	<ul style="list-style-type: none"> COVID19 	<ul style="list-style-type: none"> Drug: Hydroxychloroquine 200 Mg Oral Tablet Drug: Azithromycin 500Mg Oral Tablet Dietary Supplement: Glucose tablets 	<ul style="list-style-type: none"> Ayub Teaching Institution Abbottābād, K.p.k, Pakistan
2	<input type="checkbox"/>	Withdrawn	Hydroxychloroquine in SARS-CoV-2 (COVID-19) Pneumonia Trial	<ul style="list-style-type: none"> SARS-CoV-2 Pneumonia COVID-19 	<ul style="list-style-type: none"> Drug: Hydroxychloroquine 	<ul style="list-style-type: none"> Kootenai Health Coeur d'Alene, Idaho, United States
3	<input type="checkbox"/>	Withdrawn	Assessing Hydroxychloroquine in Patients With SARS-CoV-2 (COVID-19)	<ul style="list-style-type: none"> COVID-19 SARS-CoV-2 	<ul style="list-style-type: none"> Drug: Hydroxychloroquine Drug: Placebo 	<ul style="list-style-type: none"> Oregon Health and Science University Portland, Oregon, United States
4	<input type="checkbox"/>	Withdrawn	The University of the Philippines Hydroxychloroquine PEP Against COVID-19 Trial	<ul style="list-style-type: none"> COVID-19 	<ul style="list-style-type: none"> Drug: Hydroxychloroquine plus standard preventive measures Drug: Placebo plus standard preventive measures 	<ul style="list-style-type: none"> Philippine General Hospital - University of the Philippines Manila Manila, Philippines
5	<input type="checkbox"/>	Recruiting	Use of Bromhexine and Hydroxychloroquine for Treatment of COVID-19 Pneumonia	<ul style="list-style-type: none"> Covid-19 	<ul style="list-style-type: none"> Drug: Bromhexine Oral Tablet and/or hydroxychloroquine tablet 	<ul style="list-style-type: none"> SB Celje Celje, Slovenia
6	<input type="checkbox"/>	Active, not recruiting	Hydroxychloroquine vs. Azithromycin for Hospitalized Patients With Suspected or Confirmed COVID-19	<ul style="list-style-type: none"> COVID-19 	<ul style="list-style-type: none"> Drug: Hydroxychloroquine Drug: Azithromycin 	<ul style="list-style-type: none"> Intermountain Medical Center Murray, Utah, United States University of Utah Salt Lake City, Utah, United States
7	<input type="checkbox"/>	Withdrawn	Hydroxychloroquine Azithromycin COVID-19 Pregnancy Trial	<ul style="list-style-type: none"> COVID19 	<ul style="list-style-type: none"> Drug: Hydroxychloroquine and azithromycin treatment Other: conventional management of patients 	
8	<input type="checkbox"/>	Terminated	Hydroxychloroquine for the Treatment of Mild COVID-19 Disease	<ul style="list-style-type: none"> COVID-19 	<ul style="list-style-type: none"> Drug: Hydroxychloroquine Drug: Placebo 	<ul style="list-style-type: none"> Institute for Tropical Medicine Tübingen, Germany
9	<input type="checkbox"/>	Not yet recruiting	Hydroxychloroquine and Nitazoxanide Combination Therapy for COVID-19	<ul style="list-style-type: none"> COVID-19 	<ul style="list-style-type: none"> Combination Product: Hydroxychloroquine plus Nitazoxanide Other: Standard care 	
10	<input type="checkbox"/>	Recruiting	High-dose Hydroxychloroquine for the Treatment of Ambulatory Patients With Mild COVID-19	<ul style="list-style-type: none"> COVID-19 	<ul style="list-style-type: none"> Drug: Hydroxychloroquine 	<ul style="list-style-type: none"> University of Chicago Chicago, Illinois, United States
11	<input type="checkbox"/>	Terminated	ALBERTA HOPE COVID-19 for the Prevention of Severe COVID19 Disease	<ul style="list-style-type: none"> COVID-19 	<ul style="list-style-type: none"> Drug: Hydroxychloroquine 	<ul style="list-style-type: none"> University of Calgary/Foothills Medical Centre Calgary, Alberta, Canada University of Alberta Edmonton, Alberta, Canada
12	<input type="checkbox"/>	Active, not recruiting	Hydroxychloroquine Use in Hospitalized Patients With COVID-19: Impact on Progression to Severe or Critical Disease	<ul style="list-style-type: none"> COVID-19 	<ul style="list-style-type: none"> Drug: Hydroxychloroquine Drug: Placebo 	<ul style="list-style-type: none"> Wellstar Kennestone Hospital Marietta, Georgia, United States
13	<input type="checkbox"/>	Recruiting	Hydroxychloroquine (HCQ) as Post Exposure Prophylaxis (PEP) for Prevention of COVID-19	<ul style="list-style-type: none"> Covid19 COVID-19 Prevention 	<ul style="list-style-type: none"> Drug: Hydroxychloroquine (HCQ) Other: Standard care Other: Placebo 	<ul style="list-style-type: none"> Post Graduate Institute of Medical Education and Research Chandigarh, India
14	<input type="checkbox"/>	Recruiting	Randomized Placebo-controlled Trial of Hydroxychloroquine in Outpatient Cases With Coronavirus Disease 2019 (COVID-19)	<ul style="list-style-type: none"> COVID-19 	<ul style="list-style-type: none"> Drug: Hydroxychloroquine Drug: Placebo 	<ul style="list-style-type: none"> Centro de Pesquisas Clínicas Dr. Marco Mota HCOR Maceió, Alagoas, Brazil Hospital e Clínica São Roque Ipiaú, Bahia, Brazil Clínica Otorhinus

Therapeutics – lessons for TB

- Maintaining a drug development pipeline is critical
- Coordinating expensive and lengthy trials across multiple groups is far more efficient

Vaccines

- The good

- Incredibly rapid development and deployment
- New technologies that promise rapid development
- Extensive real-world safety testing for vaccine delivery methods
- Large RCTs done quickly while maintaining high quality

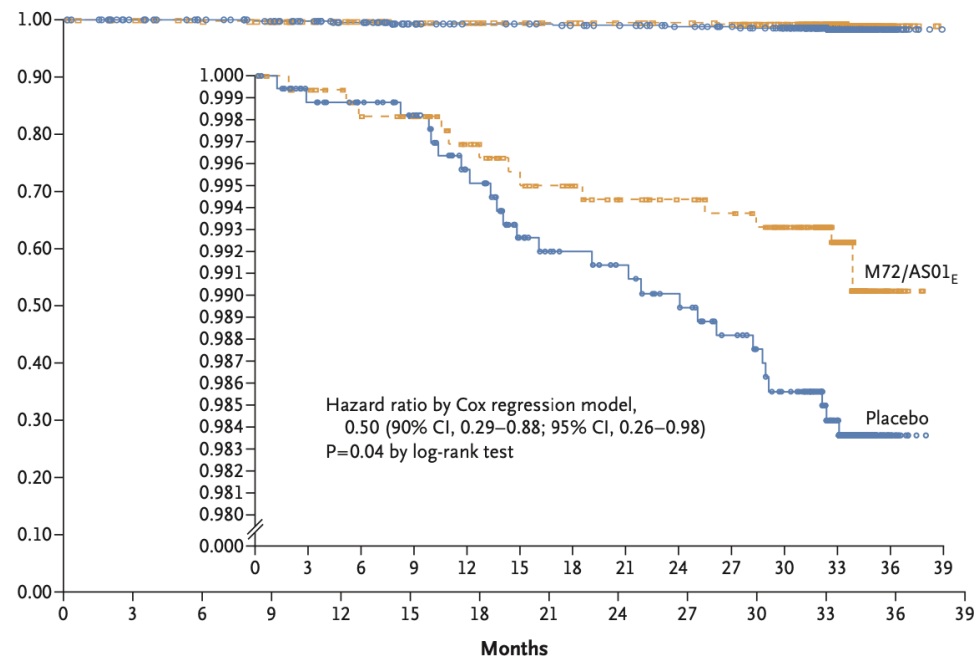
- The bad

- We still don't have a reliable biomarker of vaccine efficacy

ORIGINAL ARTICLE

Final Analysis of a Trial of M72/AS01_E Vaccine to Prevent Tuberculosis

D.R. Tait, M. Hatherill, O. Van Der Meeren, A.M. Ginsberg, E. Van Brakel, B. Salaun, T.J. Scriba, E.J. Akite, H.M. Ayles, A. Bollaerts, M.-A. Demoitié, A. Diacon, T.G. Evans, P. Gillard, E. Hellström, J.C. Innes, M. Lempicki, M. Malahleha, N. Martinson, D. Mesia Vela, M. Muyoyeta, V. Nduba, T.G. Pascal, M. Tameris, F. Thienemann, R.J. Wilkinson, and F. Roman



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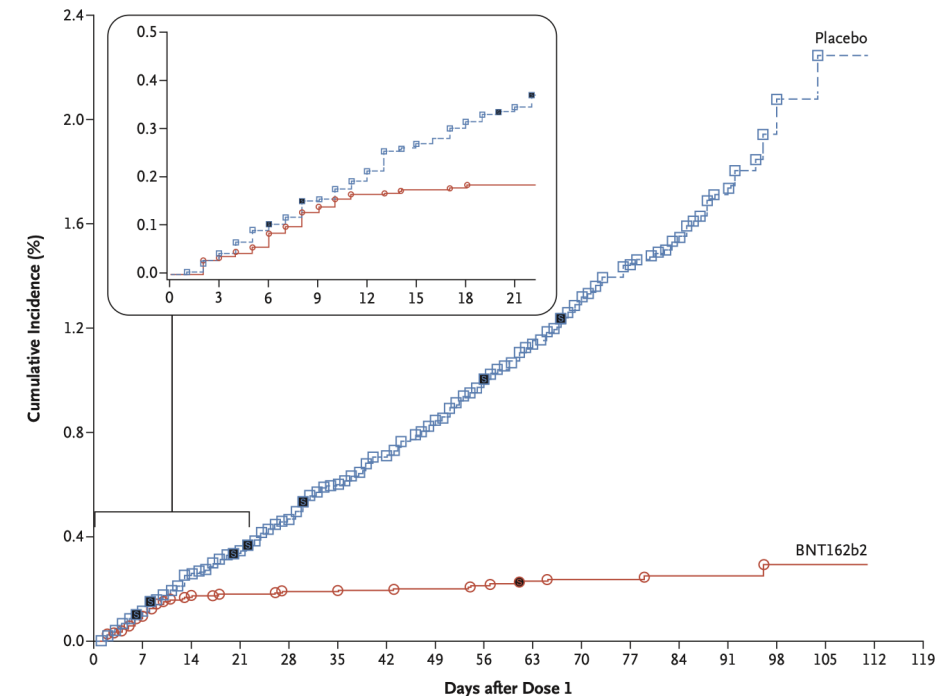
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Safety and Efficacy of the BNT162b2 mRNA Covid-19 Vaccine

Fernando P. Polack, M.D., Stephen J. Thomas, M.D., Nicholas Kitchin, M.D., Judith Absalon, M.D., Alejandra Gurtman, M.D., Stephen Lockhart, D.M., John L. Perez, M.D., Gonzalo Pérez Marc, M.D., Edson D. Moreira, M.D., Cristiano Zerbini, M.D., Ruth Bailey, B.Sc., Kena A. Swanson, Ph.D., Satrajit Roychoudhury, Ph.D., Kenneth Koury, Ph.D., Ping Li, Ph.D., Warren V. Kalina, Ph.D., David Cooper, Ph.D., Robert W. Frenc, Jr., M.D., Laura L. Hammitt, M.D., Özlem Türeci, M.D., Haylene Nell, M.D., Axel Schaefer, M.D., Serhat Ünal, M.D., Dina B. Tresnan, D.V.M., Ph.D., Susan Mather, M.D., Philip R. Dormitzer, M.D., Ph.D., Uğur Şahin, M.D., Kathrin U. Jansen, Ph.D., and William C. Gruber, M.D., for the C4591001 Clinical Trial Group*



Vaccines – the lessons for TB

- mRNA (and, possibly, adenovirus vectors) provides a rapid way to test new antigens without adjuvants with limited availability
- Without biomarkers or new and persuasive models, there is still no substitute for large phase 3 trials