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Effects of new antiepileptic drugs and progabide on the mitogen-induced proliferative activity of mouse splenocytes

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Abstract:

Classical antiepileptic drugs are known to affect immune system activity, although the effects of new generation anticonvulsants on T- and B-cell-mediated immunity remain unknown. Therefore, in the present study, we compared a selection of new antiepileptic drugs with classical ones in terms of their effects on the proliferative activity of lymphocytes stimulated by concanavalin A (Con A) and lipopolysaccharide (LPS). Felbamate $(3 \times 10^{-6} - 10^{-4} \text{ M})$ was the most potent in inhibiting [³H]-thymidine incorporation in C57BL/6 mouse spleen cells stimulated by Con A and LPS. Treatment of the cells with stiripentol $(3 \times 10^{-5} \text{ and } 10^{-4} \text{ M})$ and lorecle-zole (10^{-4} M) suppressed the proliferative activity of splenocytes both after Con A and LPS stimulation. Tiagabine $(3 \times 10^{-5} \text{ M and } 10^{-4} \text{ M})$ inhibited the Con A-induced cell proliferation, whereas the effect of LPS was attenuated only by the highest concentration of this drug (10^{-4} M) . Progabide showed immunosuppressive effects at concentrations of 3×10^{-5} and 10^{-4} M or only 10^{-4} M after LPS or Con A stimulation, respectively. No effect on the immune parameters was observed after lamotrigine treatment. On the other hand, the Con A-induced proliferation of splenocytes was enhanced by carbamazepine $(10^{-5} - 10^{-4} \text{ M})$ and sodium valproate $(5 \times 10^{-4} - 3 \times 10^{-3} \text{ M})$. Neither carbamazepine nor sodium valproate affected the LPS-induced proliferation. These data indicate that some new antiepileptic drugs, especially felbamate at pharmacological concentrations, may suppress the mitogen-stimulated proliferative activity of mouse splenocytes. In contrast, two classical anticonvulsants (carbamazepine and sodium valproate) stimulated T-cell-mediated immunity. The above differences in the effects of particular antiepileptic drugs on the immune response may play roles in both their therapeutic efficiency and undesired effects.

Key words:

antiepileptic drugs, mitogens, proliferative activity of splenocytes