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**Conferral of the Doctorate honoris causa
in Advances in Infectious Diseases,
Microbiology, Legal Medicine
and Public Health Sciences
on Anthony S. Fauci**

Thursday, January 13, 2022 - 3.00 pm
Senate Hall, Rectorate Building

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of the Doctorate honoris causa
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Senate Hall, Rectorate Building
Piazzale Aldo Moro 5, Rome

Programme

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Anthony S. Fauci, M.D., is director of the National Institute of Allergy and Infectious Diseases (NIAID) at the U.S. National Institutes of Health, where he oversees an extensive research portfolio devoted to preventing, diagnosing, and treating infectious and immune-mediated diseases.

He has been a key advisor to seven Presidents and their administrations on global AIDS issues, and on initiatives to bolster medical and public health preparedness against emerging infectious disease threats such as pandemic influenza. As an HIV/AIDS researcher he has been involved in the scientific effort since AIDS was recognized in 1981, conducting pivotal studies that underpin the current understanding of the disease and efforts to develop therapies and tools of prevention.

Anthony S. Fauci was one of the principal architects of the President's Emergency Plan for AIDS Relief (PEPFAR), which has helped save millions of lives throughout the developing world.

Opening Address of the Rector

Antonella Polimeni

Your Excellencies,
dear Colleagues,
Ladies and Gentlemen,

Sapienza University of Rome,
in compliance with the resolution
of the Academic Senate of December 13, 2021,
is honoured to confer today
the Doctorate *honoris causa*
in Advances in Infectious Diseases,
Microbiology, Legal Medicine
and Public Health Sciences on Anthony Fauci
“for his fundamental contribution
in the study of infectious diseases,
with special regard to his pioneer studies
on HIV infections and to the establishment
of effective measures of transmission control
and therapies against AIDS,
and more recently, for his strong
and competent contribution
to fight the SARS-CoV2 pandemic.”

Research on infectious diseases
and their agents has a long tradition
at Sapienza University, spanning
from clinical infectiology
across tropical medicine and public health
to virology, bacteriology and parasitology.
To mention but a few of the scientists
from Sapienza University who have made
fundamental contributions in the area,
I would like to recall Giuseppe Giunchi,
the founder of the Italian school
of modern infectious diseases at Sapienza,
further developed in recent years

by Vincenzo Vullo and co-workers,
Ferdinando Aiuti, a pioneer of HIV research
who greatly contributed also to remove
the stigma of shame and guilt afforded
to AIDS patients, Giuseppe Sanarelli
who provided relevant contributions
to studies on cholera and, together
with Angelo Celli, to the creation
of the well-known malariology school.
In this perspective, we should not forget
Mario Coluzzi, whose outstanding
contribution to studies on malaria vectors
is recognised worldwide.

Italy’s geographical position,
at the Mediterranean border of Europe
and close at the same time to North Africa
and the Middle East, makes our country
naturally sensitive to tropical diseases,
a circumstance witnessed in recent history
when many regions of our country
were heavily affected by transmissible diseases.
At the beginning of the last century,
Italian scientists like Grassi, Celli, Bignami
among others were a reference
for scientists worldwide. At that time,
Italy was a land where infectious diseases
had been endemic for many years.
Only after the Second World War
did large programmes of improvement
of the population’s social and hygienic
conditions allow such threats to be defeated.

Nowadays, Italian scientists are still engaged
in studying tropical diseases,

and many Italian research institutions have been involved in the WHO roadmap to fight Neglected Tropical Diseases; this seems especially important in a situation in which climate change and globalisation are expanding the geographic range of endemic diseases and, in this perspective, Italy is already facing the introduction of alien species potentially involved in disease transmission.

In this context, Sapienza University looks at the eminent figure of Anthony Fauci as an example of excellence and dedication in the field of transmissible diseases. Dr Fauci made an immense contribution to pioneer studies for understanding AIDS and developing new therapies and prevention tools.

Anthony Fauci is widely recognised for delineating the precise mechanisms whereby immunosuppressive agents modulate the human immune response, providing clues for understanding the pathogenesis and treatment of immune-mediated and infectious diseases.

Notably, we should also mention Dr Fauci's key role as scientific advisor to seven U.S. Presidents and their administrations, which had crucial importance in improving public health preparedness against emerging infectious disease threats such as AIDS and, more recently, the COVID-19 pandemic. In this regard, I would like to especially stress the worldwide impact and contribution of Anthony Fauci in the past two dramatic years: his firm and, at the same time, simple and clear communication

has made him one of the most listened to scientific voices, not only by ordinary people around the world but also by policymakers, helping them follow the rational way of the scientific method, even when shortcuts may have been more accessible and more appealing.

For all the above reasons, today we are conferring the Doctorate *honoris causa* in Advances in Infectious Diseases, Microbiology, Legal Medicine and Public Health Sciences on Anthony Fauci, true to our University's tradition of excellence, inclusion, internationalisation and cooperation.

Address by the Dean of the Faculty of Pharmacy and Medicine Carlo Della Rocca

Magnificent Rector,
distinguished Colleagues,
dear Students, Ladies and Gentlemen,

the members of the Faculty of Pharmacy and Medicine are delighted to approve the Doctorate *honoris causa* in Advances in Infectious Diseases, Microbiology, Legal Medicine and Public Health Sciences to Dr. Anthony Fauci, proposed by the Department of Public Health and Infectious Diseases.

The Rector has just briefly outlined the extraordinary scientific achievements of Anthony Fauci and she underlined how he served the USA and global community during the dramatic emergencies of AIDS and Covid-19 pandemic, providing an invaluable scientific and moral compass. Therefore, what I would like to do here is to recall his career and scientific achievements.

Anthony Fauci attended Cornell University's Medical College graduating first in his class with a Doctor of Medicine degree in 1966. He then did an internship and residency in internal medicine at New York Hospital-Cornell Medical Center

Then Dr. Fauci joined the NIH in 1968 as a clinical associate in the Laboratory of Clinical Investigation (LCI) at the National Institute of Allergy and Infectious Diseases (NIAID). He became Head of the Clinical Physiology Section, LCI, and was later appointed Chief of the Laboratory of Immunoregulation. Since 1984 he is Director of the NIAID.

Anthony Fauci has carried out pioneering studies that provided the basis for understanding the regulation of the human immune response. Moreover, he unraveled the mechanisms whereby immunosuppressive agents modulate the human immune response. He applied his deep understanding of these fundamental mechanisms for the development of therapies for formerly fatal inflammatory and immune-mediated diseases such as polyarteritis nodosa, Wegener's granulomatosis, and lymphomatoid granulomatosis that in 1985 were declared as "one of the most important advances in patient management in rheumatology over the previous 20 years."

In 1981 Fauci understood the significance of a new, mysterious disease and decided

to make it the focus of his research, despite contrary advice from his mentor and colleagues. His work helped change the course of the epidemic and in 1984, at the age of 43, he was named director of the NIH National Institute of Allergy and Infectious Diseases (NIAID).

That made him the youngest institute director ever at NIH. Under Fauci's leadership, NIAID became the single largest funder of HIV/AIDS research in the world.

His own lab's research also has helped clarify fundamental relationships between the virus and the immune system.

Anthony Fauci has made seminal contribution to the understanding of how the HIV virus disarms the body's immune defenses by targeting and depleting CD4 cells, thus leading to its susceptibility to deadly infections, the AIDS.

He also has delineated the mechanisms of induction of HIV expression by endogenous cytokines. Furthermore, he has been instrumental in developing highly effective strategies for the therapy of patients with this deadly disease, as well as for a vaccine to prevent HIV infection. He continues to devote much of his research efforts to identifying the nature of the immunopathogenic mechanisms of HIV infection and the scope of the body's immune responses to the AIDS retrovirus.

In 2003, an Institute for Scientific Information study indicated that in the twenty year period from 1983 to 2002, Anthony Fauci was the 13th most-cited scientist among the 2.5 to 3 million authors in all disciplines throughout the world who published articles

in scientific journals during that time frame. Dr. Fauci was the world's 10th most-cited HIV/AIDS researcher in the period 1996-2006.

Research and clinical achievements make Anthony Fauci a prominent figure in medicine, across two centuries. But he also served as head of NIH National Institute of Allergy and Infectious Diseases and as scientific advisor to seven US presidents and their administration, giving a contribution to shaping national and global policies with the integrity and principles of the best practice in science and medicine.

Anthony Fauci is an outstanding figure for scientific excellence and for commitment to medicine and principles of integrity and dedication to health for all human beings, on behalf of the entire scientific community, we are gathered here to express our gratitude for Anthony Fauci's contributions and we congratulate him on receiving the Doctorate *honoris causa* in Advances in Infectious Diseases, Microbiology, Legal Medicine and Public Health Sciences from Sapienza University.

Praise of Anthony S. Fauci

Stefano D'Amelio

Magnificent Rector,
distinguished Colleagues,
dear Students, Ladies and Gentlemen,
dear Dr. Fauci,

it is with great pleasure and pride that, as coordinator of the PhD course in Advances in Infectious Diseases, Microbiology Legal Medicine and Public Health Sciences, I have been asked to hold a praise on the occasion of the Doctorate *honoris causa* awarded to Anthony Fauci.

At its beginning, this initiative was shared with Professor Beatrice Vallone, director of the H2CU, the Honors Center of Italian Universities, an institution especially devoted to promote the scientific and academic collaboration between Italy and USA.

Our PhD course is largely interdisciplinary, spanning from infectious diseases to microbiology and parasitology, to public health, medical statistics and epidemiology, up to forensic medicine.

In the vast majority of the research and academic fields covered by our PhD course, Anthony Fauci represents an extraordinary example of scientific excellence and a worldwide recognized global expert. His scientific production is terrific, counting up to 1171 scientific papers in Scopus, with an H-index of 183.

He has been a pioneer in immunomodulation, providing fundamental contributions to the study on the complex mechanisms that contributes to define the human immune response.

This allowed relevant advances in developing treatment strategies of inflammatory and immune-mediated diseases such as polyarteritis nodosa, granulomatosis with polyangiitis (formerly Wegener's granulomatosis), and lymphomatoid granulomatosis.

In the late seventies, Anthony Fauci carried out important studies on immune regulation in infections caused by *Mycobacterium tuberculosis*. He never neglected this disease, and recently he claimed for the need of measures for the intensification of public health efforts, including improving the current diagnostic, preventive, and therapeutic tools.

Dr. Fauci has given a fundamental contribution to the understanding of how HIV destroys the immune defenses leading to its susceptibility to deadly infections. As director of NIAID, he has a longstanding commitment to discover new drugs to control HIV and AIDS-associated complications and co-infections. His studies have permitted the development of anti-HIV treatments that have radically increased life expectancy

and improved the living conditions of HIV-infected people throughout the world. As an immunologist, he contributed with fundamental findings on HIV's pathogenesis. At the same time, he was the key architect of the US President's Emergency Plan for AIDS Relief that provides anti-HIV treatments to people from the poorest countries. Dr. Fauci published over 400 papers on HIV and the most recent publications focused on the development of the long-awaited vaccine, shedding a light of hope for the future.

In the course of Ebola epidemics, along to relevant papers on the treatment of the diseases, Anthony Fauci has treated Ebola patients himself, and I want to report the following highly moral statements using his own words: "I do believe that one gets unique insights into disease when you actually physically interact with patients." Moreover, he also wanted to show his staff that he wouldn't ask them to do anything he wouldn't do himself. This is a strong moral and teaching legacy.

Dr. Fauci has been deeply involved in programs aimed to reduce the malaria burden in endemic countries. In particular, he has claimed the need for the availability of the genomic sequence data for each of the three organisms involved in malaria transmission, with the aim to clarify the pathophysiologic relationships among human host, parasite, and vector.

Another important area of research that Anthony Fauci has stimulated at his Institute regards Neglected

Tropical Diseases, by supporting and stimulating programs on vector biology, filariasis, schistosomiasis and by funding eight Tropical Research Medicine Centers to support studies on NTDs in endemic areas.

In the last two years characterized by the COVID-19 pandemics, Dr. Fauci has represented for the United States citizens and for all of us a firm point of reference both for his scientific authority and for his honesty and clarity in communicating the severity of the disease to the large public and to policymakers. He advocated the three-ways approach for the control of the pandemics, represented by prevention, vaccination and therapies. In this context, Dr. Fauci recently reviewed the efficacy of several therapeutic strategies, identifying pitfalls in approaches that failed to demonstrate efficacy following validation based on randomized controlled clinical trials. He always underlined the need of a data-driven approach to evaluate eventual benefits in treating hospitalized COVID-19 patients.

His ability and dedication to the research in all these fields has been always characterized by a "bench to bed" approach, as he understood the necessary connection between basic and applied science, and on how science represents not only a value for the universal knowledge but also an essential tool for the improvement of humanity, with a strong and passionate eye to the patient well-being. Nevertheless, he recently claimed that mRNA-based vaccines against COVID-19, which marked a critical milestone in reducing the impact of the pandemic, are the fruits of years of basic research.

We all know that the strong links of Anthony Fauci with our country have their roots also on his Italian origin, in part from Sicily and for the other side from Naples and Irpinia. But the connections went over in all these years, as he has mentored a large number of Italian researchers, now spread in numerous laboratories in the world.

With the conferring of the Doctorate *honoris causa* to Anthony Fauci, we want not only to renew this connection and to acknowledge his universal contribution in the field of infectious disease, but also recognize an example that will stimulate our PhD students to pursue their scientific objectives with dedication, commitment and creativity, in the belief that basic or applied research will make the world a better place to live.

Stefano D'Amelio
Professor in Parasitology
and Parasitic Diseases of Animals

Lectio magistralis

Anthony S. Fauci

COVID-19: Lessons Learned and Remaining Challenges

Summary

Just over two years ago, a novel coronavirus causing severe pneumonia was first recognized in Wuhan, China. Later named SARS-CoV-2, this virus has since spread throughout the world and has caused several hundred million cases of coronavirus disease 2019 (COVID-19) and more than five million deaths. The COVID-19 pandemic has been characterized by the emergence and spread of SARS-CoV-2 variants, including the highly transmissible and severe delta variant and the highly transmissible omicron variant now predominant in many parts of the world. Early data suggest that the disease severity of COVID-19 caused by infection with the omicron variant is lower than with previous variants. However, the increased transmissibility and immune evasion of omicron may override any impact lower disease severity may have on the burden of hospitalizations and death, particularly among unvaccinated individuals at highest risk.

Infection with SARS-CoV-2 is associated with a wide spectrum of a disease, from asymptomatic infections to severe pneumonia and organ failure.

Numerous non-pulmonary complications of COVID-19 have been reported, including neurologic disorders, hyperinflammation, cardiac dysfunction, hypercoagulability, acute kidney injury, and multisystem inflammatory syndrome in children (MIS-C). Risk factors for severe COVID-19 illness include being older than 65 years, and having underlying conditions such as obesity, hypertension, diabetes, chronic lung disease, or immunosuppression.

The public health response to COVID-19 has required an unprecedented public-private research effort. Clinical trials launched very early in the pandemic demonstrated the benefit of several repurposed therapeutics – including remdesivir, corticosteroids, and other immunomodulators – in mitigating severe outcomes. The development of monoclonal antibody therapies, delivered as infusions, later proved highly effective at preventing progression to advanced disease if given early in infection. However, some of these products have little or no effectiveness against the omicron variant. Recently, two effective oral drugs (Paxlovid and Molnupirivir) have been authorized for outpatient use in individuals with SARS-CoV-2 at high risk for progression to severe disease. Research programs have been launched

to stimulate the discovery and development of additional antivirals for COVID-19 and future viral pandemics.

The notable extraordinary scientific success story of the pandemic has been vaccines. Sustained research investments prior to the emergence of SARS-CoV-2 and the formation of robust public-private partnerships enabled the rapid pace of COVID-19 vaccine development. The Pfizer-BioNTech and Moderna mRNA vaccines and the Johnson & Johnson adenovirus-vectored vaccine now are authorized or approved for use in the United States and many other countries. Other countries including the UK, Russia, China, India among others have developed effective vaccines with varying degrees of effectiveness. With regard to the vaccines utilized in the USA, real-world effectiveness studies have shown that these vaccines greatly reduce symptomatic infection, severe disease, and death from COVID-19. However, protection (particularly against symptomatic infection) wanes over time, especially with the immune-evasive omicron variant. A booster (third mRNA or second adenovirus) dose has been shown to restore high levels of protection, even against omicron. Looking to the future, the development of broadly protective coronavirus vaccines that are impervious to viral variants is a major research priority.

As we enter the third year of the COVID-19 pandemic, we face significant challenges

with the surging omicron variant and perhaps other variants to follow. Fortunately, we have the tools we need to bring the SARS-CoV-2 virus under control. Using these tools, particularly vaccines and therapies, in conjunction with proven public health measures such as masking, ventilation, and social distancing we can help subdue outbreaks. Vaccination, including booster doses, can turn a potentially deadly pneumonia into a mild respiratory illness. Rapid antigen and molecular diagnostic tests can identify infected individuals to break the chain of transmission and enable isolation and early treatment. Therapeutics can prevent progression to severe disease for those at high risk. In 2022 and beyond, we will apply what we have learned during the first two years of the COVID-19 response to prepare for future SARS-CoV-2 variants and other viral pathogens with pandemic potential.

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