

Esposizione alla nicotina, invecchiamento, machine learning e blockchain solution: un approccio multilivello basato su modelli in vitro e pazienti per costruire una road map del rischio perCovid-19 (IFT)

Nicotine Changes Airway Epithelial Phenotype and May Increase the SARS-COV-2 Infection Severity

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Abstract:

Background: Nicotine is implicated in the SARS-COV-2 infection through activation of the $\alpha 7$ -nAChR and over-expression of ACE2. Our objective was to clarify the role of nicotine in SARS-CoV-2 infection exploring its molecular and cellular activity.

Methods: HBEpC or si-mRNA- $\alpha 7$ -HBEpC were treated for 1 h, 48 h or continuously with 10–7 M nicotine, a concentration mimicking human exposure to a cigarette. Cell viability and proliferation were evaluated by trypan blue dye exclusion and cell counting, migration by cell migration assay, senescence by SA- β -Gal activity, and anchorage-independent growth by cloning in soft agar. Expression of Ki67, p53/phospho-p53, VEGF, EGFR/pEGFR, phospho-p38, intracellular Ca²⁺, ATP and EMT were evaluated by ELISA and/or Western blotting.

Results: nicotine induced through $\alpha 7$ - nAChR

- (i) increase in cell viability,
- (ii) cell proliferation,
- (iii) Ki67 over-expression,
- (iv) phospho- p38 up-regulation,
- (v) EGFR/pEGFR over-expression,
- (vi) increase in basal Ca²⁺ concentration,
- (vii) reduction of ATP production,
- (viii) decreased level of p53/phospho-p53,
- (ix) delayed senescence,
- (x) VEGF increase,
- (xi) EMT

and consequent

- (xii) enhanced migration,

and

- (xiii) ability to grow independently of the substrate.

Conclusions: Based on our results and on evidence showing that nicotine potentiates viral infection, it is likely that nicotine is involved in SARS-CoV-2 infection and severity.

References: <https://doi.org/10.3390/molecules26010101>

Keywords: Cell Proliferation; EMT; Mitochondrial Dysfunction; nAChR; Nicotine; SARS-CoV-2

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