

MEETING

LO STATO DELL'ARTE DEL PNRR: TELEMEDICINA E FORMAZIONE D'ECCELLENZA

VENERDÌ **21 FEBBRAIO** 2025
ore **9:00 - 13:00**

Aula Magna "Aldo Cossu"
Università degli Studi di Bari Aldo Moro
Via Scipione Crisanzio, 1 - Bari



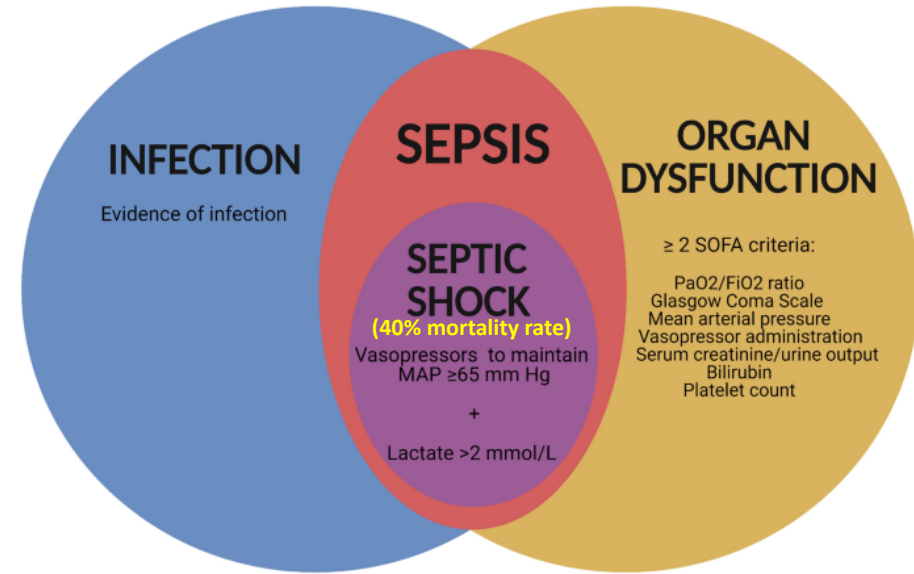
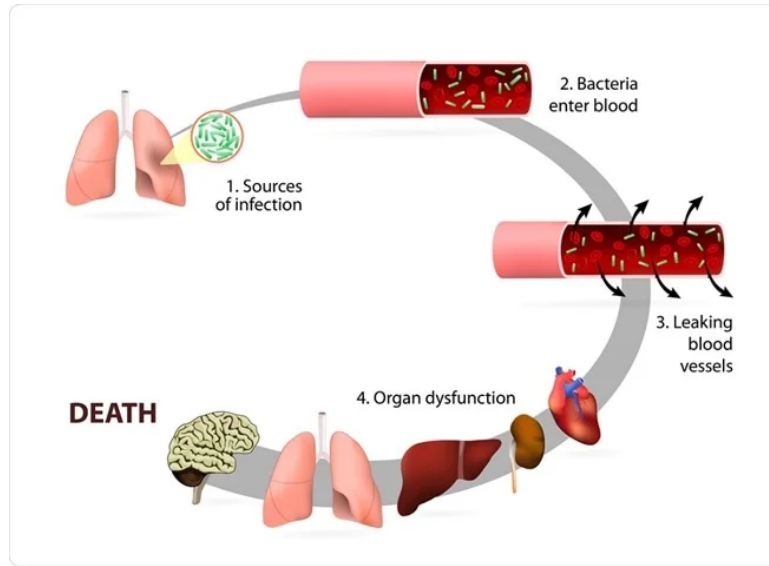
GIUSEPPE CALAMITA

**AQUAPORINA-9 E INTELLIGENZA
ARTIFICIALE NELLA PREDIZIONE
PROGNOSTICA DELLA SEPSI**

Sepsis, one of the oldest and most ambiguous syndromes in medicine

Sepsis

When the immune system becomes overactive in response to an infection, causing inflammation that can affect the functioning of other tissues and organs



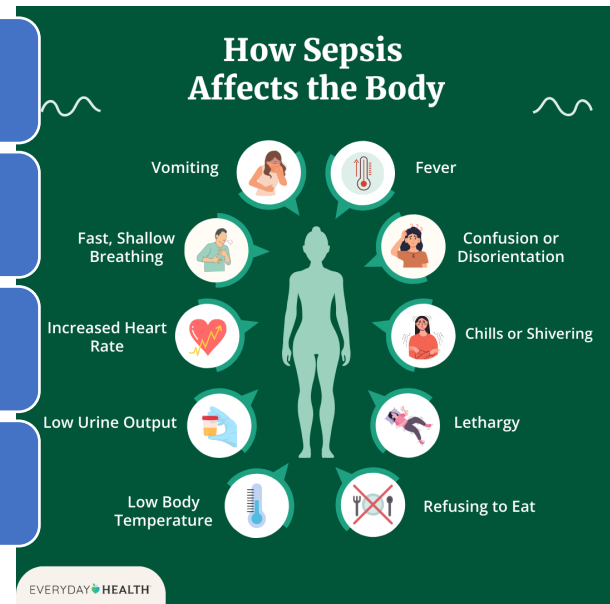
Sepsis, a silent killer and leading cause of death

40-70 million cases and 11 million deaths per year in the world (associated with one in five deaths on the planet)

Recognized as an acute inflammatory condition with the activation of the innate immune system triggered by infection

Main etiological causes: bacterial, fungal and viral infections

Different categories of subjects at risk



➤ It **develops quickly**, and it can be **difficult to identify in its early stage**

- Diagnostic and prognostic predictions are fundamental
- Need of novel drug targets



Scoring systems for the characterization of sepsis and associated outcomes

SOFA score

Sequential Organ Failure Assessment (SOFA)

- Performance of six organ systems in the body:
- Neurologic
- Blood
- Liver
- Kidney
- Blood pressure
- Blood hemodynamics

qSOFA score

Quick SOFA (qSOFA)

Three items:

- respiratory rate ≥ 22 breaths per minute
- altered mentation [GCS] < 15
- systolic blood pressure (SBP) < 100 mmHg

MEWS score

Modified Early Warning Score (MEWS)

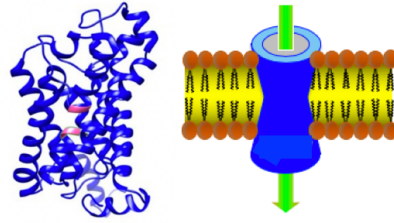
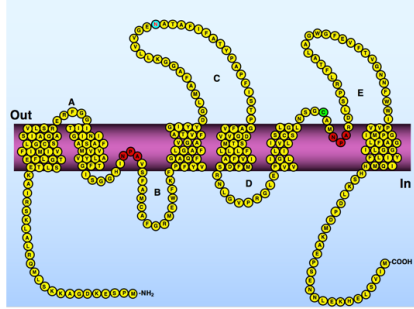
Four items:

- respiratory rate > 22 breaths per minute
- state of consciousness
- oxygen saturation
- heart rate

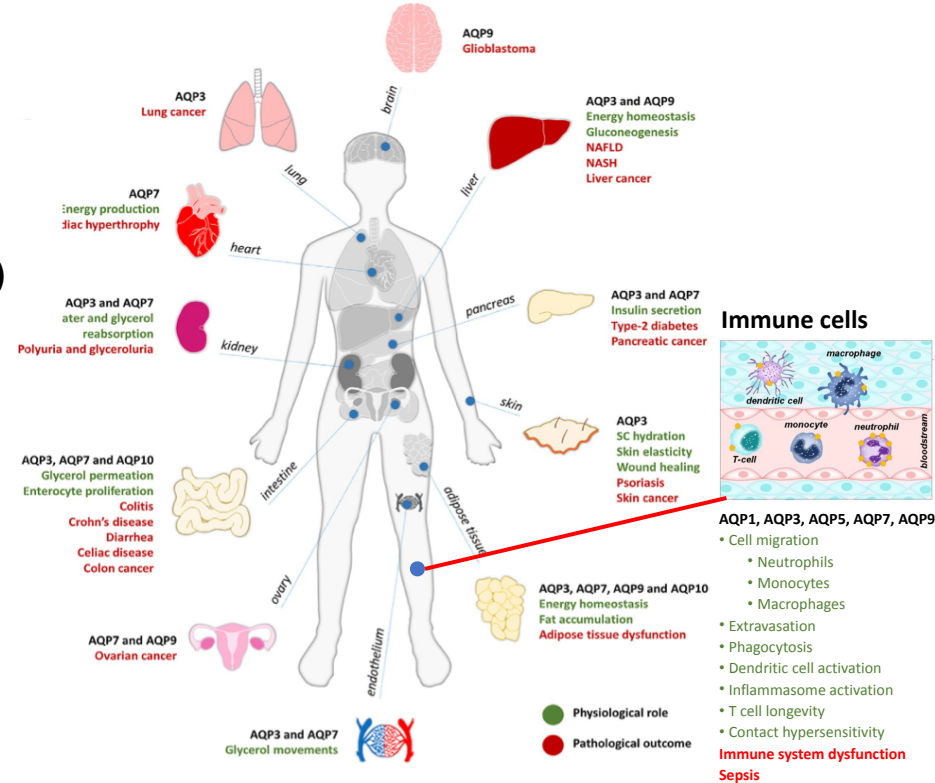
Groundbreaking advances in personalizing treatment of sepsis **are needed** to save lives

- Inclusion of state-of-the art **biomarkers**, replacement of time-consuming blood cultures (days) by a **rapid (hours) test kit** and **AI-supported methods of diagnostic and prognostic prediction**

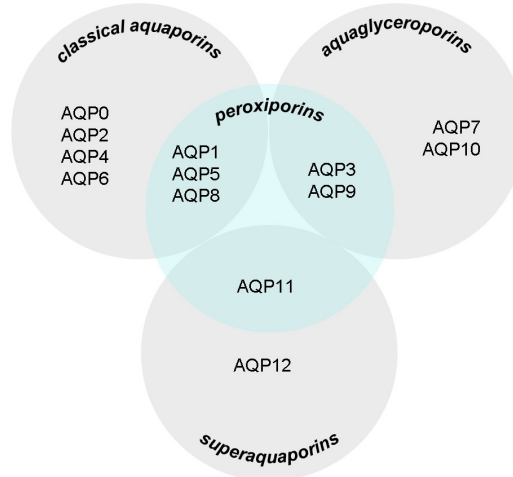
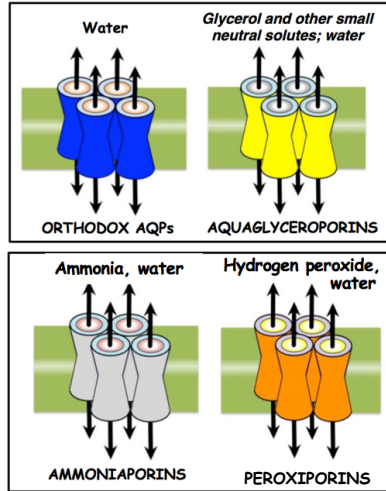
The Aquaporin family of membrane channels



Human Aquaporins in health and disease



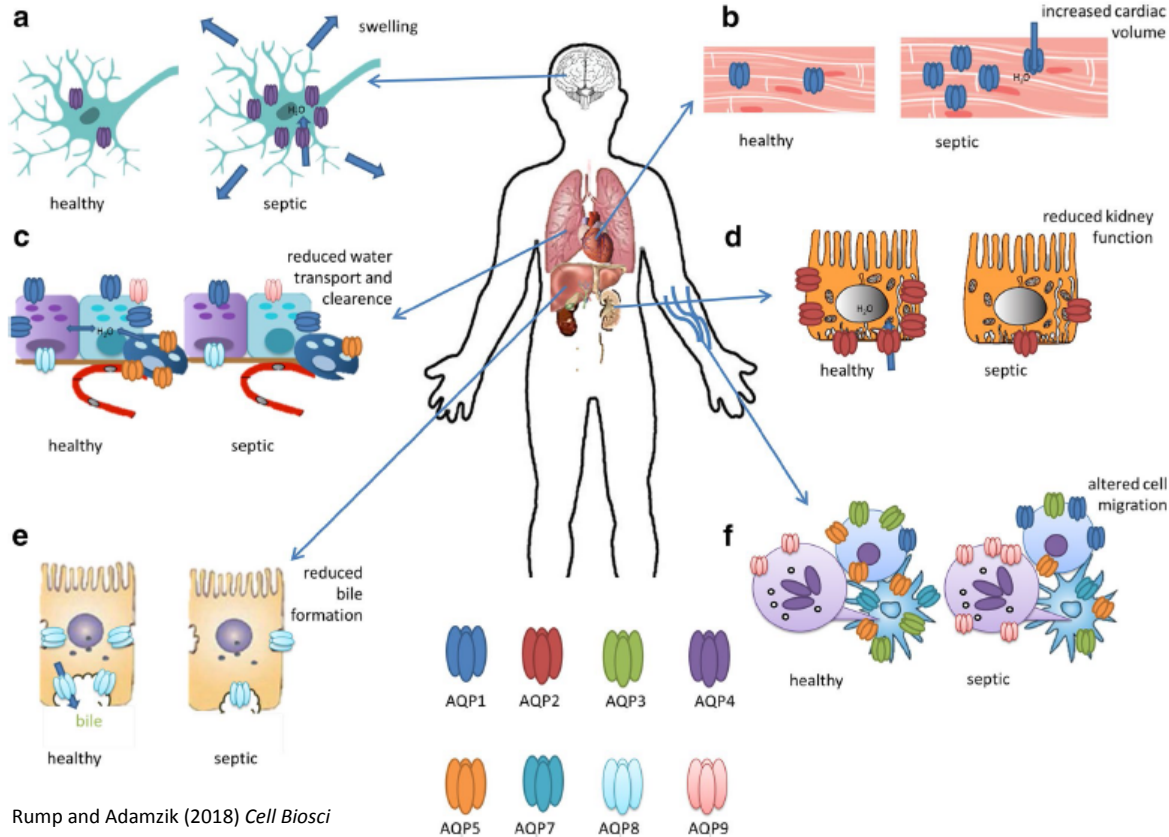
Humans have 13 AQPs (AQP0-AQP12)



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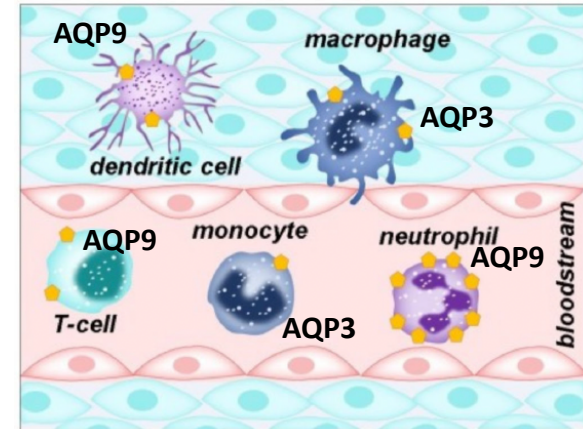
Lo stato dell'arte del PNRR: telemedicina e formazione d'eccellenza
Università degli Studi di Bari – 21 Febbraio 2025

Expression and proposed roles of AQPs in sepsis



Rump and Adamzik (2018) *Cell Biosci*

AQP3 and AQP9 in immune cells



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Lo stato dell'arte del PNRR: telemedicina e formazione d'eccellenza
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AQP3 and AQP9 have roles in the immune system

The Rockefeller University Press \$30.00
J. Exp. Med. 2012 Vol. 209 No. 10 1743–1752
www.jem.org/cgi/doi/10.1084/jem.20112398

Chemokine-dependent T cell migration requires aquaporin-3–mediated hydrogen peroxide uptake

Mariko Hara-Chikuma,^{1,2,4} Shunsuke Chikuma,³ Yoshinori Sugiyama,⁴ Kenji Kabashima,¹ Alan S. Verkman,^{5,6} Shintaro Inoue,⁴ and Yoshiki Miyachi¹

Cellular and Molecular Life Sciences
<https://doi.org/10.1007/s00018-020-03708-3>

Cellular and Molecular Life Sciences

ORIGINAL ARTICLE



Aquaporin-3 is involved in NLRP3-inflammasome activation contributing to the setting of inflammatory response

Inês Vieira da Silva^{1,2} · Carlos Cardoso^{2,3} · Helios Martínez-Banaclocha⁴ · Angela Casini⁵ · Pablo Pelegrin⁴ · Graça Soveral^{1,2}

 **frontiers**
in Microbiology

ORIGINAL RESEARCH
published: 03 September 2015
doi: 10.3389/fmicb.2015.00915

Pseudomonas aeruginosa lasI/rhlI quorum sensing genes promote phagocytosis and aquaporin 9 redistribution to the leading and trailing regions in macrophages

Angelika Holm, Thommie Karlsson and Elena Vikström*

Int. J. Mol. Sci. 2024, 25, 1209. <https://doi.org/10.3390/ijms25021209>



International Journal of
Molecular Sciences



Article

AQP3 and AQP9—Contrary Players in Sepsis?

Patrick Thon¹, Tim Rahmel¹, Dominik Ziehe¹, Lars Palmowski¹, Britta Marko¹, Hartmuth Nowak^{1,2}, Alexander Wolf¹, Andrea Witowski¹, Jennifer Orłowski¹, Björn Ellger³, Frank Wappler⁴, Elke Schwier⁵, Dietrich Henzler⁵, Thomas Köhler⁵, Alexander Zarbock⁶, Stefan Felix Ehrentauf⁷, Christian Putensen⁷, Ulrich Hermann Frey⁸, Moritz Anft⁹, Nina Babel⁹, Barbara Sitek¹, Michael Adamzik¹, Lars Bergmann¹, Matthias Unterberg¹, Björn Koos¹ and Katharina Rump^{1,*} on behalf of the SepsisDataNet.NRW Research Group

Our studies on AQP9 in inflammation and sepsis



Aquaporin-9 Contributes to the Maturation Process and Inflammatory Cytokine Secretion of Murine Dendritic Cells

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OPEN ACCESS

Stefania De Santis^{1,2,3†}, Grazia Serino^{1†}, Maria R. Fiorentino⁴, Vanessa Galleggiante¹, Patrizia Gena⁵, Giulio Verna¹, Marina Liso¹, Monica Massaro¹, Jinggang Lan⁴, Jacopo Troisi^{6,7}, Iliaria Cataldo⁵, Alessia Bertamino², Aldo Pinto², Pietro Campiglia^{2,8}, Angelo Santino⁹, Gianluigi Giannelli¹, Alessio Fasano^{4,7}, Giuseppe Calamita^{5†} and Marcello Chieppa^{1,2†}

Frontiers in Immunology | www.frontiersin.org | June 2022 | Volume 13 | Article 900906

RG100204, A Novel Aquaporin-9 Inhibitor, Reduces Septic Cardiomyopathy and Multiple Organ Failure in Murine Sepsis

Shireen Mohammad^{1*}, Caroline E. O'Riordan¹, Chiara Verra¹, Eleonora Aimaretti², Gustavo Ferreira Alves³, Klaus Dreisch⁴, Johan Evenäs⁴, Patrizia Gena⁵, Angela Tesse⁶, Michael Rützler^{7,8}, Massimo Collino³, Giuseppe Calamita⁵ and Christoph Thiemermann^{1*}

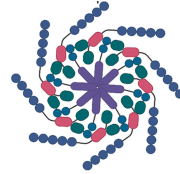
Cells 2021, 10, 435. <https://doi.org/10.3390/cells10020435>



Article

Ablation of Aquaporin-9 Ameliorates the Systemic Inflammatory Response of LPS-Induced Endotoxic Shock in Mouse

Angela Tesse^{1*}, Patrizia Gena², Michael Rützler^{3,4} and Giuseppe Calamita^{2*}



International Immunopharmacology 137 (2024) 112450

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journal homepage: www.elsevier.com/locate/intimp



Inhibition of aquaporin-9 ameliorates severe acute pancreatitis and associated lung injury by NLRP3 and Nrf2/HO-1 pathways

Jiawei Chen^{6,1}, Xiangdong Zhu^{6,1}, Ziqiong Wang⁸, Michael Rützler⁶, Qiaohong Lu⁸, Hongjie Xu⁸, Roland Andersson⁶, Yinwei Dai⁶, Zouwen Shen⁸, Giuseppe Calamita⁴, Shangjing Xie⁶, Yongheng Bai⁶, Bicheng Chen⁶

Cells 2022, 11, 3118. <https://doi.org/10.3390/cells11193118>



Article

Characterization of the Aquaporin-9 Inhibitor RG100204 In Vitro and in db/db Mice

Marilina Florio^{1,†}, Angelica Engfors^{2,†}, Patrizia Gena¹, Jessica Larsson³, Alessandro Massaro^{4,5}, Stella Timpka³, Martina Kvist Reimer³, Per Kjellbom², Eric Beitz⁶, Urban Johanson², Michael Rützler^{2,7,*} and Giuseppe Calamita^{1,*}

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