SYNTHESIS OF SOME N'-n-DIMETHYLPHENYL-N,N-DIMETHYLTHIOUREAS WITH ANTIPROLIFERATIVE PROPERTIES

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Methods for synthesis of newcompounds of thioureeaderivatives are described. These compounds are of both theoretically and practically interest through heir aplications in various fields, as they posses valuable biological activit ysuch as antifungal, antitumour, antiviral, antibacterial, pharmacological, herbicidal, and insecticidal properties. Thioureas have suitable C=S function groups, and they can be considered as useful chelating agents due to their ability to encapsulate into their coordinating moiety metal ions[1].

An efficient method for the synthesis of unsymmetrical substituted thiourea derivatives by means of simple condensation between available building blocks such as amines and carbon disulfide in aqueous medium, but this method uses a toxic reactive as carbon disulfide so we tried to substituteit with the accessible and nontoxic reactive as tetramethyldithiocarbamate (DTMT). Synthesis of N¹-n-dimethylphenyl-N, N- dimethylthioureas was carried between n-dimethylaniline and (DTMT). The ends of the reactions were checked by thin layer chromatography (TLC). N¹-n-dimethylphenyl-N, N-dimethylthioureas presents white crystalline products, their structure was confirmed by ¹H and ¹³C NMR Spectra and are presented in table below:

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Nr.	Name	Melting point, °C	H ¹ NMR	C ¹³ NMR
			DMSO-d ₆	$DMSO-d_6$
1	N ¹ -2,3-dimetilfenil-N,N-	170-172	8.8(s.1H,NH),7-6.8(m, Ar-H), 3.45(s,	14.7, 20.6, 41.1, 125.3,
	dimetiltiourea		N-CH ₃),	127.2, 128.0, 134.7, 137.1,
			2.03, 2.24(s, CH ₃ -Ar)	140.0, 181.9
2	N ¹ -2,4-dimetilfenil-N,N-	160-162	8.76(s.1H,NH),7.01-6.9(m, Ar-H),	18.3, 21.05, 41.1 126.9,
	dimetiltiourea		3.26(s, N-CH ₃),	129.3, 131.0, 135.7, 135.8,
			2.27,2.12(s, CH ₃ -Ar)	137.7, 182.07
3	N ¹ -2,5-dimetilfenil-N,N-	145-147	8.75(s.1H,NH),7.12-6.8(m, Ar-H),	17.9, 20.8, 41.1, 127.0,
	dimetiltiourea		3.45(s, N-CH ₃),	129.0, 130.0, 133.0, 135.0,
			2.26, 2.12(s, CH ₃ -Ar)	140.0, 181.9
4	N ¹ -2,6-dimetilfenil-N,N-	140-142	8.80(s.1H,NH),7.72-7.48(m, Ar-H),	19.0, 40.6, 125.6, 126.7
	dimetiltiourea		3.42(s, N-CH ₃),2.27(s, CH ₃ -Ar)	128.3, 128.8, 129.6, 131.5,
				176.8
5	N ¹ -3,4-dimetilfenil-N,N-	180-182	8.88(s.1H,NH),7.2-6.8(m, Ar-H),	19.32, 19.9, 41.2, 123.0,
	dimetiltiourea		3.25(s, N-CH ₃),	127.0, 129.0, 132.0, 135.0,
			2.18(s, CH ₃ -Ar)	139.0, 181.8
6	N ¹ -3,5-dimetilfenil-N,N-	190-192	8.80(s.1H,NH),6.89-6.74(m, Ar-	21.4, 41.4, 123.0, 126.1,
	dimetiltiourea		H),3.25(s, N-CH ₃), 2.24(s, CH ₃ -Ar)	126.4, 136.0, 137.0, 141.0,
				181.6

These compounds were tested as anti proliferative agents.

^{1.} Gulea A., Poirier D., Roy J., Stavila V., Bulimestru I., Țapcov V., Bârcă M., Popovschi L. In vitro antileukemia, antibacterial and antifungal activities of some 3d metal complexes: Chemical synthesis and structure – activity relationships // Journal of Enzyme Inhibition and Medicinal Chemistry, 2008; V. 23. Nr.6, pp.806-818.