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For Sustainable Development

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PREFACE

Assalamu'alaikum Wr. Wb.

All praises are due to ALLAH SWT, God Almighty, Who made this International Conference of successful. The International Conference on Natural, Mathematical and Environmental Sciences (NAMES) 2015 is organized by the Faculty of Mathematics and Natural Sciences (FMIPA), Lambung Mangkurat University, Banjarbaru, South Kalimantan. This conference covered a wide range of topics, including biology, chemistry, physics, mathematics, pharmacy, computer science, material Science, and environmental science.

All papers were compiled into the proceedings book which had six sections, namely Biology, Pharmacy, Chemistry, Mathematics, Physics, and Computer Sciences. This book was also published in the NAMES Website <http://names.fmipa.unlam.ac.id>. I am glad that for the first time both types of books can be realized.

The seminar took a theme of "Sustainable Development" as a hot issue in Banjarbaru. Banjarbaru, is a fast growing city in the province of South Kalimantan, Indonesia and famously known as an urban city with a unique natural landscape, a cultural diversity, and a friendly welcoming citizen. Moreover, Banjarbaru becomes the centre of provincial government that its government is located in Banjarmasin today. The conference provided an ideal platform to share information and discuss their scientific results and experiences, with particular references to sustain development.

I was fully satisfied to all the members of the program committee who contributed for the success in framing the program and the books. My appreciation was especially from all delegates who contributed to the success of this conference by accepting our invitation and submitting articles for presentation in the scientific program.

I can guarantee you that this book provides a full of intellectual scientific research activities. I do hope the next conference will pick up similar success, and even better.

Wassalamualaikum Wr. Wb.
Banjarbaru, March 2016

Dr. Krisdianto
Chief of Executive

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The effect of Cl substituent position in benzoylchloride reagent towards the yield of synthesis of the novel lead and substituted compounds of 1-benzoyl-3-benzylurea as prospective anticancers

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ABSTRACT

This study is related to design and synthesize of a novel lead compound 1-benzoyl-3-benzylurea and the compounds which are substituted with Cl substituent, as prospective anticancers with antiproliferation mechanism. The purpose of this study is to obtain a novel lead compound 1-benzoyl-3-benzylurea and the compounds which are substituted with Cl substituent at a different position. In order to reach such purpose, the synthesis with nucleophilic substitution reaction and modified Schotten Baumann method was done. Benzylurea as a starting material is reacted with the lead and substituted benzoylchloride reagents (2-Cl, 3-Cl and 4-Cl). Four compounds are yielded from the synthesis, which are 1-benzoyl-3-benzylurea 36%, 1-benzoyl-3-(2-chlorobenzyl)urea 8%, 1-benzoyl-3-(3-chlorobenzyl)urea 22% and 1-benzoyl-3-(4-chlorobenzyl)urea 36%. The position of substituent towards the carbonyl group affects the compound's electrophilic force. The closer the substituent to the carbonyl group, the less the electrophilic force is. Due to the fact that the synthesis yields are relatively low, it is highly recommended to do further optimization in synthesis to all of the lead and substituted compounds 1-benzoyl-3-benzylurea as prospective anticancers.

Keywords: the lead and substituted compounds 1-benzoyl-3-benzylurea, design, synthesize

Introduction

Cancer is a health issue all over the world currently, including in Indonesia. In 2013, the data of the World Health Organization, abbreviated as WHO, shows that cancer is in the 2nd place of most common cause of death, with cardiovascular as the 1st. In between 2008-2014, the number of cancer cases increases from 12.7 million to 14.1 million, followed by the number of death from 7.6 million in 2008 to 8.2 million in 2014. Based on the Basic Health Research (*Riset Kesehatan Dasar*, abbreviated to *Riskedas*) by the research and development division of Indonesian Republic's Health Department in 2007, cancer sits in the 6th place of the biggest cause of death in Indonesia. According to the 2013th *Riskedas*, cancer can strike any person in any ages, regardless the economic status and the education status of the person. The national cancer number in Indonesia is 1.4 per 1000 citizens, with the higher number on female victims of 5.7 per 1000 citizens compared to male victims of 2.9 per 1000 citizens.

In order to control cancer, the Health Ministry has done an effort through risk factor management and early detection approach. Besides such approach, efforts of patient medicating in hospitals and palliative clinics have also been done. However, all of those efforts will not work efficiently without the supports from every related factor, along with every single layer of society (www.depkes.go.id). Due to such need, the role of pharmacists as the medication provider is essential, as a factual example, in the chemotherapy medication. Chemotherapy is one of the cancer medication procedures by using anti-cancer drugs to kill or inhibit the activity of cancer cell while it is exercising its fission through various mechanisms.

Hydroxyurea is an anti-cancer drug which has been used for some decades to medicate some types of cancer. This compound dissolves in water which is distributed equally all around the body through the body fluids (Saban and Bujak, 2009). The best *Ribonucleotide reductase* inhibitor, abbreviated as RNR, is hydroxyurea, which will be absorbed after oral administration, transportation through cells, and activation of RNR enzyme, by tying the radical tyrosil in the RNR's active side.

Researches regarding the urea derivate have been done many times with urea group as the group which influences the anti-cancer activity (pharmacophore). Lokwani et al. (2011) and Lu et al. (2012) stated