



# CYTOKINE EXPRESSION IN CIRCULATING MONOCYTES DURING THE ACUTE PHASE OF ISCHEMIC STROKE

A. Semerano, L. Peruzzotti-Jametti G. Giacalone, D. Strambo, G. Comi, M. Sessa, M. Bacigaluppi  
Neurology Department, San Raffaele Scientific Institute, Via Olgettina 58, 20132, Milan, Italy  
e-mail: semerano.aurora@hsr.it

## BACKGROUND

Complex interactions between the CNS and the immune peripheral system in cerebrovascular diseases are increasingly recognized and appear to be bidirectional<sup>1</sup>. Monocytes have been recognized as important players, being both rapidly recruited in the infarcted area<sup>2</sup> and increased in peripheral blood within the first 24 hours after stroke<sup>3</sup>. Furthermore circulating monocytes have been advocated as key players in the prognosis and risk of infections after stroke<sup>3,4</sup>.

## AIM

We aimed to examine the cytokine expression profile of circulating monocytes in acute ischemic stroke and its correlation with the patient outcome.

## METHODS

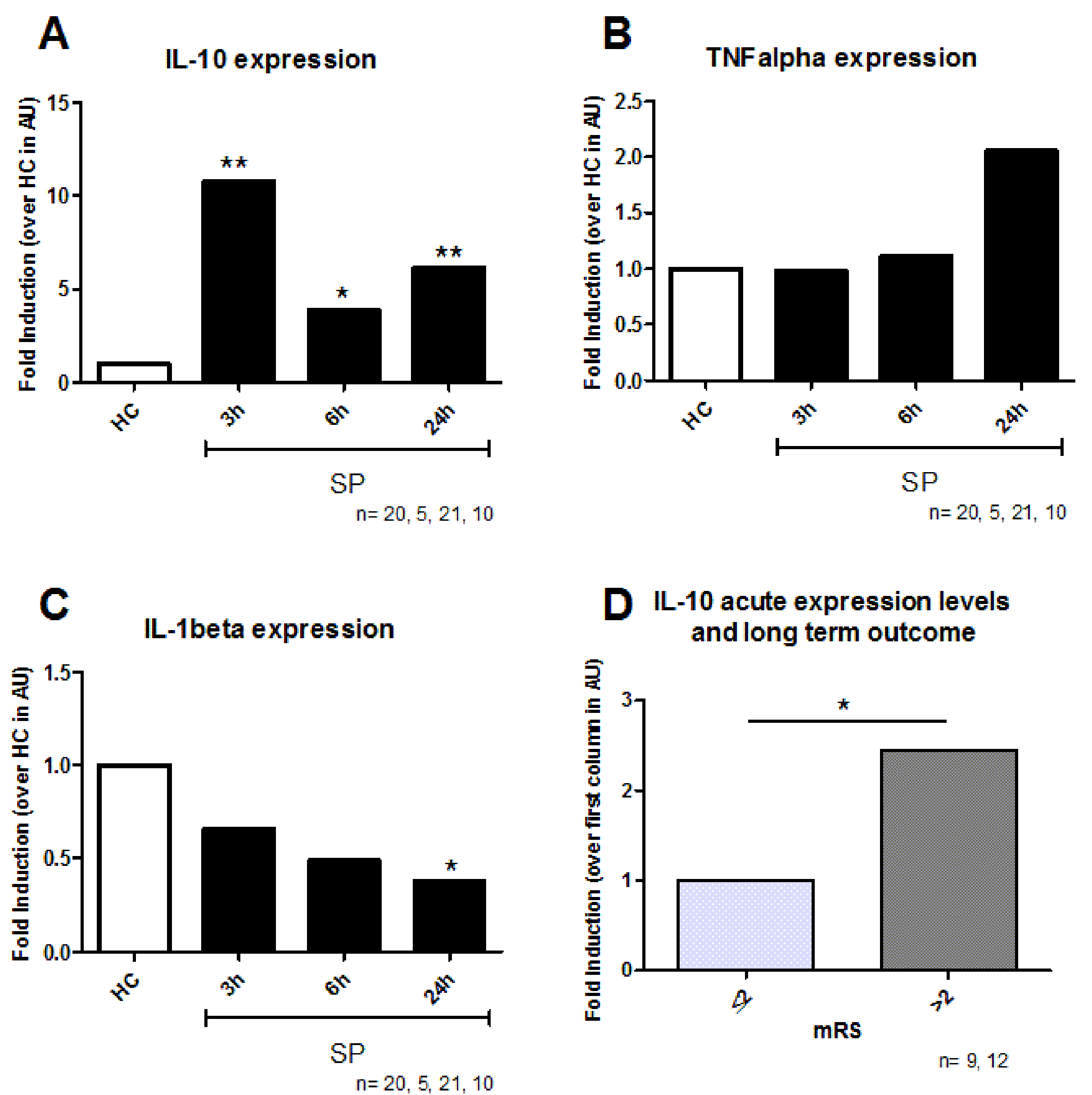
We collected blood samples from 21 stroke patients (SP) at 3, 6 and 24 hours after symptom onset and from 20 healthy controls (HC). From each sample we measured the mRNA levels of IL-10, TNFalpha and IL-1beta expressed by circulating monocytes.

## RESULTS

Compared to HC, in SP the IL-10 expression significantly increased at 3h after symptom onset ( $p < 0.01$ ) and high levels persisted at 6h ( $p < 0.05$ ) and 24h ( $p < 0.01$ ). IL-1beta was reduced ( $p < 0.05$ ) whereas TNFalpha was increased within the first 24h after stroke, though last finding was not statistically significant. High levels of IL-10 expression during the acute phase were associated with poor outcome at 3 months ( $mRS > 2$ ), while association with infections after stroke was not significant.

## REFERENCES.

- <sup>1</sup>Kamel H, Iadecola C, *Brain-immune interactions and ischemic stroke: clinical implications*, Arch Neurol. 2012 69(5):576-81
- <sup>2</sup>Iadecola C., *The immunology of stroke: from mechanisms to translation*, Nat. Med. 2011 7;17(7):796-808
- <sup>3</sup>Urra X et al., *Monocytes are major players in the prognosis and risk of infection after acute stroke*, Stroke 2009 40(4):1262-8
- <sup>4</sup>Meisel A et al., *Predicting post-stroke infections and outcome with blood-based immune and stress markers*, Cerebrovasc Dis. 2012 33(6):580-8.



**FIGURE. (A, B, C) IL-10, TNFalpha and IL-1beta gene expression in stroke patients and healthy controls.** Data are expressed as Fold Inductions (over healthy controls in arbitrary units). Statistics: Kruskal-Wallis test followed by Dunn's Multiple Comparison Test. \*= $p < 0.05$ ; \*\*= $p < 0.01$ . **(D) IL-10 gene expression at 6h from symptom onset in patients with good ( $mRS \leq 2$ ) and poor ( $mRS > 3$ ) three-month outcome.** Data are expressed as Fold Inductions (over first column in arbitrary units). Statistics: Unpaired t Test. \*= $p < 0.05$ . Abbreviations: IL-10, interleukin-10; TNFalpha, Tumour Necrosis Factor alpha; IL-1beta, interleukin-1 beta; HC, healthy controls; SP, stroke patients; AU, arbitrary units; mRS, modified Rankin Scale.

## CONCLUSIONS

Circulating monocytes modify their pattern of cytokine expression acutely after stroke, mainly resulting in increased IL-10 expression. High levels of IL-10 showed correlation with 3-month outcome, suggesting that peripheral immune activation during acute stroke might have possible clinical implications.