Professor Vincenzo Nicola TALESA

Department of -Medicine and Surgery University of Perugia, Perugia, Italy

Biography: Vincenzo Nicola Talesa was born in Vibo Valentia (VV), Italy, on 8 December 1959. He is internationally recognized in the area of toxicology, cancer and neurodegenerative research fields in which his major interest is on the comprehension of the mechanisms that lead to the genesis and progression of the related diseases in the bio-medical area. Vincenzo Talesa has published more than 120 publications in high-ranked scientific journals. He has been involved in the EU projects "Multidisciplinary approach to structure and functions of Cholinesterases", "New Biosensors for improved detection of environmental contamination by anticholinesterase pesticides" and "Development of targeted nutrition for prevention of undernutrition for older adults". Reviewer of international journals with impact factor. Scientific leader of the research group "Glyoxalases inhibition and expression *in vivo* and *in vitro* in the control of tumor growth". Organizer of the International Meetings "New Biosensors for improved detection of environmental contamination by anticholinesterase pesticides", Perugia, Italy, and VIIIth International Meeting on Cholinesterases, Perugia, Italy. Organizer of two AIBG Meetings, in 2001 and 2012.

Curriculum Vitae:

- -1982: Graduated with honors in Biological Sciences from the University of Perugia (Perugia, Italy)
- -1985-1986: Visiting Scientist at the Department of Medical Chemistry, University of Helsinki (Finland);
- -1989: Visiting Research at North Texas State University, Denton, Texas (USA);
- -1993-1994: Research Assistant at the Laboratoire de Differenciation cellulaire et Croissance, INRA, Montpellier (France) under the project "Multidisciplinary approach to structure and functions of cholinesterases" funded by the European Economic Community (Human Capital and Mobility);
- -1998: Research Assistant at the Laboratoire de Differenciation cellulaire et Croissance INRA, Montpellier (France) under the project "Galileo."
- -1999–2004 Associate Professor of Applied Biology, University of Perugia Medical School (Perugia, Italy)
- -2004-2020 Full Professor of Applied Biology, University of Perugia Medical School (Perugia, Italy)

COORDINATION OF RESEARCH ACTIVITY

- Biological, biochemical and molecular characterization of Acetylcholinesterases (AChEs) in a) human diseases and b) mechanisms involved in insecticide resistance.
- a) Since the discovery of the cholinergic deficit in Alzheimer disease (AD), acetylcholinesterase (AChE) has been widely investigated in tissues involved in the disease. These studies showed modifications in AChE activity and changes in its polymorphism in brain as well as in cerebro-spinal fluid (CSF) and blood. The co-localization of the enzyme in the senile plaque provided evidence of its anomalous features. It has been also shown that AChE forms a stable complex with senile plaque components through its peripheral anionic site. Moreover, the neurotoxicity of amyloid components is increased by the presence of AChE. The occurrence of an altered glycosylation of some AChE forms in AD is closely related to the presence of amyloid formations. In this ambit, Vincenzo Talesa studied the effect of different AChE inhibitors as potential agents in the treatment of AD.
- b) Cholinesterase (ChE) is one of the most employed biomarkers in environmental analysis. Among ChEs, potentially the most significant in environmental terms is AChE, the enzyme that terminates the nerve impulse. Because of its physiological role, AChE has long been considered a highly specific biomarker for organisms exposed to anticholinesterases agents, primarily agro-chemicals (organophosphate and carbamate pesticides). The effects of these pesticides depends upon their selective inhibition of AChE. In this ambit, Vincenzo Talesa performed the kinetic and molecular characterization, particularly focusing on the kinetic and inhibition of this enzyme in different phyla of Vertebrates and Invertebrates. Of particular relevance is the identification (molecular cloning and expression) of a full-length cDNA encoding a new AChE resistant to organophosphate.
- Glyoxalases inhibition and expression, in vivo and in vitro, in the control of tumor growth.

Glyoxalase 1 (Glo1), together with glyoxalase 2 (Glo2), forms the main scavenging system of methylglyoxal, a potent pro-apoptotic agent mainly generated by glycolysis. An increased rate of glycolysis is a well-known signature of cancer cells. As a survival strategy, Glyoxalases are overexpressed in many human malignant cells. Hence, targeting

these enzymes represents a strategy to selectively inhibit cancer. In this ambit, the research group directed by Vincenzo Talesa identified some major molecular mechanisms based on glyoxalase system and the related glycative and oxidative stress in the genesis, progression and response to therapy in prostate and breast cancers. In the same malignancies, studies have been performed in order to evaluate the association of a specific Glo1 SNP (loss-of-function mutation) with the risk of genesis, progression and survival of tumor-bearing patients. A similar study was performed in the autoimmune disease, multiple sclerosis, and in cerebral cavernous malformations (CCM), a major cerebrovascular disease of proven genetic origin.

- In Vitro study of the role of Natriuretic peptides (NPs) in inflammation and immunity

NPs (ANP and BNP) are hormone/paracrine/autocrine factors released by the heart in response to myocardial stretch and overload, modulating body fluid homeostasis. However, a role of NPs in inflammation and immunity is emerging. Interleukin- 1β (IL- 1β) is a potent pro-inflammatory cytokine involved in a wide range of biological responses, including the immunological one. Unlike other cytokines, IL- 1β production is rigorously controlled. Primarily, NF-kB activation is required to produce pro-IL- 1β ; subsequently, NALP3 inflammasome/caspase-1 activation is required to cleave pro-IL- 1β into the active secreted protein. NALP3 is a molecular platform capable of sensing a large variety of signals and a major player in innate immune defense. Due to their pleiotropism, IL- 1β and NALP3 dysregulation is a common feature of a wide range of diseases. Identifying molecules regulating IL- 1β /NALP3/caspase-1 expression is an important step in the development of new potential therapeutic agents. In this ambit, the research group directed by Vincenzo Talesa provided new evidence of the direct involvement of NPs on NF-kB/NALP3/caspase-1-mediated IL- 1β release and NF-kB-mediated pro-IL- 1β production. We suggest a possible employment of NPs for the treatment of inflammatory/immune-related diseases and IL- 1β /NALP3-associated disorders, affecting millions of people worldwide.

TEACHING ACTIVITY

- -Professor of Cell Biology and Genetics in the the MD Program at Perugia University Medical School
- -Professor of Cell Biology and Genetics in the Biotechnology Degree Program
- -Professor of Cell Biology and Genetics in the Nursing Degree at Perugia University Medical School
- -Professor of Cell Biology and Genetics in Postgraduate Specializations of Medical School

COORDINATION OF TEACHING ACTIVITY

- -2002-2010 Director of the nursing Degree Program at the University of Perugia
- -Director of the University Master in Planning, management and evaluation of integrated health promotion actions for the community

MANAGEMENT AND ADMINISTRATION

- -Vice Dean of Faculty of Medicine
- -Deputy Rector for Teaching Division, University of Perugia
- -2011-2013 Member of the Board of Governors, University of Perugia
- -Board member and Vice-coordinator of the "Evaluation team" of the University of Perugia;
- -Member of the Academic Board of the PhD in Systems Biology in Immune and Infectious Pathologies;
- -Board member of the Genomic Center
- Director and Vice- Director of the Center for Advanced Medical Simulation
- -2014-2019 Director of the Department of Experimental Medicine, University of Perugia (Perugia, Italy)
- -Chief of the Library of the Faculty of Medicine
- -Member of the University Senate, University of Perugia (20014-2019);
- --2020 Member of the Board of Governors, University of Perugia
- -2020-2025, Director of the Department of Medicine and Surgery, University of Perugia (Perugia, Italy)

List of of publications in recent years

PTEN/PKM2/ERα-Driven Glyoxalase 1 Overexpression Sustains PC3 Prostate Cancer Cell Growth Through MG-H1/RAGE Pathway Desensitization Leading to H_2O_2 -Dependent KRIT1 Downregulation.

Manfredelli D, Torcoli C, Pariano M, Bellezza G, Baroni T, **Talesa VN**, Sidoni A, Antognelli C.Antioxidants (Basel). 2025 Sep 15;14(9):1120. doi: 10.3390/antiox14091120.

Natriuretic peptides as novel regulators of dendritic cells-mediated inflammation.

Manni G, Barcelos ECS, Ricciuti D, Pieroni B, Gargaro M, Mencarelli G, Acha-Orbea H, **Talesa VN**, Mezzasoma L, Fallarino F.Cell Mol Life Sci. 2025 Jul 11;82(1):273. doi: 10.1007/s00018-025-05769-8.

Acetylcholine Sustains LNCaP Prostate Cancer Cell Migration, Invasion and Proliferation Through Glyoxalase 1/MG-H1 Axis with the Involvement of Osteopontin.

Manfredelli D, Armeni T, de Bari L, Scirè A, **Talesa VN**, Antognelli C, Pariano M.Int J Mol Sci. 2025 Apr 25;26(9):4107. doi: 10.3390/ijms26094107.

Amniotic fluid stem cell-derived extracellular vesicles educate type 2 conventional dendritic cells to rescue autoimmune disorders in a multiple sclerosis mouse model.

Manni G, Gargaro M, Ricciuti D, Fontana S, Padiglioni E, Cipolloni M, Mazza T, Rosati J, di Veroli A, Mencarelli G, Pieroni B, Silva Barcelos EC, Scalisi G, Sarnari F, di Michele A, Pascucci L, de Franco F, Zelante T, Antognelli C, Cruciani G, Talesa VN, Romani R, Fallarino F.J Extracell Vesicles. 2024 Jun;13(6):e12446. doi: 10.1002/jev2.12446.PMID: 38844736

SARS-CoV-2 Infection is Associated with Age- and Gender-Specific Changes in the Nasopharyngeal Microbiome.

Bozza S, Nunzi E, Frias-Mazuecos A, Pieraccini G, Pariano M, Renga G, Mencacci A, **Talesa VN**, Antognelli C, Puccetti P, Romani L, Costantini C.Front Biosci (Landmark Ed). 2024 Feb 6;29(2):59. doi: 10.31083/j.fbl2902059.PMID: 38420819

Severe Acute Respiratory Syndrome Coronavirus 2 (SARS-CoV-2) Spike Protein S1 Induces Methylglyoxal-Derived Hydroimidazolone/Receptor for Advanced Glycation End Products (MG-H1/RAGE) Activation to Promote Inflammation in Human Bronchial BEAS-2B Cells.

Manfredelli D, Pariano M, Costantini C, Graziani A, Bozza S, Romani L, Puccetti P, **Talesa VN**, Antognelli C.Int J Mol Sci. 2023 Oct 3;24(19):14868. doi: 10.3390/ijms241914868.PMID: 37834316

Expression of CD13 and CD26 on extracellular vesicles in canine seminal plasma: preliminary results.

Troisi A, Schrank M, Bellezza I, Fallarino F, Pastore S, Verstegen JP, Pieramati C, Di Michele A, **Talesa VN**, Martìnez Barbitta M, Orlandi R, Polisca A.Vet Res Commun. 2024 Feb;48(1):357-366. doi: 10.1007/s11259-023-10202-1. Epub 2023 Sep 14.PMID: 37707657

Microbiota-Associated HAF-EVs Regulate Monocytes by Triggering or Inhibiting Inflammasome Activation.

Nunzi E, Mezzasoma L, Bellezza I, Zelante T, Orvietani P, Coata G, Giardina I, Sagini K, Manni G, Di Michele A, Gargaro M, **Talesa VN**, Di Renzo GC, Fallarino F, Romani R.Int J Mol Sci. 2023 Jan 28;24(3):2527. doi: 10.3390/ijms24032527.PMID: 36768851

Extracellular Vesicles and the Inflammasome: An Intricate Network Sustaining Chemoresistance.

Mezzasoma L, Bellezza I, Romani R, **Talesa VN.**Front Oncol. 2022 Apr 22;12:888135. doi: 10.3389/fonc.2022.888135. eCollection 2022.PMID: 35530309 Review.

Amniotic fluid stem cell-derived extracellular vesicles are independent metabolic units capable of modulating inflammasome activation in THP-1 cells.

Mezzasoma L, Bellezza I, Orvietani P, Manni G, Gargaro M, Sagini K, Llorente A, Scarpelli P, Pascucci L, Cellini B, **Talesa VN**, Fallarino F, Romani R.FASEB J. 2022 Apr;36(4):e22218. doi: 10.1096/fj.202101657R.PMID: 35218567

Anakinra restores cellular proteostasis by coupling mitochondrial redox balance to autophagy.

van de Veerdonk FL, Renga G, Pariano M, Bellet MM, Servillo G, Fallarino F, De Luca A, Iannitti RG, Piobbico D, Gargaro M, Manni G, D'Onofrio F, Stincardini C, Sforna L, Borghi M, Castelli M, Pieroni S, Oikonomou V, Villella VR, Puccetti M, Giovagnoli S, Galarini R, Barola C, Maiuri L, Della Fazia MA, Cellini B, **Talesa VN**, Dinarello CA, Costantini C, Romani L.J Clin Invest. 2022 Jan 18;132(2):e144983. doi: 10.1172/JCI144983.PMID: 34847078

<u>Defective Glyoxalase 1 Contributes to Pathogenic Inflammation in Cystic Fibrosis.</u>

Pariano M, Costantini C, Santarelli I, Puccetti M, Giovagnoli S, **Talesa VN**, Romani L, Antognelli C.Vaccines (Basel). 2021 Nov 11;9(11):1311. doi: 10.3390/vaccines9111311.PMID: 34835243

Metastatic Prostate Cancer Cells Secrete Methylglyoxal-Derived MG-H1 to Reprogram Human Osteoblasts into a Dedifferentiated, Malignant-like Phenotype: A Possible Novel Player in Prostate Cancer Bone Metastases.

Antognelli C, Marinucci L, Frosini R, Macchioni L, **Talesa VN.**Int J Mol Sci. 2021 Sep 22;22(19):10191. doi: 10.3390/ijms221910191.PMID: 34638532

Glyoxalase-1-Dependent Methylglyoxal Depletion Sustains PD-L1 Expression in Metastatic Prostate Cancer Cells: A Novel Mechanism in Cancer Immunosurveillance Escape and a Potential Novel Target to Overcome PD-L1 Blockade Resistance.

Antognelli C, Mandarano M, Prosperi E, Sidoni A, **Talesa VN.**Cancers (Basel). 2021 Jun 13;13(12):2965. doi: 10.3390/cancers13122965.PMID: 34199263

Natriuretic Peptides Regulate Prostate Cells Inflammatory Behavior: Potential Novel Anticancer Agents for Prostate Cancer.

Mezzasoma L, **Talesa VN**, Costanzi E, Bellezza I.Biomolecules. 2021 May 25;11(6):794. doi: 10.3390/biom11060794.PMID: 34070682

Methylglyoxal-Dependent Glycative Stress Is Prevented by the Natural Antioxidant Oleuropein in Human Dental Pulp Stem Cells through Nrf2/Glo1 Pathway.

Delle Monache S, Pulcini F, Frosini R, Mattei V, **Talesa VN**, Antognelli C.Antioxidants (Basel). 2021 May 1;10(5):716. doi: 10.3390/antiox10050716.PMID: 34062923

Pharyngeal Microbial Signatures Are Predictive of the Risk of Fungal Pneumonia in Hematologic Patients.

Costantini C, Nunzi E, Spolzino A, Palmieri M, Renga G, Zelante T, Englmaier L, Coufalikova K, Spáčil Z, Borghi M, Bellet MM, Acerbi E, Puccetti M, Giovagnoli S, Spaccapelo R, **Talesa VN**, Lomurno G, Merli F, Facchini L, Spadea A, Melillo L, Codeluppi K, Marchesi F, Marchesini G, Valente D, Dragonetti G, Nadali G, Pagano L, Aversa F, Romani L.Infect Immun. 2021 Jul 15;89(8):e0010521. doi: 10.1128/IAI.00105-21. Epub 2021 Jul 15.PMID: 33782152

ANP and BNP Exert Anti-Inflammatory Action via NPR-1/cGMP Axis by Interfering with Canonical, Non-Canonical, and Alternative Routes of Inflammasome Activation in Human THP1 Cells.

Mezzasoma L, **Talesa VN**, Romani R, Bellezza I.Int J Mol Sci. 2020 Dec 22;22(1):24. doi: 10.3390/ijms22010024.PMID: 33375031

Antognelli C, Frosini R, Santolla MF, Peirce MJ, **Talesa VN.**Oxid Med Cell Longev. 2020 Sep 9;2020:3045908. doi: 10.1155/2020/3045908. eCollection 2020.PMID: 32963691

Exploring the radiosensitizing potential of AZD8931: a pilot study on the human LoVo colorectal cancer cell line.

Antognelli C, Palumbo I, Piattoni S, Calzuola M, Del Papa B, **Talesa VN**, Aristei C.Int J Radiat Biol. 2020 Nov;96(11):1504-1512. doi: 10.1080/09553002.2020.1820610. Epub 2020 Sep 22.PMID: 32910714

JNK signaling regulates oviposition in the malaria vector Anopheles gambiae.

Peirce MJ, Mitchell SN, Kakani EG, Scarpelli P, South A, Shaw WR, Werling KL, Gabrieli P, Marcenac P, Bordoni M, **Talesa V**, Catteruccia F.Sci Rep. 2020 Sep 1;10(1):14344. doi: 10.1038/s41598-020-71291-5.PMID: 32873857

 $\underline{\textit{Redox-Sensitive Glyoxalase 1 Up-Regulation Is Crucial for Protecting Human Lung Cells from Gold Nanoparticles Toxicity}.$

Gambelunghe A, Giovagnoli S, Di Michele A, Boncompagni S, Dell'Omo M, Leopold K, Iavicoli I, **Talesa VN**, Antognelli C.Antioxidants (Basel). 2020 Aug 3;9(8):697. doi: 10.3390/antiox9080697.PMID: 32756399

Spectrophotometric Method for Determining Glyoxalase 1 Activity in Cerebral Cavernous Malformation (CCM) Disease.

Antognelli C, **Talesa VN**, Retta SF.Methods Mol Biol. 2020;2152:445-449. doi: 10.1007/978-1-0716-0640-7 33.PMID: 32524572

Pharmacologic Induction of Endotoxin Tolerance in Dendritic Cells by L-Kynurenine.

Manni G, Mondanelli G, Scalisi G, Pallotta MT, Nardi D, Padiglioni E, Romani R, **Talesa VN**, Puccetti P, Fallarino F, Gargaro M.Front Immunol. 2020 Mar 11;11:292. doi: 10.3389/fimmu.2020.00292. eCollection 2020.PMID: 32226425

Extracellular Vesicles from Human Advanced-Stage Prostate Cancer Cells Modify the Inflammatory Response of Microenvironment-Residing Cells.

Mezzasoma L, Costanzi E, Scarpelli P, **Talesa VN**, Bellezza I.Cancers (Basel). 2019 Aug 30;11(9):1276. doi: 10.3390/cancers11091276.PMID: 31480312

Oleuropein-Induced Apoptosis Is Mediated by Mitochondrial Glyoxalase 2 in NSCLC A549 Cells: A Mechanistic Inside and a Possible Novel Nonenzymatic Role for an Ancient Enzyme.

Antognelli C, Frosini R, Santolla MF, Peirce MJ, **Talesa VN.**Oxid Med Cell Longev. 2019 Jul 22;2019:8576961. doi: 10.1155/2019/8576961. eCollection 2019.PMID: 31428230

Methylglyoxal Acts as a Tumor-Promoting Factor in Anaplastic Thyroid Cancer.

Antognelli C, Moretti S, Frosini R, Puxeddu E, Sidoni A, **Talesa VN.**Cells. 2019 Jun 6;8(6):547. doi: 10.3390/cells8060547.PMID: 31174324

<u>Testosterone and Follicle Stimulating Hormone-Dependent Glyoxalase 1 Up-Regulation Sustains the Viability of Porcine</u> Sertoli Cells through the Control of Hydroimidazolone- and Argpyrimidine-Mediated NF-κB Pathway.

Antognelli C, Mancuso F, Frosini R, Arato I, Calvitti M, Calafiore R, **Talesa VN**, Luca G.Am J Pathol. 2018 Nov;188(11):2553-2563. doi: 10.1016/j.ajpath.2018.07.013. Epub 2018 Aug 18.PMID: 30125541

S1P promotes migration, differentiation and immune regulatory activity in amniotic-fluid-derived stem cells.

Romani R, Manni G, Donati C, Pirisinu I, Bernacchioni C, Gargaro M, Pirro M, Calvitti M, Bagaglia F, Sahebkar A, Clerici G, Matino D, Pomili G, Di Renzo GC, **Talesa VN**, Puccetti P, Fallarino F.Eur J Pharmacol. 2018 Aug 15;833:173-182. doi: 10.1016/j.ejphar.2018.06.005. Epub 2018 Jun 7.PMID: 29886240

Eicosapentaenoic acid induces DNA demethylation in carcinoma cells through a TET1-dependent mechanism.

Ceccarelli V, Valentini V, Ronchetti S, Cannarile L, Billi M, Riccardi C, Ottini L, **Talesa VN**, Grignani F, Vecchini A.FASEB J. 2018 May 14:fj201800245R. doi: 10.1096/fj.201800245R. Online ahead of print.PMID: 29757674

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Deficiency of immunoregulatory indoleamine 2,3-dioxygenase 1in juvenile diabetes.

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Antognelli C, Trapani E, Delle Monache S, Perrelli A, Fornelli C, Retta F, Cassoni P, **Talesa VN**, Retta SF.Data Brief. 2017 Dec 13;16:929-938. doi: 10.1016/j.dib.2017.12.026. eCollection 2018 Feb.PMID: 29511711

Glyoxalase 1 sustains the metastatic phenotype of prostate cancer cells via EMT control.

Antognelli C, Cecchetti R, Riuzzi F, Peirce MJ, **Talesa VN.**J Cell Mol Med. 2018 May;22(5):2865-2883. doi: 10.1111/jcmm.13581. Epub 2018 Mar 5.PMID: 29504694

Glyoxalases in Urological Malignancies.

Antognelli C, **Talesa VN.**Int J Mol Sci. 2018 Jan 31;19(2):415. doi: 10.3390/ijms19020415.PMID: 29385039 Review.

Nicotine induces apoptosis in human osteoblasts via a novel mechanism driven by H_2O_2 and entailing Glyoxalase 1-dependent MG-H1 accumulation leading to TG2-mediated NF-kB desensitization: Implication for smokers-related osteoporosis.

Marinucci L, Balloni S, Fettucciari K, Bodo M, **Talesa VN**, Antognelli C.Free Radic Biol Med. 2018 Mar;117:6-17. doi: 10.1016/j.freeradbiomed.2018.01.017. Epub 2018 Jan 31.PMID: 29355739

KRIT1 loss-of-function induces a chronic Nrf2-mediated adaptive homeostasis that sensitizes cells to oxidative stress: Implication for Cerebral Cavernous Malformation disease.

Antognelli C, Trapani E, Delle Monache S, Perrelli A, Daga M, Pizzimenti S, Barrera G, Cassoni P, Angelucci A, Trabalzini L, **Talesa VN**, Goitre L, Retta SF.Free Radic Biol Med. 2018 Feb 1;115:202-218. doi: 10.1016/j.freeradbiomed.2017.11.014. Epub 2017 Nov 21.PMID: 29170092

<u>Influence of chemotherapeutic drug-related gene polymorphisms on toxicity and survival of early breast cancer patients receiving adjuvant chemotherapy.</u>

Ludovini V, Antognelli C, Rulli A, Foglietta J, Pistola L, Eliana R, Floriani I, Nocentini G, Tofanetti FR, Piattoni S, Minenza E, **Talesa VN**, Sidoni A, Tonato M, Crinò L, Gori S.BMC Cancer. 2017 Jul 26;17(1):502. doi: 10.1186/s12885-017-3483-2.PMID: 28747156

An intensive lifestyle intervention reduces circulating oxidised low-density lipoprotein and increases human paraoxonase activity in obese subjects.

Russo A, Pirisinu I, Vacca C, Reginato E, Tomaro ES, Pippi R, Aiello C, **Talesa VN**, De Feo P, Romani R.Obes Res Clin Pract. 2018 Jan-Feb;12(Suppl 2):108-114. doi: 10.1016/j.orcp.2016.11.006. Epub 2016 Dec 10.PMID: 27956218