

Antiproliferative Activity of Methylated Analogues of *E*- and *Z*-Resveratrol

Venera Cardile^a, Rosa Chillemi^b, Laura Lombardo^a, Sebastiano Sciuto^b, Carmela Spatafora^b, and Corrado Tringali^{b,*}

^a Dipartimento di Scienze Fisiologiche, Università di Catania, Viale A. Doria 6, I-95125, Catania, Italy

^b Dipartimento di Scienze Chimiche, Università di Catania, Viale A. Doria 6, I-95125, Catania, Italy. Fax: +39095 580138. E-mail: ctringali@unict.it

* Author for correspondence and reprint requests

Z. Naturforsch. **62c**, 189–195 (2007); received November 16/December 13, 2006

The stilbenoids *E*-resveratrol (*E*-3,5,4'-trihydroxystilbene, **1**), *E*-3,5,4'-trimethoxystilbene (**2**), *E*-3,4,4'-trimethoxystilbene (**3**) and *E*-3,4'-dimethoxy-5-hydroxystilbene (**4**) were converted by photoisomerization to their corresponding *Z*-isomers **5–8**. Compounds **1–8** were subjected to antiproliferative activity bioassays towards a set of four different human cancer cell lines, namely DU-145 (androgen not responsive human prostate tumor), LNCaP (androgen responsive human prostate tumor), M-14 (human melanoma) and KB (human mouth epidermoid carcinoma). The methylated analogues of **1** are more active than the natural lead in the majority of bioassays. The most active compound was *Z*-3,5,4'-trimethoxystilbene (**6**), which showed against DU-145 and LNCaP cells GI₅₀ values close to those of the anticancer drug vinorelbine; **6** resulted more active than its *E*-isomer **2** towards DU-145, LNCaP and especially KB cell lines. A number of methylated *Z*-isomers displayed a higher activity than their *E*-isomers, but *E*-resveratrol (**1**) was more active than *Z*-resveratrol (**5**) towards all the tested cell lines.

Key words: Resveratrol Analogues, Antiproliferative Activity