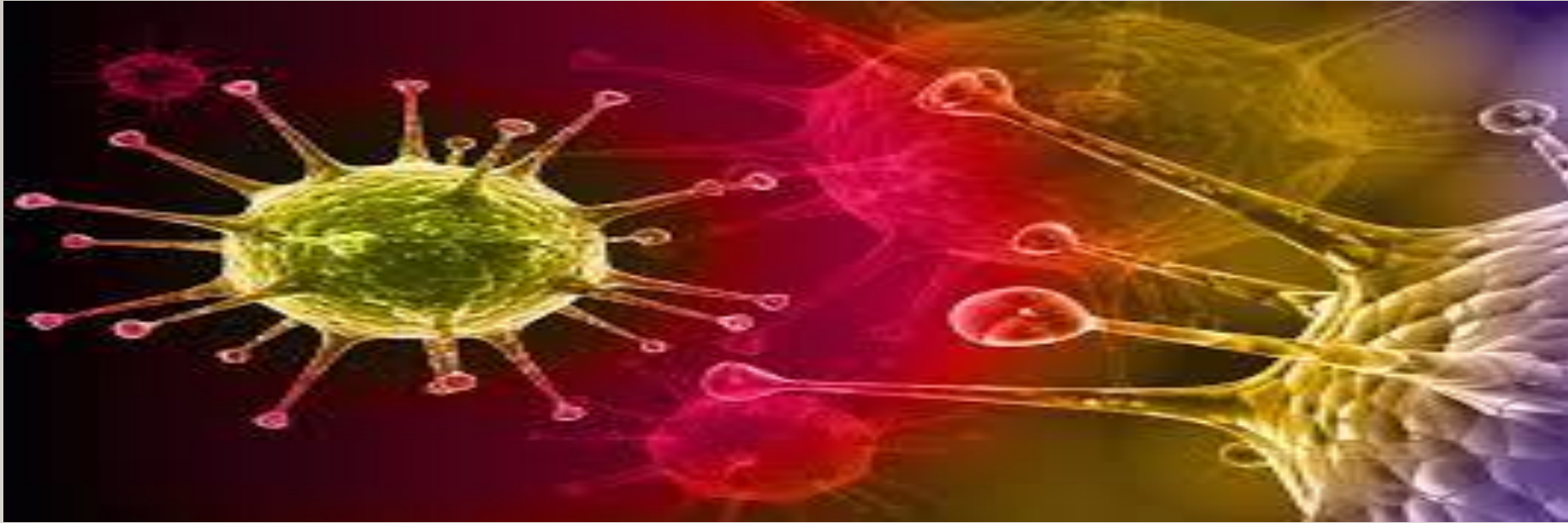




TOP 10 INFECTIOUS DISEASES ARTICLE IN **2023**

MASOUD MARDANI, MD, MPH, FIDSA, FESCMID

PROFESSOR OF INFECTIOUS DISEASES , SBMU, TEHRAN ,IRAN



In 2023 at least 10 practice changing studies on prevention and treatment of a broad range of infectious diseases has been published.

These research projects have been lead to advancement of clinical medical practice of infectious diseases specially in immunocompromised cancer patient.

Due to change of practice of infectious diseases in recent years we preferred to have a brief presentation of 10 top infectious diseases article in 2023.

BIVALENT PREFUSION F VACCINE IN PREGNANCY TO PREVENT RSV ILLNESS IN INFANTS

- A 3, double-blind trial conducted in 18 countries, pregnant women at 24 through 36 weeks' gestation to receive a *single intramuscular injection of 120 µg of a bivalent RSV prefusion F protein-based (RSVpreF)* vaccine or placebo.
- The two primary efficacy end points were medically attended severe RSV-associated lower respiratory tract illness and medically attended RSV-associated lower respiratory tract illness in infants within *90, 120, 150, and 180* days after birth.
- Efficacy of a *bivalent prefusion-stabilized F glycoprotein RSV vaccine* was evaluated in pregnant women (to prevent disease in their infants) and in older adults, which lead to which led to *FDA approval* of, and *Advisory Committee on Immunization Practices recommendations* for, these vaccines.

The NEW ENGLAND JOURNAL of MEDICINE

ESTABLISHED IN 1812 APRIL 20, 2023 VOL. 384 NO. 16

Bivalent Prefusion F Vaccine in Pregnancy to Prevent RSV Illness in Infants

B. Ewerghoan, S.A. Madhi, I. Munjal, E.A.F. Simões, B.A. Fahud, C. Ulagar, J. Baber, G. Pérez Marc, D. Radley, E. Shaha, J. Glennerink, H. Snugga, J. Baber, P. Zachariah, S.L. Sarmalata, M. Fouzati, T. Adani, N. Perrotin, M.A. Van Houten, A. Karimik, L.-M. Hwang, L.J. Bost, T. Otsuki, S.L. Vargas, J. Gullatt, B. Tapiero, R.T. Stein, F.P. Polack, H.J. Zar, N.B. Shafer, M. Duran Padilla, P.C. Richmond, K. Knury, K. Schneider, E.V. Kalinova, D. Cooper, K.U. Jansen, A.S. Anderson, K.A. Swanson, W.C. Gruber, and A. Gurtman, for the MATISSE Study Group*

ABSTRACT

BACKGROUND
Whether vaccination during pregnancy could reduce the burden of respiratory syncytial virus (RSV)-associated lower respiratory tract illness in newborns and infants is uncertain.

METHODS
In this phase 3, double-blind trial conducted in 18 countries, we randomly assigned, in a 1:1 ratio, pregnant women at 24 through 36 weeks' gestation to receive a single intramuscular injection of 120 µg of a bivalent RSV prefusion F protein-based (RSVpreF) vaccine or placebo. The two primary efficacy end points were medically attended severe RSV-associated lower respiratory tract illness and medically attended RSV-associated lower respiratory tract illness in infants within 90, 120, 150, and 180 days after birth. A lower boundary of the confidence interval for vaccine efficacy (95% confidence interval [CI] at 90 days; 97.58% CI at later intervals) greater than 20% was considered to meet the success criterion for vaccine efficacy with respect to the primary end points.

RESULTS
At this prespecified interim analysis, the success criterion for vaccine efficacy was met with respect to one primary end point. Overall, 3682 maternal participants received vaccine and 3676 received placebo; 3570 and 3558 infants, respectively, were evaluated. Medically attended severe lower respiratory tract illness occurred within 90 days after birth in 6 infants of women in the vaccine group and 33 infants of women in the placebo group (vaccine efficacy, 81.8%; 95.5% CI, 40.6 to 96.3); 19 cases and 62 cases, respectively, occurred within 180 days after birth (vaccine efficacy, 69.4%; 97.58% CI, 44.5 to 84.1). Medically attended RSV-associated lower respiratory tract illness occurred within 90 days after birth in 34 infants of women in the vaccine group and 56 infants of women in the placebo group (vaccine efficacy, 57.1%; 93.9% CI, 34.7 to 79.8); these results did not meet the statistical success criterion. No safety signals were detected in maternal participants or in infants and toddlers up to 36 months of age. The incidences of adverse events reported within 1 month after injection or within 1 month after birth were similar in the vaccine group (13.8% of women and 37.1% of infants) and the placebo group (13.1% and 34.9%, respectively).

CONCLUSIONS
RSVpreF vaccine administered during pregnancy was effective against medically attended severe RSV-associated lower respiratory tract illness in infants, and no safety concerns were identified. (Funded by Pfizer; MATISSE ClinicalTrials.gov number, NCT04424116.)

The authors' full names, academic degrees, and affiliations are listed in the Appendix. Dr. Madhi can be contacted at isma.madhi@pfi.com or at Vaccine Research and Development, Pfizer, 400 N. Middletown Rd., Pearl River, NY 10968.

*The members of the MATISSE Study Group are listed in the Supplementary Appendix, available at [NEJM.org](https://doi.org/10.1056/NEJMoa2214440).

Dr. Kanyemitsu, Madhi, and Murali were included equally to this article.

This article was published on April 3, 2023, at [NEJM.org](https://doi.org/10.1056/NEJMoa2214440).

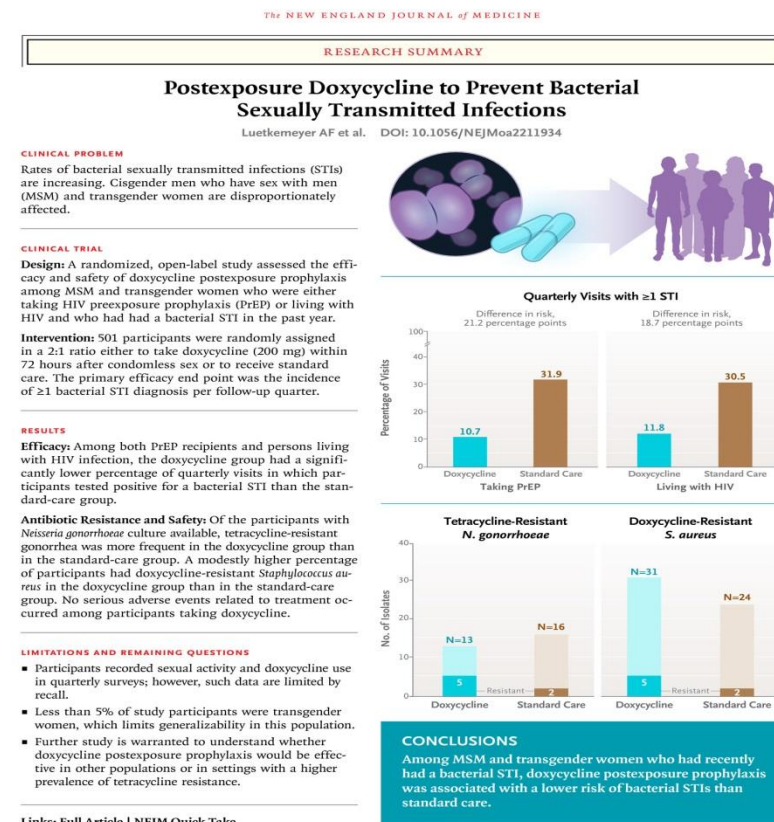
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NEJM JUNE 18, 2023 | [NEJM.org](https://doi.org/10.1056/NEJMoa2214440) | 2451

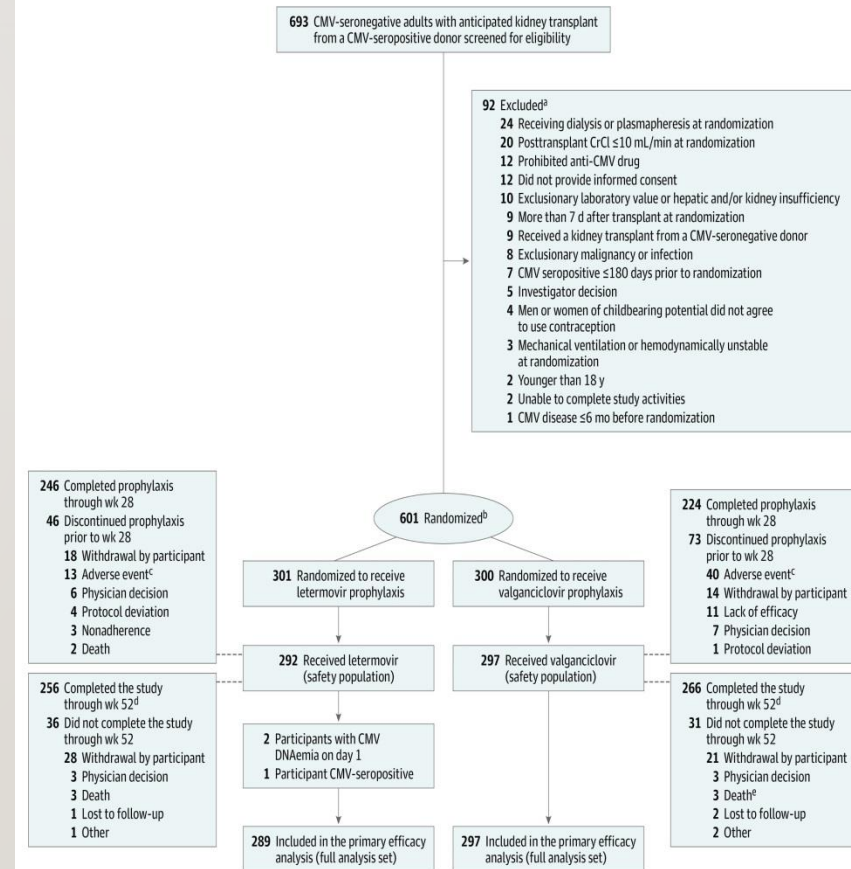
POSTEXPOSURE PROPHYLAXIS WITH DOXYCYCLINE LOWERED INCIDENCE OF BACTERIAL SEXUALLY TRANSMITTED INFECTIONS

- This trial demonstrating that *postexposure prophylaxis with doxycycline* prevents bacterial *sexually transmitted infections (STIs)* , With the rising incidence of bacterial sexually transmitted infections in many countries.
- In all, 501 participants (327 [PrEP group] and 174 [PLWH group]) were randomized to a *single 200-mg dose of DOXY PEP* or standard of care (SOC) without DOXY.
- They found that *postexposure doxycycline significantly reduced bacterial STI incidence in at-risk MSM and transgender women* who were on PrEP or were living with HIV



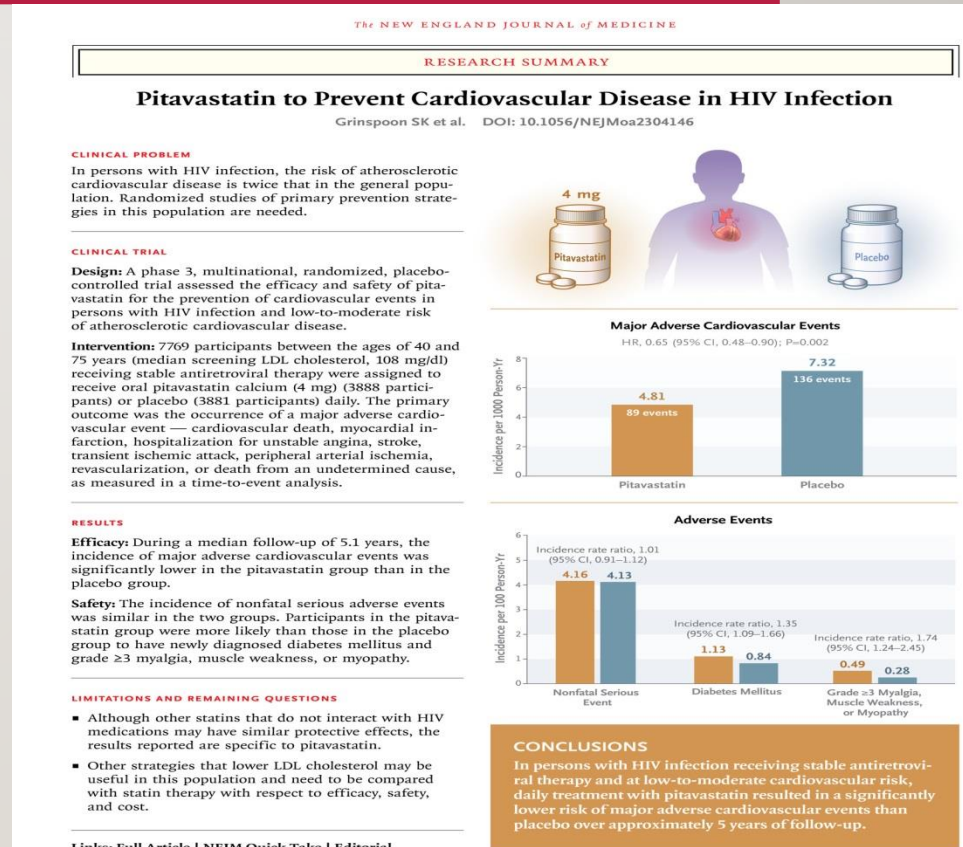
LETERMIVIR VS VALGANCICLOVIR FOR PROPHYLAXIS OF CYTOMEGALOVIRUS IN HIGH-RISK KIDNEY TRANSPLANT RECIPIENTS

- This study led the **FDA** to approve **letermovir** for prevention of **CMV** disease in high-risk kidney transplant recipients.
- Randomized, double-masked, double-dummy, noninferiority, phase 3 trial in adult CMV-seronegative kidney transplant recipients who received an organ from a CMV-seropositive donor at 94 participating sites.
- Participants were randomized in a 1:1 ratio to receive **letermovir, 480 mg**, orally daily (with acyclovir) or **valganciclovir, 900 mg**, orally daily (adjusted for kidney function) for **up to 200 days** after transplant, with matching placebos.
- Universal prophylaxis with valganciclovir is highly effective in preventing CMV disease during the early post-transplant period, but **leukopenia is common**, resulting in discontinuation of prophylaxis or changes to the immunosuppressive regimen.
- **Letermovir is a CMV-specific antiviral approved** for prophylaxis after stem cell transplant.



PITAVASTATIN TO PREVENT CARDIOVASCULAR DISEASE IN HIV INFECTION

- *People with HIV (PWH)* are at higher risk for cardiovascular disease (CVD) and ASCVD risk score than those without HIV.
- In phase 3 trial, randomly assigned **7769 participants with HIV infection** with a low-to-moderate risk of cardiovascular disease who were receiving antiretroviral therapy to receive **daily pitavastatin calcium (at a dose of 4 mg) or placebo**.
- The primary outcome was the occurrence of a major adverse cardiovascular event, which was defined as a composite **of cardiovascular death, myocardial infarction, hospitalization for unstable angina, stroke, transient ischemic attack, peripheral arterial ischemia, revascularization, or death from an undetermined cause**.
- The incidence of a **major adverse cardiovascular event** was **4.81 per 1000** person-years in the pitavastatin group and **7.32 per 1000** person-years in the placebo group.
- Participants with HIV infection who received pitavastatin **had a lower risk of a major adverse cardiovascular event** than those who received placebo over a median follow-up of 5.1 years





Dr. Hamid

Badali

SULBACTAM-DURLOBACTAM: A NEW WEAPON TO ATTACK ACINETOBACTER BAUMANNII

- This study has been lead [to FDA approval](#),
- *Acinetobacter baumannii* Infections caused by carbapenem-resistant *Acinetobacter baumannii* (CRAB) *have limited therapeutic options*.
- *Sulbactam*, a BLI developed >40 years ago, was often combined with ampicillin,
- *Durlobactam* is a novel non- β -lactam BLI that is a potent inhibitor of β -lactamases found in CRAB
- In an international phase 3 trial across **59 sites**, investigators randomized **177 hospitalized adults** with documented HABP/VABP due to *Acinetobacter calcoaceticus* to receive sul-dur or colistin.
- Both groups also received *imipenem/cilastatin*.
- sulbactam–durlobactam was non-inferior to colistin, both agents given in combination with imipenem–cilastatin, for the primary endpoint of 28-day all-cause mortality
- *Sulbactam–durlobactam* was well tolerated and could be an effective intervention to **reduce mortality** from serious infections caused by **carbapenem-resistant ABC, including multidrug-resistant strains**.

Efficacy and safety of sulbactam–durlobactam versus colistin for the treatment of patients with serious infections caused by *Acinetobacter baumannii*–*calcoaceticus* complex: a multicentre, randomised, active-controlled, phase 3, non-inferiority clinical trial (ATTACK)

Prof Keith S Kaye, MD ✉ • Prof Andrew F Shorr, MD • Prof Richard G Wunderink, MD • Prof Bin Du, MD •

Gabrielle E Poirier, BS • Khurram Rana, PharmD • et al. [Show all authors](#)



FDA-APPROVED ORAL FECAL MICROBIOTA THERAPY FOR PREVENTION OF RECURRENT *C. DIFFICILE* INFECTION

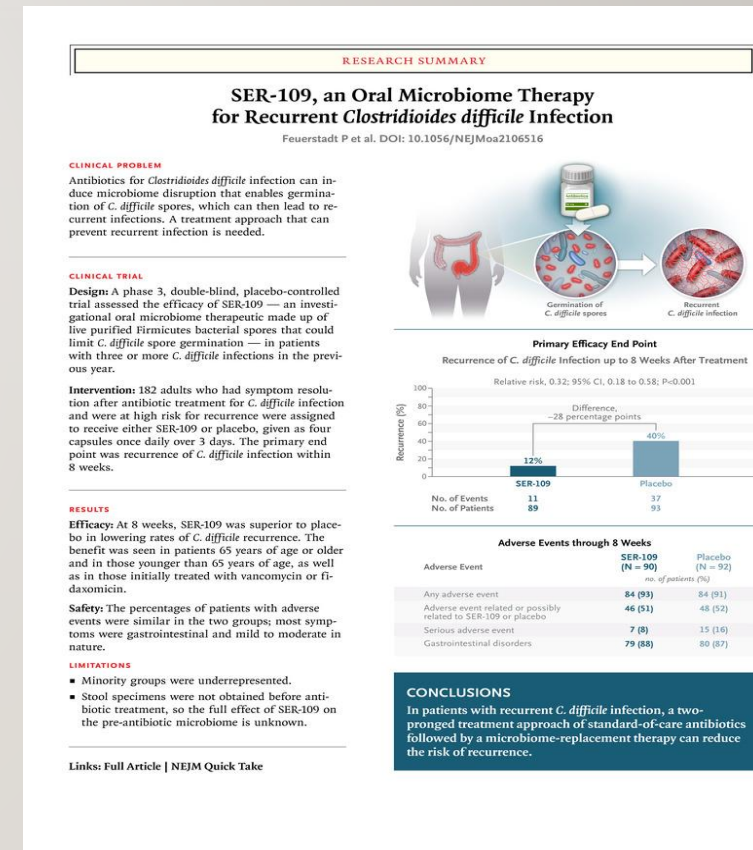
Standard treatments for *Clostridioides difficile* infection (CDI), while highly effective for resolving the initial episode, are limited by frequent recurrence.

Fecal microbiota therapy (FMT) after CDI treatment is recommended for patients with multiple episodes.

In a phase 3, double-blind, randomized, placebo-controlled trial in which patients who had had three or more episodes of *C. difficile* infection received **SER-109** or **placebo** after standard-of-care antibiotic treatment.

Among the **281 patients screened**, **182 were enrolled**. The percentage of patients with recurrence of *C. difficile* infection was 12% in the SER-109 group and 40% in the placebo group.

In patients with symptom resolution of *C. difficile* infection after treatment with standard-of-care antibiotics, **oral administration of SER-109 was superior to placebo in reducing the risk of recurrent infection.**



REZAFUNGIN FOR INVASIVE CANDIDIASIS

- Another new advance is *Rezafungin, a once-weekly* echinocandin for treating invasive candidiasis.
- In a prospective, double-blind, noninferiority phase 3 study assessed weekly *IV Rezafungin* compared with daily *IV Caspofungin* followed by *optional step-down oral fluconazole* in adult patients with candidemia or invasive candidiasis.
- Participants received treatment for a minimum of *14 days*.
- In a phase 3 trial, *weekly Rezafungin was noninferior to daily Caspofungin followed by optional fluconazole regarding 14-day global cure and 30-day all-cause mortality*.
- The availability of Rezafungin as a new antifungal is a valuable addition to candidaemia and invasive candidiasis treatments and may help to address the growing challenge of treatment-resistant *Candida* strains and species.

Efficacy and safety of rezafungin and caspofungin in candidaemia and invasive candidiasis: pooled data from two prospective randomised controlled trials

George R Thompson III, MD   • Alex Soriano, PhD • Patrick M Honore, PhD • Matteo Bassetti, MD • Oliver A Cornely, MD • Marin Kollef, MD • et al. [Show all authors](#)

[Open Access](#) • Published: November 23, 2023 • DOI: [https://doi.org/10.1016/S1473-3099\(23\)00551-0](https://doi.org/10.1016/S1473-3099(23)00551-0)





VaxigripTetra
DO YOU HAVE ASTHMA? YOU'RE AT RISK.
Asthma can prevent you from doing what you love to do. It can even get worse. Don't risk it. Get a flu jab today.

VaxigripTetra
Quadrivalent influenza vaccine (split virus, inactivated)
Supporting your practice's most vulnerable against influenza







MORE GOOD NEWS IN THE TREATMENT OF MULTIDRUG-RESISTANT PULMONARY TUBERCULOSIS

- In a spotlighted study, studies on the efficacy of an ***all-oral bedaquiline-containing regimen*** for treatment of multidrug-resistant tuberculosis.
- The incidence of ***resistant tuberculosis (TB)*** continues to rise worldwide, in part because treatment protocols are prolonged, complex, and painful, all potentially leading to medication failures
- Several small single-nation studies have found that bedaquiline-based regimens appear effective for these infections.
- In a multinational open-label trial, an ***all-oral 9-month bedaquiline-containing regimen and a 6-month bedaquiline regimen with injectable kanamycin were superior to a 9-month control injectable regimen.***

Evaluation of two short standardised regimens for the treatment of rifampicin-resistant tuberculosis (STREAM stage 2): an open-label, multicentre, randomised, non-inferiority trial

Ruth L Goodall, PhD   • Prof Sarah K Meredith, MBBS • Prof Andrew J Nunn, MSc • Adamu Bayissa, MD
Anuj K Bhatnagar, MD • Gay Bronson, MSc • et al. [Show all authors](#) • [Show footnotes](#)

ORAL SIMNOTRELVIR FOR ADULT PATIENTS WITH MILD-TO-MODERATE COVID-19

- *Simnotrelvir* is an oral 3-chymotrypsin-like protease inhibitor that has been found to have in vitro activity against severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) and potential efficacy in a phase 1B.
- In this phase 2–3, double-blind, randomized, placebo-controlled trial, we assigned patients who had mild-to-moderate coronavirus disease 2019 (Covid-19) and onset of symptoms within the past 3 days in a 1:1 ratio to *receive 750 mg of simnotrelvir plus 100 mg of ritonavir or placebo twice daily for 5 days*. The primary efficacy end point was the time to sustained resolution of symptoms, defined as the absence of 11 Covid-19–related symptoms for 2 consecutive days. Safety and changes in viral load were also assessed.
- *Early administration of simnotrelvir plus ritonavir* shortened the time to the resolution of symptoms among adult patients with Covid-19, without evident safety concerns.

Oral Simnotrelvir for Adult Patients with Mild-to-Moderate Covid-19

Bin Cao, M.D., Yeming Wang, M.D., Hongzhou Lu, M.D., Chaolin Huang, M.D., Yumei Yang, M.D., Ph.D., Lianhan Shang, M.D., Zhu Chen, M.D., Rongmeng Jiang, M.D., Yihe Liu, M.D., Ling Lin, M.D., Ping Peng, M.D., Fuxiang Wang, M.D., Fengyun Gong, M.D., Honglin Hu, M.S., Cong Cheng, M.D., Xiangyang Yao, M.D., Xianwei Ye, M.D., Hourong Zhou, M.D., Yinzhong Shen, M.D., Chenfan Liu, M.D., Chunying Wang, M.D., Zhennan Yi, M.D., Bijie Hu, M.D., Jiuyang Xu, M.D., Xiaoying Gu, Ph.D., Jingshan Shen, Ph.D., Yechun Xu, Ph.D., Leike Zhang, Ph.D., Jia Fan, M.D., Renhong Tang, Ph.D., and Chen Wang, M.D.

ABSTRACT

BACKGROUND

Simnotrelvir is an oral 3-chymotrypsin-like protease inhibitor that has been found to have in vitro activity against severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) and potential efficacy in a phase 1B trial.

METHODS

In this phase 2–3, double-blind, randomized, placebo-controlled trial, we assigned patients who had mild-to-moderate coronavirus disease 2019 (Covid-19) and onset of symptoms within the past 3 days in a 1:1 ratio to receive 750 mg of simnotrelvir plus 100 mg of ritonavir or placebo twice daily for 5 days. The primary efficacy end point was the time to sustained resolution of symptoms, defined as the absence of 11 Covid-19–related symptoms for 2 consecutive days. Safety and changes in viral load were also assessed.

RESULTS

A total of 1208 patients were enrolled at 35 sites in China; 603 were assigned to receive simnotrelvir and 605 to receive placebo. Among patients in the modified intention-to-treat population who received the first dose of trial drug or placebo within 72 hours after symptom onset, the time to sustained resolution of Covid-19 symptoms was significantly shorter in the simnotrelvir group than in the placebo group (180.1 hours [95% confidence interval (CI), 162.1 to 201.6] vs. 216.0 hours [95% CI, 203.4 to 228.1]; median difference, –35.8 hours [95% CI, –60.1 to –12.4]; $P=0.006$ by Peto–Prentice test). On day 5, the decrease in viral load from baseline was greater in the simnotrelvir group than in the placebo group (mean difference [\pm SE], -1.51 ± 0.14 log₁₀ copies per milliliter; 95% CI, -1.79 to -1.24). The incidence of adverse events during treatment was higher in the simnotrelvir group than in the placebo group (29.0% vs. 21.6%). Most adverse events were mild or moderate.

CONCLUSIONS

Early administration of simnotrelvir plus ritonavir shortened the time to the resolution of symptoms among adult patients with Covid-19, without evident safety concerns. (Funded by Jiangsu Sincere Pharmaceutical; ClinicalTrials.gov number, NCT05506176.)

DIAGNOSIS AND MANAGEMENT OF DIABETES-RELATED FOOT INFECTIONS

- In a new guideline for diagnosis and management of *diabetes-related foot infections (DFI)* its incidence is increasing in parallel with the rising global prevalence of diabetes and is associated with significant *morbidity and mortality*.
- Recommendations were recently updated by *the International Working Group on the Diabetic Foot/Infectious Diseases Society of America (IWGDF/IDSA)* to guide the management and diagnosis of DFI.

JOURNAL ARTICLE CORRECTED PROOF GUIDELINES

IWGDF/IDSA Guidelines on the Diagnosis and Treatment of Diabetes-related Foot Infections (IWGDF/IDSA 2023)

Éric Senneville, Zaina Albalawi, Suzanne A van Asten, Zulfiqarali G Abbas, Geneve Allison, Javier Aragón-Sánchez, John M Embil, Lawrence A Lavery, Majdi Alhasan, Orhan Oz ...

[Show more](#)

[Author Notes](#)

Clinical Infectious Diseases, ciad527, <https://doi.org/10.1093/cid/ciad527>

Published: 02 October 2023 **Article history** ▼

KEY RECOMMENDATION

- Severity and diagnosis of DFI both depend on local and systemic symptoms.
- Consider culturing tissue aseptically sampled by wound curettage or biopsy.
- If plain x-rays and probe-to-bone testing are inconclusive for suspected osteomyelitis, magnetic resonance imaging should be performed.
- Bone sampling (intraoperative or percutaneous) should be obtained for culture in osteomyelitis cases.
- Antibiotics should be avoided in the absence of signs or symptoms of infection in diabetic foot ulcers.
- For DFI involving skin and soft tissue, treatment duration is typically 1–2 weeks (up to 4 weeks if improvement is slow).

- Empiric treatment should focus on gram-positive bacteria, including *Staphylococcus aureus*.
- Empiric coverage of *Pseudomonas aeruginosa* is suggested for those living in Asia or North Africa.
- In patients with DFI-associated osteomyelitis and amputation with positive bone margins, antibiotics are suggested for 3 weeks; for those patients without amputation, 6 weeks are recommended.
- Surgical management should be considered in patients with moderate to severe DFI.
- Adjunctive therapies (e.g., G-CSF, topical antiseptics, silver, honey, bacteriophages, topical antibiotics, hyperbaric oxygen) are not recommended.

CONCLUSION

As in the past, the breathtaking advances during 2023 highlight the dynamism of the field of infectious diseases while presenting that we are likely to see continued and accelerating progress in the years to come.

فلوشیپ بیماریهای عفونت در نقص ایمنی و پیوند دانشگاه علوم پزشکی شهید بهشتی



برای اولین بار در ایران در سال **۱۳۸۹** دوره تکمیل تخصصی (فلوشیپ) "بیماریهای عفونی در بیماران مبتلا به نقص ایمنی و پیوند"

(Infectious Diseases in Immunocompromised Hosts & Transplantation fellowship)

به همت دکتر مسعود مردانی، مدیر گروه بیماریهای عفونی دانشگاه علوم پزشکی شهید بهشتی، زیر نظر این گروه و با ظرفیت پذیرش ۲ دستیار در بیمارستان طالقانی راه اندازی شد. در حال حاضر ظرفیت پذیرش دستیار این دوره به ۴ نفر افزایش یافته است.

تعداد فارغ التحصیلان و دستیاران در حال تحصیل رشته
فلوشیپ بیماریهای عفونت در نقص ایمنی و پیوند دانشگاه علوم
پزشکی شهید بهشتی: **۳۶ نفر**

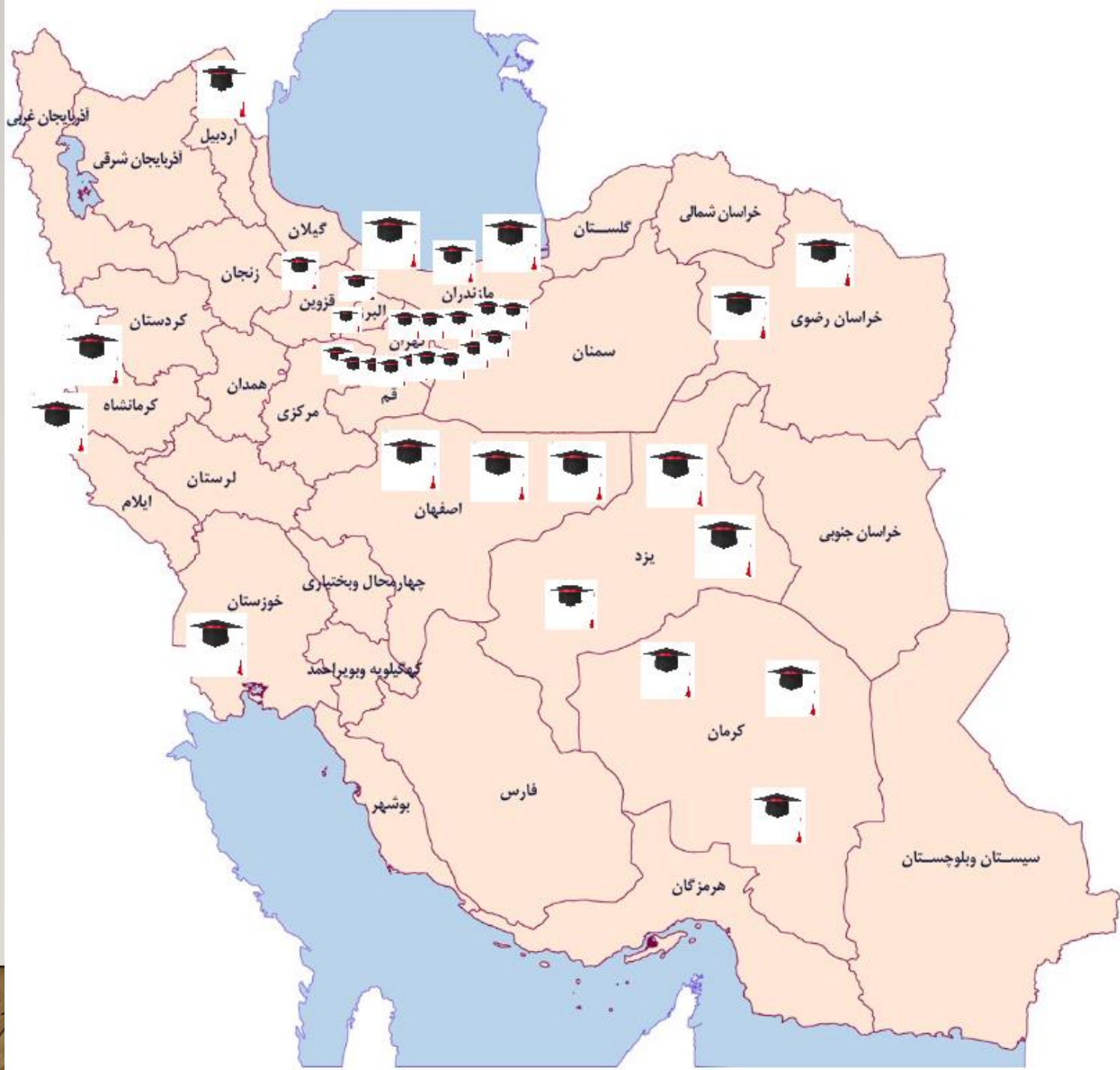
مدیر برنامه: دکتر مسعود مردانی

طول دوره

فلوشیپ بیماریهای عفونی در بیماران مبتلا به نقص ایمنی

و پیوند ۱۸ ماه به شرح جدول ذیل است:

ردیف	نام بخش	مرکز پزشکی، آموزشی، درمانی	مدت رویش (ماه)
۱	آنکولوژی/هماتولوژی	طالقانی	۶
۲	بیماریهای عفونی	لقمان حکیم	۳
۳	روماتولوژی	لقمان حکیم	۱
۴	پیوند کبد	طالقانی	۱
۵	پیوند کلیه	لبافی نژاد	۲
۶	پیوند قلب و ریه و عفونت نقص ایمنی مادرزادی	مسیح دانشوری	۳
۷	رادیوتراپی	شهدای تجریش	۱
۸	اطفال	کودکان مفید	۱



Dear Massoud,

I hope this mail finds you well. I am pleased to inform you that I had a special rate for our Iranian colleagues for registration of the ICBS Symposium. It will be 100 Euro per person irrespective of submitting and abstract or not. Please forward this information your colleagues.

See you in Antalya.

Best regards,

Murat

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IRANIAN

IMMUNOCOMPROMISED

HOST SOCIETY

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THANK YOU