(p = 0.03, r² = 0.49) and higher NPI total scores (p = 0.01, r² = 0.63) (see Figures S1A and S1B, respectively; available as supplementary material attached to the electronic version of this letter at www.journals.cambridge.org/jid_IPG). Similarly, there were trend level correlations of IL-6 release with worse ratings on the CDR (p = 0.08, r² = 0.33) and MMSE (p = 0.05, r² = 0.39).

In this pilot project, we sought to replicate prior findings suggesting that IL-6 release from LPS-stimulated PBMCs is higher in AD than cognitively normal controls. We found increased IL-6 release ratios in AD, with trend level significance at the higher LPS concentration only. This is not surprising given the small sample; since the results are in the predicted direction they offer support for hypothesis 1. We sought to extend prior results by adding an AD-specific stimulus (A-β1-42), in an attempt to more closely model the neuroinflammatory processes of AD. Modest support was seen for hypothesis 2 with A-β1-42 increasing the IL-6 release ratio by a factor of 2 in AD patients but not controls, although this was not statistically significant; these results are in the direction predicted by hypothesis 2.

Additionally we found a significant trend correlation between IL-6 release ratio and several clinical variables, including verbal category fluency (animals), NPI, CDR and MMSE. A higher IL-6 release ratio was associated with worse illness severity (higher NPI total and CDR, lower MMSE and category fluency. This suggests that the IL-6 release ratio may have value as a marker of AD disease severity, especially severity of neuropsychiatric symptoms (NPS). The latter appears to be a unique finding, and may reflect the involvement of inflammatory mechanisms (peripheral and central) in the development of NPS.

This study is limited by small sample size, assessment of only one candidate cytokine, and limited capacity to assess multiple reaction conditions. The strengths of the study include a well-characterized cohort of AD and control participants and the extension of prior results to neuropsychiatric and functional as well as cognitive findings.

These preliminary results suggest that IL-6 release from PBMCs may be a useful marker of disease severity and NPS severity in AD, thus meriting further study.

References


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The relationship between annual predicted future population growth rates and elderly suicide rates

Two paradoxical hypotheses – (i) that countries with low population growth rates or a decline in population growth rate will have high elderly dependency ratios leading to high elderly suicide rates, and (ii) that countries with high population growth rates will have high elderly suicide rates because of Durkheim’s hypothesis that the overall cohort size may influence suicide rates due to competition for scarce resources – were supported by a recent study (Shah, 2008). The relationship between average annual population growth rates and elderly suicide rates was shown to be curvilinear (U-shaped curve) fitting the quadratic equation \( y = a + bx + cx^2 \) (where \( y \) is the elderly suicide rate, \( x \) is the average

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Population growth rate is a function of life expectancy and birth rate. Thus, in countries with low average annual population growth rates or decline in average annual population growth rates, elderly dependency ratios may be high because of increased life expectancy and low birth rate. This, in turn, may result in increased elderly suicide rates. The average annual growth rate may begin to increase in some countries due to an increase in birth rates. This, in turn, may lead to a decline in the elderly dependency ratios. Moreover, in turn, this may result in a decline in elderly suicide rates. As the average annual population growth rate continues to increase, at a critical point the composite influence of increased life expectancy, increased elderly population size and the increase in the proportion of elderly in the general population on elderly suicide rates may become more prominent. This may, in part, reflect Durkheim’s hypothesis that the overall cohort size may influence suicide rates due to competition for scarce resources (Shah and De, 1998).

The same curvilinear (U-shaped curve) relationship between elderly suicide rates and the average annual predicted future population growth rate was examined. Data on elderly suicide rates for males and females in the age-bands 65–74 years and 75+ years for each listed country were ascertained from the World Health Organization (WHO) website (www.who.int/whosis/mort/table1.cfm). The median (range) of the latest available year for data on elderly suicide rates was 2000 (1985–2003). Data on the average annual predicted future population growth rate until 2015 for each listed country were ascertained from the United Nations Development Programme (UNDP) website (www.hdr.undp.org/reports/global/2005/pdf/hdr05_HDI.pdf). Curve estimation regression models were used to examine the curvilinear relationship between elderly suicide rates and the average annual predicted population population growth fitting the quadratic equation $y = a + bx + cx^2$.

Full data sets for elderly suicide rates and the average annual predicted future population growth were available for 80 countries. Table S1 (available online at www.journals.cambridge.org/jid_IPG as supplementary material to the electronic version of this letter) illustrates the curve estimation regression models, whereby the relationships between suicide rates in both sexes in both elderly age-bands and the average annual predicted future population growth rates were curvilinear (U-shaped curve) and fitted the quadratic equation $y = a + bx + cx^2$; the significance level was at least at the 0.05 level.

How can the average annual predicted future population growth rates predict current suicide rates given that this population growth has not yet occurred? It is possible that this observed relationship was purely due to chance. Also, the average annual predicted future population growth rate may be a proxy measure for other correlates of elderly suicides or other variables may predict both elderly suicide rates and the average annual predicted future population growth rates (epiphenomena). The findings of this brief study illustrate that considerable caution and care are required in interpreting findings from cross-sectional ecological studies exploring potential etiological relationships.

References


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Cultural aspects of the patient–doctor relationship

The patient–doctor relationship is central to medicine. This relationship has two fundamental components (Calman and McLean, 1984). The first is the doctor’s care, skill and knowledge; the second is information-giving to help the individual make decisions. Hence, communication and trust are essential in this relationship.

The problem arises when doctors see patients as sick people from whom to extract information or to whom to impart advice, and they therefore ignore a vital purpose of communication, which is to initiate and enhance the relationship with their patients (Persaud, 2005). Long and Jiwa (2004) found that in 25% of medical consultations, the chief concerns of patients had not been elicited and 40% of cancer