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Research Article

48. Total Flavonoid Levels of Ethanol Extract and Ethyl Acetate Fraction Dry Shallots (*Allium cepa* L. var. Garden Onion of Brebes) with Maceration Methods Using UV-Vis Spectrophotometry

Heru Nurcahyo, Sri Adi Sumiwi, Eli Halimah, Gofarana Wilar

SRP. 2020; 11(10): 286-289

» Abstract (?mno=25386) » PDF (index.php?fulltxt=25386&fulltxtj=196&fulltxtp=196-1604396654.pdf) » doi: 10.31838/srp.2020.10.48 (<http://dx.doi.org/10.31838/srp.2020.10.48>)

49. In Silico Study on Antibacterial Activity and Brazilein ADME of Sappan Wood (*Caesalpinia Sappan L.*) Against *Escherichia coli* (Strain K12)

Dwi Krihariyani, Eddy Bagus Wasito, **Isnaeni Isnaeni**, Siswandono Siswodihardjo, Wiwik Misaco Yuniarti, Entuy Kurniawan

SRP. 2020; 11(10): 290-296

» Abstract (?mno=25708) » PDF (index.php?fulltxt=25708&fulltxtj=196&fulltxtp=196-1604435838.pdf) » doi: 10.31838/srp.2020.10.49 (<http://dx.doi.org/10.31838/srp.2020.10.49>)

Regular Article

50. Did Trilogy Leadership Style, Organizational Citizenship Behaviour (OCB) and Organizational Commitment (OCO) Influence Financial Performance? Evidence from Pharmacy Industries

Harjoni Desky, Mukhtasar, Muhammad Istan, Yeni Ariesa, Inge Bunga Mira Dewi, Mochammad Fahlevi, Muhammad Nur Abdi, Rinto Noviantoro, Agus Purwanto

SRP. 2020; 11(10): 297-305

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51. Adaptive Strategy of Women`s Leprosy in Indonesia: Psychic Experience of Women with Leprosy in Living a Community Life

Abd. Nasir, Ah Yusuf, Muhammad Yulianto Listiawan, Susilo Harianto, Nuruddin, Nuh Huda

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Research Article

52. Anticancer Activity of Phyllanthus Niruri Linn Extract in Colorectal Cancer Patients: A phase II Clinical Trial

Muhammad Sayuti, Ignatius Riwanto, B. Parish Boediono, Teuku Ilhami Surya Akbar

SRP. 2020; 11(10): 313-317

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53. The Effect of Anthocyanin of Whole-Grain Pigmented Rice Attenuated Visceral Fat, Cholesterol, LDL and PPAR γ Gene Cascade in Dyslipidemia Rat

Fatchiyah Fatchiyah, Anna Safitri, Rista Nikmatu Rohmah, Lidwina Faraline Triprisila, Nia Kurnianingsih, Yudhistira Nugraha, Sisca Fajriani, Hazna Noor Meidinna, James Ketudat Robert-Cairns

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Regular Article**54. Public Management in the Activity of Educational Institutions in the Context of Autonomy of their Work (Under the Conditions of Crises and Covid-19)**

Budanova Liana, Filyanina Nelya, Nosyk Oksana, Andriyenko Mykola, Slepchenko Anzhela, Zastrozhnikova Irina

SRP. 2020; 11(10): 328-334» Abstract (?mno=27400) » PDF (index.php?fulltxt=27400&fulltxtj=196&fulltxtp=196-1604667828.pdf) » doi:
10.31838/srp.2020.10.54 (<http://dx.doi.org/10.31838/srp.2020.10.54>)**Research Article****55. Combination Therapy of Eurycomanone and Doxorubicin As Anticancer on T47D and MCF-7 Cell Lines**

Hanifah Yusuf, Denny Satria, Suryawati Suryawati, Marhami Fahriani

SRP. 2020; 11(10): 335-341» Abstract (?mno=27468) » PDF (index.php?fulltxt=27468&fulltxtj=196&fulltxtp=196-1604676311.pdf) » doi:
10.31838/srp.2020.10.55 (<http://dx.doi.org/10.31838/srp.2020.10.55>)**Review Article****56. Notch Pathway and its Role in Cardiovascular System: Review**

Amira M. Badr, Yasmen F. Mahran

SRP. 2020; 11(10): 342-349» Abstract (?mno=27474) » PDF (index.php?fulltxt=27474&fulltxtj=196&fulltxtp=196-1604678144.pdf) » doi:
10.31838/srp.2020.10.56 (<http://dx.doi.org/10.31838/srp.2020.10.56>)**Regular Article****57. CERDIK Behavior as a Risk Factor for Individuals with Non-Communicable Diseases**

Suprajitno, Sri Mugianti

SRP. 2020; 11(10): 350-360» Abstract (?mno=27927) » PDF (index.php?fulltxt=27927&fulltxtj=196&fulltxtp=196-1604855786.pdf) » doi:
10.31838/srp.2020.10.57 (<http://dx.doi.org/10.31838/srp.2020.10.57>)**Review Article****58. Implementation of Health Promotion Methods for the Prevention of Mother-to-Child HIV Transmission among Pregnant Women: A Literature Review**

Dhesi Ari Astuti, Mohammad Hakimi, Yayi Suryo Prabandari, Ida Safitri Laksanawati, Atik Triratnawati

SRP. 2020; 11(10): 361-366» Abstract (?mno=27940) » PDF (index.php?fulltxt=27940&fulltxtj=196&fulltxtp=196-1604857923.pdf) » doi:
10.31838/srp.2020.10.58 (<http://dx.doi.org/10.31838/srp.2020.10.58>)**Research Article****59. The Effect of Curcumin in Core-Shell Nanoparticle as Therapy in Radiotherapy-Induced Hyposalivation**

Nanda Rachmad Putra Gofur, Aisyah Rachmadani Putri Gofur, Kemal Alif Athallandi, Ayu Anggraini Broto Nagoro,

Soesilaningtyas, Rizki Nur Rachman Putra Gofur, Mega Kahdina, Hernalia Martidal Putri

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10.31838/srp.2020.10.59 (<http://dx.doi.org/10.31838/srp.2020.10.59>)

Review Article

60. Effect of Leadership Style Toward Indonesian Education Performance in Education 4.0 Era: A Schematic Literature Review

Suyudi, Budi Sulistiyo Nugroho, Minnah El Widdah, Aep Tata Suryana, Tatang Ibrahim, Megan Asri Humaira, Moh. Nasrudin, Muhammad Sultan Mubarak, Muhammad Taufiq Abadi, Aprilian Ria Adisti, Silvy Sondari Gadzalia, Muhammad Rikza Muqtada, Agus Purwanto, Mochammad Fahlevi, Yuli Sudargini

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10.31838/srp.2020.10.60 (<http://dx.doi.org/10.31838/srp.2020.10.60>)

Research Article

61. Prediction of Flick Density in the Rainy and Dry Seasons Based on Health Services, Behavior, Environmental Conditions, and Breeding Place in Banjarbaru City Using Partial Least Square

Isnawati, Bambang W. Otok

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10.31838/srp.2020.10.61 (<http://dx.doi.org/10.31838/srp.2020.10.61>)

Review Article

62. A Review of an Important Medicinal Plant: *Alpinia galanga* (L.) Willd

Aswin Rafif Khairullah, Tridiganita Intan Solikhah, Arif Nur Muhammad Ansori, Amaq Fadholly, Sancaka Cashyer Ramandinianto, Ribby Ansharieta, Agus Widodo, Katty Hendriana Priscilia Riwu, Naimah Putri, Annise Proboningrat, Muhammad Khaliim Jati Kusala, Briantono Willy Rendragraha, Akyun Rozaqi Syah Putra, Azharuddin Anshori

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10.31838/srp.2020.10.62 (<http://dx.doi.org/10.31838/srp.2020.10.62>)

Regular Article

63. Developing Model of Halal Food Purchase Intention among Indonesian Non-Muslim Consumers: An Explanatory Sequential Mixed Methods Research

Hery Purwanto, Muchamad Fauzi, Ratna Wijayanti, Khothibul Umam Al Awwaly, Imam Jayanto, Mahyuddin, Agus Purwanto, Mochammad Fahlevi, Hendri Hermawan Adinugraha, Rahmi Andini Syamsudin, Angga Pratama, Nurmin Ariyanto, Denok Sunarsi, Elizabeth Tika Kristina Hartuti, Jasmani

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10.31838/srp.2020.10.63 (<http://dx.doi.org/10.31838/srp.2020.10.63>)

- 64. Effect of Ecological, Servant dan Digital Leadership Style Influence University Performance? Evidence from Indonesian Universities**
Abdul Quddus, Budi Sulistiyo Nugroho, Lukman Hakim, M. Sidi Ritaudin, Een Nurhasanah, Abin Suarsa, Umum Budi Karyanto, Rahman Tanjung, Hendar, Versiandika Yudha Pratama, Husni Awali, Abdul Mufid, Agus Purwanto, Mochammad Fahlevi, Yuli Sudargini
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- 65. The Effect of Double Role Conflict (Work Family Conflict) on Female Worker's Performance with Work Stress as the Intervening Variable**
Helmi Buyung Aulia Safrizal, Anis Eliyana, Kurnia Lail Febriyanti
SRP. 2020; 11(10): 418-428
» Abstract (?mno=29409) » PDF (index.php?fulltxt=29409&fulltxtj=196&fulltxtp=196-1605130453.pdf) » doi: 10.31838/srp.2020.10.65 (<http://dx.doi.org/10.31838/srp.2020.10.65>)
- 66. The Effect of Work-Family Conflict on Job Satisfaction with Organizational Commitment as the Moderator Variable**
Diana, Anis Eliyana, Inanta Indra Pradana
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» Abstract (?mno=29629) » PDF (index.php?fulltxt=29629&fulltxtj=196&fulltxtp=196-1605175168.pdf) » doi: 10.31838/srp.2020.10.66 (<http://dx.doi.org/10.31838/srp.2020.10.66>)
- 67. The Engagement and Working Satisfaction of Millennial Lecturers During the COVID-19 Pandemic: Differences in Gender Identity Perspectives**
Despinur Dara, Anis Eliyana, Hamidah
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» Abstract (?mno=29650) » PDF (index.php?fulltxt=29650&fulltxtj=196&fulltxtp=196-1605177840.pdf) » doi: 10.31838/srp.2020.10.67 (<http://dx.doi.org/10.31838/srp.2020.10.67>)
- 68. The Effect of Leadership Style, Organizational Culture and Job Satisfaction on Employee Performance with Organizational Commitment as the Intervening Variable**
Bambang Raditya Purnomo, Anis Eliyana, Elvina Dyah Pramesti
SRP. 2020; 11(10): 446-458
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- 69. The Effect of Work Family Conflict on Job Performance Through Emotional Exhaustion**
Yos Horta Meliala, Anis Eliyana, Hamidah, Agung Dharmawan Buchdadi, M. Burhanudin Habibi
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- 70. The Effect of Good Corporate Governance and Leadership in Applying Operations Readiness**
Maya Rezeki Angriani, Anis Eliyana, Dyah Sri Wulandari
SRP. 2020; 11(10): 466-471

» Abstract (?mno=29863) » PDF (index.php?fulltxt=29863&fulltxtj=196&fulltxtp=196-1605202318.pdf) » doi:
10.31838/srp.2020.10.70 (<http://dx.doi.org/10.31838/srp.2020.10.70>)

71. Effect of e-Leadership Style, Organizational Commitment and Service Quality towards Indonesian School Performance

Denok Sunarsi, Nani Rohaeni, Retno Wulansari, Jeni Andriani, Ade Muslimat, Zackharia Rialmi, Endang Kustini, Lily Setyawati Kristianti, Dian Rostikawati, Aidil Amin Effendy, Agus Purwanto, Mochammad Fahlevi

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Research Article

72. The Behavioral Effect of Anthocyanin from Purple Sweet Potatoes on Prenatally Stressed Offspring Mice

Nia Kurnianingsih, Retty Ratnawati, Tommy Alfandy Nazwar, Mulyohadi Ali, Fatchiyah Fatchiyah

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» Abstract (?mno=29901) » PDF (index.php?fulltxt=29901&fulltxtj=196&fulltxtp=196-1605206821.pdf) » doi:
10.31838/srp.2020.10.72 (<http://dx.doi.org/10.31838/srp.2020.10.72>)

73. The Application of Geographic Information System for Assessing the Risk Level of Hypertension in Samut Songkhram Province

Jatuporn Ounprasertsuk, Tipvarin Benjanirat, Pongsak Jaroengngarmsamer, Wanich Suksatan, Phannee Rojanabenjakun, Sasipen Krutchangthong, Sunatcha chaowai

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74. Nutritional Value and Acceptability from Drink Probiotic Yogurt with Sago Flour (Metroxylon Sagu Rottb) with Sexual Dysfunction in Postpartum Women

Mustamir Kamaruddin, Anjar Briannita, Sriyanti, Pika Ayu Rahmadewi, Anwar Mallongi

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10.31838/srp.2020.10.74 (<http://dx.doi.org/10.31838/srp.2020.10.74>)

75. The Efficacy of Mindfulness Therapy with Family Centered Care Approach on Mental Recovery of People Living with HIV/AIDS (PLWHA) in Sorong City

Norma, Butet Agustarika, Anwar Mallongi

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76. Ultraviolet Light Application Model in Lowering Germs on Food Snacks at Elementary School in Makassar

Zaenab, Rafidah, Anwar Mallongi, Ashari Rasjid

SRP. 2020; 11(10): 508-513

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10.31838/srp.2020.10.76 (<http://dx.doi.org/10.31838/srp.2020.10.76>)

77. Patient Health Information System for Discharge Planning in Nursing Services in Hospitals

Abdul Aziz Alimul Hidayat, Musrifatul Uliyah, Sukadiono, Nahdah Aulia Aziz Taufiqurrahman

SRP. 2020; 11(10): 514-518» Abstract (?mno=30137) » PDF (index.php?fulltxt=30137&fulltxtj=196&fulltxtp=196-1605253239.pdf) » doi:
10.31838/srp.2020.10.77 (<http://dx.doi.org/10.31838/srp.2020.10.77>)**Review Article****78. Duck Meat, Delicious yet low in Unsaturated Fat; Comparative Study between Duck Meat Consumption in ASEAN toward the Number of COVID Cases**

Maslichah Mafruchati

SRP. 2020; 11(10): 519-523» Abstract (?mno=30148) » PDF (index.php?fulltxt=30148&fulltxtj=196&fulltxtp=196-1605254726.pdf) » doi:
10.31838/srp.2020.10.78 (<http://dx.doi.org/10.31838/srp.2020.10.78>)**Research Article****79. Non-Toxic Fractions of Streptomyces hygroscopicus Subsp. Hygroscopicus Metabolite Suppressed the Growth of Plasmodium Falciparum in Vitro Possibly through L-malate: Quinone Oxidoreductase (PfMQO) Mitochondrial Enzyme Inhibition**

Alfian Wika Cahyono, Loeki Enggar Fitr, Nafisatuzzamrudah, Rivo Yudhinata Brian Nugraha, Rara Aulia, Fitria Febriliani, Dio Giovanni Ariel, Kana Mardhiyyah, Sri Winarsih, Suciati, Dian Japany, Erwahyuni Endang, Danang Waluyo

SRP. 2020; 11(10): 524-531» Abstract (?mno=226) » PDF (index.php?fulltxt=226&fulltxtj=196&fulltxtp=196-1605295619.pdf) » doi:
10.31838/srp.2020.10.79 (<http://dx.doi.org/10.31838/srp.2020.10.79>)**Regular Article****80. Factors Affecting Entrepreneurial Student Intention in Learning Technology**

Bambang Leo Handoko, Septi Wifasari, Fahry Priandhana

SRP. 2020; 11(10): 532-536» Abstract (?mno=339) » PDF (index.php?fulltxt=339&fulltxtj=196&fulltxtp=196-1605305753.pdf) » doi:
10.31838/srp.2020.10.80 (<http://dx.doi.org/10.31838/srp.2020.10.80>)**Research Article****81. In Silico Prediction of Malayan Krait (Bungarus candidus) PLA2 Epitope**

Nia Kurniawan, Coni Anggie Kurniasari, Fatchiyah

SRP. 2020; 11(10): 537-548» Abstract (?mno=475) » PDF (index.php?fulltxt=475&fulltxtj=196&fulltxtp=196-1605329798.pdf) » doi:
10.31838/srp.2020.10.81 (<http://dx.doi.org/10.31838/srp.2020.10.81>)**Regular Article****82. Interactive Teaching Methods as a Change in the Purpose of Modern Education**

Leonid Orshanskyi, Volodymyr Krasnopolskyi, Iryna Fednova, Lesia Vysochan, Tetiana Novalska, Oksana Ivantsiv

SRP. 2020; 11(10): 549-555

» Abstract (?mno=663) » PDF (index.php?fulltxt=663&fulltxtj=196&fulltxtp=196-1605341160.pdf) » doi:
10.31838/srp.2020.10.82 (<http://dx.doi.org/10.31838/srp.2020.10.82>)

Research Article**83. The Effect of Sea Grapes (*Caulerpa cylindrica*) to Gastric Inflammatory Cell Infiltration Score and Catalase Activity in Indomethacin-induced Wistar Rats**

Fitri Handajani, Sulistiana Prabowo

SRP. 2020; 11(10): 556-563

» Abstract & References (?mno=1206) » PDF (index.php?fulltxt=1206&fulltxtj=196&fulltxtp=196-1605381483.pdf) » doi: 10.31838/srp.2020.10.83 (<http://dx.doi.org/10.31838/srp.2020.10.83>)

84. Evaluation of *Agaricus* sp. and *Pleurotus* sp. Extracts Efficiency in *Aspergillus Flavus* Growth Inhibition and Aflatoxin B1 Reduction

Alia Haikal Hussain, Halima Zugher Hussein

SRP. 2020; 11(10): 564-569

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10.31838/srp.2020.10.84 (<http://dx.doi.org/10.31838/srp.2020.10.84>)

85. The Effect of Silver Nanoparticles on BRAF Gene Expression

Gulboy Abdolmajeed Nasir, Mohammed Ayyed Najm, Ammar Lateef Hussein

SRP. 2020; 11(10): 570-575

» Abstract (?mno=2772) » PDF (index.php?fulltxt=2772&fulltxtj=196&fulltxtp=196-1605544267.pdf) » doi:
10.31838/srp.2020.10.85 (<http://dx.doi.org/10.31838/srp.2020.10.85>)

86. Aqueous Extract of *Boswellia* against Rifampicin Toxicity in rats

Amal Umran Mosa, Ban Hoshi khalaf and Salam Ahmed Abed

SRP. 2020; 11(10): 576-583

» Abstract (?mno=2782) » PDF (index.php?fulltxt=2782&fulltxtj=196&fulltxtp=196-1605547529.pdf) » doi:
10.31838/srp.2020.10.86 (<http://dx.doi.org/10.31838/srp.2020.10.86>)

87. Detection of the Toxin Associated Genes of Methicillin-Resistant *Staphylococcus Aureus* Using a Multiplex PCR Assay in Wasit General Hospitals

Rana Hussein Raheema and Ban Hamid Qaddoori

SRP. 2020; 11(10): 584-589

» Abstract (?mno=2796) » PDF (index.php?fulltxt=2796&fulltxtj=196&fulltxtp=196-1605551785.pdf) » doi:
10.31838/srp.2020.10.87 (<http://dx.doi.org/10.31838/srp.2020.10.87>)

Regular Article**88. Internet Addiction among University Students and its Associated Factors: A Cross-Sectional Study among College Students in Hanoi, Vietnam**

Nguyen Thi Mai Lan, Lee Kyesun, Vu Dung, Nguyen Thi Thanh Huyen, Huynh Van Chan, Nguyen Thi Quy, Tran Thu Huong, Nguyen Thi Hoa Mai, Vu Thu Trang, Nguyen Van Hieu

SRP. 2020; 11(10): 590-596

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10.31838/srp.2020.10.88 (<http://dx.doi.org/10.31838/srp.2020.10.88>)

Research Article

89. The Effect of Low-Calorie High Protein Diet on Insulin, TNF- α and P38MAPK Levels in Insulin-Resistant PCOS Mice Models

Hany Puspita Aryani, Budi Santoso, Bambang Purwanto, Sony Wibisono Mudjanarko, Budi Utomo

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» Abstract (?mno=2910) » PDF (index.php?fulltxt=2910&fulltxtj=196&fulltxtp=196-1605591446.pdf) » doi:
10.31838/srp.2020.10.89 (<http://dx.doi.org/10.31838/srp.2020.10.89>)

90. Mother's Height and Calcium Intake Against Stunting among Children Aged 3-5 Years and The Impact on Child Development

Suryana, Andi Eka Yuniato, Yulia Fitri, Silvia Wagustina, Eva Fitriyaningsih, Nunung Sri Mulyani, Sanya Anda Lusiana, I Rai Ngardita

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10.31838/srp.2020.10.90 (<http://dx.doi.org/10.31838/srp.2020.10.90>)

91. Sensory Acceptance and Influence of Pumpkins (*Cucurbita moschata*) Flour in Making Crispy Noodles Toward Primary School Children of Bengkulu

Emy Yuliantini, Kamsiah, Andi Eka Yuniato

SRP. 2020; 11(10): 612-616

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10.31838/srp.2020.10.91 (<http://dx.doi.org/10.31838/srp.2020.10.91>)

Review Article

92. Pharmaceutical Business Competition in Indonesia: A Review

Heri Erlangga, Wa Ode Sifatu, Dimas Wibisono, Ade Onny Siagian, Rudi Salam, Mahnun Mas'adi, Gunartin, Riri Oktarini, Cornelia Dumarya Manik, Nani, Ahmad Nurhadi, Denok Sunarsi, Agus Purwanto, Gatot Kusjono

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Regular Article

93. Health-Promoting Behaviors and Related Factors in Patients with Chronic Diseases in a Rural Community

Wanich Suksatan, Jatuporn Ounprasertsuk

SRP. 2020; 11(10): 624-627

» Abstract (?mno=3477) » PDF (index.php?fulltxt=3477&fulltxtj=196&fulltxtp=196-1605633035.pdf) » doi:
10.31838/srp.2020.10.93 (<http://dx.doi.org/10.31838/srp.2020.10.93>)

Research Article

94. Analysis of Current Models of the Palliative Medical Care at the level of Separate Subjects of the Russian Federation

Lyudmila Alexandrovna Ertel, Marina Filippovna Mikaelyan, Irina Nikolaevna Iyro, Gasbulla Suleymanovich Barkaev, Andrey Borisovich Goryachev, Tatyana Gennadyevna Mogilenko, Stella Vazgenovna Mirzoyan, Madina & 1040;liyevna Garumova

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95. The Identification of Pork Contamination on Beef by Polymerase Chain Reaction (PCR)

Mustofa Helmi Effendi, Shelma Warda Afdilah, Dhandy Koesoemo Wardhana, Fredy Kurniawan, Rurini Retnowati

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96. A Systematic Literature Review of Education Financing Model in Indonesian School

Gunartin, Ade Onny Siagian, Khayatun Nufus, Nur'aini Yusuf, Hadi Supratikta, Ali Maddinsyah, Awaluddin Muchtar, Widya Intan Sari, Denok Sunarsi, Irfan Rizka Akbar, Nurmin Arianto, Agus Purwanto, Noryani, Hadion Wijoyo

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97. The Effect of Adsorbent Composition: Quartz Sand/Andisol Soil/Zeolite/Activated Carbon Against Mn, Fe, BOD, and COD in Citarum River Eater Cleaning Progress

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99. Prevalence of Refractive Errors Among Primary School Children (6-15 Yrs) In Al-Khartoum- Sudan

Noaman Mukbel Ghalib, Samira Mohammed Ibrahim, Nasraddin Othman Bahakim

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100. Development of Performance Appraisal Sustainable Theory of Sharia Banks in Indonesia

Ahmad Sodik, Ririn Tri Ratnasari, Imron Mawardi

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101. Effectiveness of Fucoidan Extract from Brown Algae to Inhibit Bacteria Causes of Oral Cavity Damage

Nurlindah Hamrun, Sri Oktawati, Asmawati, Irene, Hardianti Maulidita Haryo, Ira Farwiany Syafar, Andi Nurazizah Almaidah

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102. Water and Microbial Contents in Moringa Oleifera Seed Flour as Food Supplement to Prevent Stunting

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103. Formation of Learning Research Skills Through Solving Arithmetic Problems

Rabiga I. Kenzhebekova, Saule S.Kozhadeldiyeva, Kulahmet Moldabek, Louise A. Rizaeva, Kenzhegul U. Kazybayeva

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104. Coronavirus Disease (COVID-19): A Regional Autonomy Point of View

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105. The Problem Of Health Quality Of Medical Personnel In The Context Of Social Reform

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107. Nutritional Education Model Through Crossword Puzzles Toward Knowledge And Macro Nutrient Intake Of Primary School Student In Bengkulu City

Kamsiah, Emy Yuliantini, Andi Eka Yunianto

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109. Why Do People Fail to Comply with the Smoking Ban in Public Places? (The Case of Jayapura City, Indonesia)

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Review Article**111. AITCS as a Reliable Instrument for Evaluating IPC (Interprofessional Collaboration): A Systematic Review**

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114. The Hepatoprotection Effect of the Asymmetric Curcumin Analogue Synthetic Product in Male Rat Abstract (Rattus norvegicus L.)

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115. Pharmacological Evaluation of Antidiabetic Activity of Chromolaena Odorata Leaves Extract in Streptozotocin-Induced Rats

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10.31838/srp.2020.10.118 (<http://dx.doi.org/10.31838/srp.2020.10.118>)**Review Article****119. Broiler Chicken vs. Turkey Meat; which One Has the Least Bad Fat to Avoid Positive Case of COVID-19?**

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10.31838/srp.2020.10.119 (<http://dx.doi.org/10.31838/srp.2020.10.119>)**Regular Article****120. Protection of the Right to Information on One's Health by Authorized State Bodies**

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10.31838/srp.2020.10.120 (<http://dx.doi.org/10.31838/srp.2020.10.120>)**Review Article****121. Recent Updates on COVID-19 Vaccine Platforms and Its Immunological Aspects: A Review**

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10.31838/srp.2020.10.121 (<http://dx.doi.org/10.31838/srp.2020.10.121>)**122. A Review of Salmonella sp. in Tilapia fish (Oreochromis niloticus) : Public Health Importance**

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10.31838/srp.2020.10.122 (<http://dx.doi.org/10.31838/srp.2020.10.122>)**Regular Article****123. Subjects of the Right to Information on One's Health in Private and Public Law**

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10.31838/srp.2020.10.124 (http://dx.doi.org/10.31838/srp.2020.10.124)**125. Mechanism of Bone Metabolism Interruption Due to High Intensity Physical Exercise**

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10.31838/srp.2020.10.125 (http://dx.doi.org/10.31838/srp.2020.10.125)**126. A Situation Analysis on the Use of Standardized-Herbal Medicines as Supportive Therapies for Dengue Hemorrhagic Fever (DHF) patients in Indonesia**

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10.31838/srp.2020.10.126 (http://dx.doi.org/10.31838/srp.2020.10.126)**127. Overweight and OBES: The Impact of Lactobacillus Casei Strain Shirota on Cholesterol and Weight of Children in the Pandemic Time COVID-19**

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10.31838/srp.2020.10.129 (http://dx.doi.org/10.31838/srp.2020.10.129)**Review Article**

130. Microorganism-Based β -Glucan Production and their Potential as Antioxidant

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10.31838/srp.2020.10.130 (http://dx.doi.org/10.31838/srp.2020.10.130)**Regular Article****131. Did PDCA Cycle, Service Quality and Innovation Capability Influence Private Universities Performance?**

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10.31838/srp.2020.10.131 (http://dx.doi.org/10.31838/srp.2020.10.131)**Research Article****132. The Effect of Gel *Abelmoschus Manihot* (L.) Medik Leaf Extract on Second Degree Burn Wound Healing Process**

Wiwik Misaco Yuniarti, Erva Fatmareta, Laila Nur Hidayati, Nadhila Arieska, Bambang Sektiari Lukiswanto

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10.31838/srp.2020.10.132 (http://dx.doi.org/10.31838/srp.2020.10.132)**Regular Article****133. The Role and Place of Information and Communication Technologies in the Formation of Professional Competencies of Higher Education**

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10.31838/srp.2020.10.133 (http://dx.doi.org/10.31838/srp.2020.10.133)**134. Effect of ISO 9001, ISO 45001 and ISO 14000 toward Financial Performance of Indonesian Manufacturing**

Mukhlisotul Jannah, Mochammad Fahlevi, Julinta Paulina, Budi Sulistiyo Nugroho, Agus Purwanto, Milana Abdillah Subarkah, E. Kurniati, Teguh Setiawan Wibowo, Kasbuntoro, Nawang Kalbuana, Yoyok Cahyono

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10.31838/srp.2020.10.134 (http://dx.doi.org/10.31838/srp.2020.10.134)**135. Sustainable, Pro-Poor and Humane Waqf Management: A Literature Study of Evidence-Based Success Criteria of Waqf Managers**

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136. **Anti-Corruption Policy under the Conditions of Overcoming the Consequences of the Coronavirus Pandemic**
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137. **Modern Problems of Medical and Pharmacological Education**
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138. **The Impact of Using Interactive Boards on Secondary School Students' Learning Outcomes in Egypt in the Computer Skills Course**
Mohammed Elmetwali Mohammed Amer
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10.31838/srp.2020.10.138 (<http://dx.doi.org/10.31838/srp.2020.10.138>)
139. **Protection of the Rights of Public Utilities Market Participants (In the Context of Poverty Prevention)**
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140. **Molecular Detection of Parvovirus B19 and Immunohistochemical Localization of Interleukin 6 of Tissues from Thyroid Cancer**
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10.31838/srp.2020.10.140 (<http://dx.doi.org/10.31838/srp.2020.10.140>)
141. **Isolation and Diagnosis of Bacteria from Women with Urinary Tract Infection and Study of Antibiotic Susceptibility**
Rafal Ahmed Lilo, Dalal mohammed, zahraa M. Al-Taee, Zeena Hadi Obaid Alwan and Rajaa Mahmoud Ibrahim AL-Jasim
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142. **Legal Reforms of the Land Market in the Countries of Central and Eastern Europe and the Development of Their Financial and Credit Institutions**

Oleksandr Tereshchuk, Mykola Yasynok, Volodymyr & 1050;roitor, Alla Zemko, Dmytro Yasynok, Mykola Danevych

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143. The Effectiveness of Giving Cork Fish Nugget to Increase Endurance of Toddlers at Community Health Centers

Sentani Jayapura Regency

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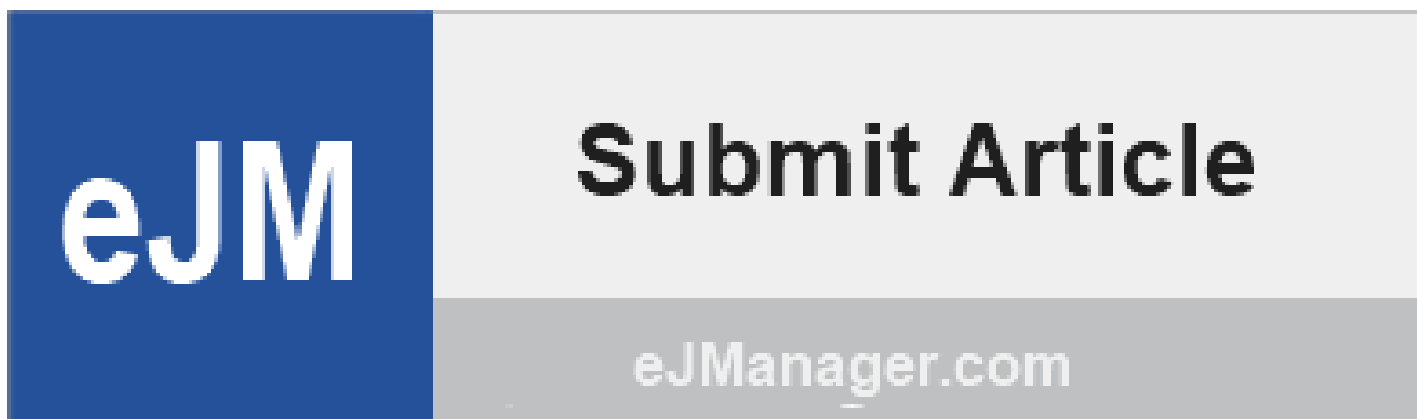
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In Silico Study on Antibacterial Activity and Brazilein ADME of Sappan Wood (*Caesalpinia Sappan* L.) Against *Escherichia coli* (Strain K12)

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ABSTRACT

This study aims to perform in silico analysis for revealing the antibacterial activity of brazilein compounds contained in sappan wood against the ESBL enzyme derived from *Escherichia coli* multiresistant isolates of UTI patients together with the antibacterial activity of meropenem as the comparative compound. The in silico test was conducted to predict antibacterial activity by docking using the Molegro Virtual Docker computer program. The receptor used was beta-lactamase AmpC, PDB code: 1LL5, together with an imipenem ligand (IM2-370). In addition to predicting the antibacterial activity, this study also aims to predict physicochemical and pharmacokinetic properties (ADME) as well as toxicity of brazilein and meropenem with the pkCSM online tool program. Next, the data obtained were analyzed by comparing the docking bond energy between brazilein, imipenem ligand, and meropenem with the target receptor. The lower the ligand bond energy than the target receptor is, the more stable the bond is formed. Hence, this can be used to predict the biological activity of compounds. In silico test results showed that the bond energy of brazilein was -77.4202 kcal/mol, -98.2425 kcal/mol for meropenem, and -85.8187 kcal/mol for imipenem. Brazilein has potential as an antimicrobial even though it is lower than meropenem and imipenem. Based on the results of in silico test using the pkCSM online tool program, the brazilein compound also has good pharmacokinetic properties causing relatively low toxicity.

Keywords: ADME, AmpC beta-lactamase, brazilein, meropenem, in silico test

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INTRODUCTION

Urinary tract infection (UTI) is an infection caused by microorganisms in the urinary tract, which starts from an infection in the urinary tract, and then infects the genitalia and even the kidneys[1]. *Escherichia coli* is a gram-negative bacteria in the Enterobacteriaceae group that dominates the cause of UTI. *Escherichia coli* isolated from urine cultures are generally Multiple Drug Resistant (MDR), but are still sensitive to the antimicrobials of carbapenem and aminoglycosides. Carbapenem antimicrobials, nevertheless, are still considered as the last option in treatments of infection cases caused by *Escherichia coli* multiresistant[2].

MDR, moreover, usually occurs because of the resistant coding gene in the plasmid, so the bacteria can produce the enzyme Extended Spectrum Beta-Lactamase (ESBL). ESBL is an enzyme that has an ability to hydrolyze the antibiotics of penicillin, cephalosporins of generation I, II, and III, as well as aztreonam[3]. Besides, ESBL is most widely produced by Enterobacteriaceae, especially *Escherichia coli* and *Klebsiella pneumoniae*. ESBL is also derived from the mutated beta-lactamase enzyme, triggering an increase in the enzymatic activity of beta-lactamase so that this enzyme can hydrolyse generation III and IV cephalosporins and aztreonam[4]. Furthermore, the Canadian External Quality Assessment Advisory Group for Antibiotics states that genes controlling the production of beta-lactamase are located in the plasmid or chromosome. This facilitates the ability of the ESBL gene

to move from one organism to another so that the spread of resistance is very easy between strains and even between species. Plasmids are also responsible for coding genes that carry resistance genes for other classes of drugs, such as aminoglycosides. This situation makes the choice of antibiotics against organisms that produce ESBL very limited[5].

Secang wood (*Caesalpinia sappan* L.) is one of the medicinal plants in Indonesia that can act as an antibacteria, antioxidant, and anti-inflammatory[6,7]. Secang wood contains five active compounds related to flavonoids. The five compounds are brazilin, brazilein, 3'-O-methylbrazilin, sappanin, chalcone, and sappanchalcone. Brazilein is a major compound contained in sappan wood. Secang wood, according to some studies, reveal some benefits of sappan wood, such as inhibiting Mesenchymal Stem Cells Senescence[8], reducing liver damage level caused by excessive amounts of iron[9], increasing motility, percentage viability, and sperm concentration of male wistar rats[10], providing antigenotoxic effects on exposure to cyclophosphamide mutagen compounds based on micronucleus assays[11], increasing cytotoxicity and induction of apoptosis in T47D cells[12], inhibiting the activity of rheumatoid arthritis in rats induced by collagen arthritis type-II[13], and inhibiting catabolic processes in human osteoarthritic chondrocytes through inhibition of NFκB1 / p50[14]. Hence, to determine the activity of brazilein as an antibacterial MDR, it is necessary to perform an in silico

test with a computer program, both offline and online. In silico test is a test approach to predict physicochemical properties of molecules, pharmacokinetic properties (absorption, distribution, metabolism and excretion = ADME) of molecules, interaction of compounds with receptors, prediction mechanism of action, as well as selectivity and toxicity of compounds. This test also has several advantages, such as safe, free from chemical waste, easy, cost-effective, and fast.

In addition, as a comparison of the potential of brazilein against *Escherichia coli* MDR, a sensitive antibiotic, meropenem, was used. For ligands or molecules that have shown good biological activity and are able to bind the desired biological targets (receptors) (docking process) on protein data banks are imipenem (PDB ID: 1LL5)[15,16].

Brazilein secang wood actually can be developed in a drug design as an antibacterial drug. Drug design is often described as a systematic elaboration process to further develop existing drugs with the aims of obtaining new drugs with better activity and reducing or eliminating side effects through molecular manipulation. Molecular manipulation or structural modification is to synthesize a number of parent strains, identify structures, and test their biological activities. Changes in the structure of a compound can change the physicochemical properties of compounds, including lipophilic, electronic and steric properties, and also cause changes in the biological activity of compounds. To be more effective and efficient in modifying the structure, the physicochemical and pharmacokinetic properties (ADME) as well as toxicity of an synthesized compound should be predicted first to get the picture of drug interactions with receptors. As a result, this study aims to reveal the potential of brazilein as an antibacterial MDR using in silico method[17].

MATERIALS AND METHOD

Materials

The 3-dimensional (PDB ID: 1LL5) X-ray crystal structure of AmpC WT beta-lactamase in complex with covalently bound imipenem was downloaded from <http://www.rcsb.org/pdb/home.do>. Next, the 3-dimensional structures of brazilein, meropenem, and imipenem were downloaded from <https://pubchem.ncbi.nlm.nih.gov/compound/>.

Tools

A set of computers with Windows 8 64 bit specifications and ChemDraw Professional 16.0, Chem3D 16.0, and Molegro Virtual Docker 5 programs.

Procedure

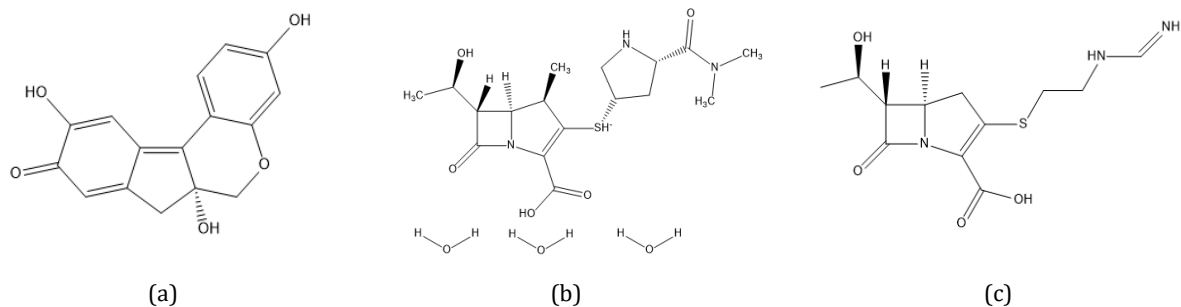


Figure 1. The 2D structures of (a) Brazilein; (b) Meropenem; (c) Imipenem compounds

Activity prediction (molecular docking)

Compounds to be docked are brazilein, meropenem, and imipenem which 2-D structures were drawn using ChemDraw Professional 16.0., and then converted to the 3-D ones using Chem3D 16.0. determined as the most stable conformation. After measuring their minimum energy, they then were stored in the form of mol2 {SYBYL2 (*. Mol2)}. Meanwhile, the structure of the *Escherichia coli* ESBL protein (PDB ID: 1LL5) was obtained from the Protein Data Bank. The results obtained were in the form of Rerank Score (RS), namely energy required in the process of ligand-receptor interaction, and then based on these values the antibacterial activity of secang wood (*Caesalpinia sappan L.*) against *Escherichia coli* ESBL could be predicted.

Prediction of physicochemical and pharmacokinetic properties as well as toxicity of compounds (pkCSM)

Prediction of physicochemical properties, such as: Molecular Weight (BM), logarithm of octanol / water partition coefficient (Log P), number of bonds between rotating atoms (Torsion); Hydrogen Bond Acceptors (HBA), Hydrogen Bond Donors (HBD), and Polar Surface Activity (PSA), were performed using the pkCSM online tool. Similarly, prediction of pharmacokinetic properties (ADME: absorption, distribution, metabolism and excretion) as well as toxicity of brazilein, meropenem, and imipenem compounds were conducted using the pkCSM online tool. First, the 2-D molecular structures of brazilein, meropenem, and imipenem compounds were drawn with the ChemDraw Professional 16.0 program, and then copied in the Chem3D 16.0 program to make their 3-D structures before stored in *.sdf files. Second, the structures of brazilein, meropenem, and imipenem were translated into the SMILES format using the Online SMILES Translator (<https://cactus.nci.nih.gov/translate/>). In the SMILES format, those compounds then were processed using the online pkCSM tool (<http://biosig.unimelb.edu.au/pkcsm/prediction>) to predict ADME and compound toxicity. Afterwards, to predict oral toxicity (LD50) in the globally harmonized system (GSH), Protox online tool was also used (<http://tox.charite.de/tox/>)[17,18].

RESULTS AND DISCUSSION

The 2-D structures made with ChemDraw Professional 16.0 were shown in (Figure 1). Next, the 2D structures then were altered into the 3-D ones with Chem3D 16.0. The 3D structures then were used at all docking stages as illustrated in (Figure 2).

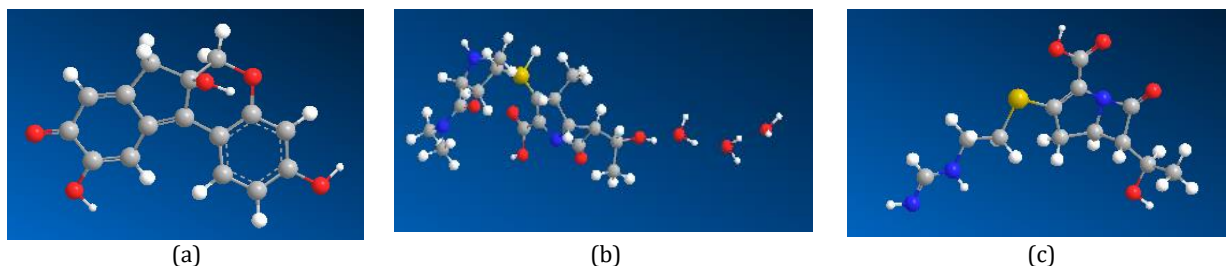


Figure 2. The 3-D structures of (a) Brazilein; (b) Meropenem; (c) Imipenem compounds stored in SYBYL2

Activity prediction with docking and amino acid analysis

Protease receptors already downloaded in Protein Data Bank with code 1LL5 and then imported in the Molegro Virtual Docker program were depicted in (Figure 3). The detection results of the place where ligands and receptors

(cavity) were interacting on the Protease 1LL5 receptor [A] were shown in (Figure 4). The Cavity used was Cavity 1 (volume 433,152) with active Ligand IM2_370 [A] since it has an area where the original ligand interacts with the protease enzyme.

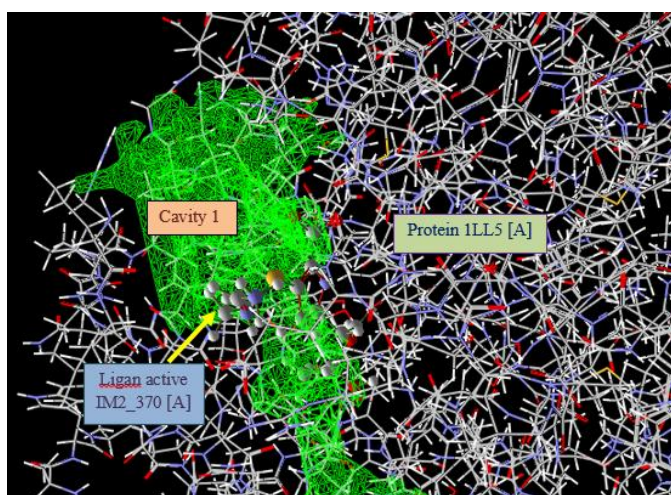


Figure 3. Protease Receptors (PDB code 1LL5) with cavity 1 (volume 433,152) and active Ligand IM2_370 [A]

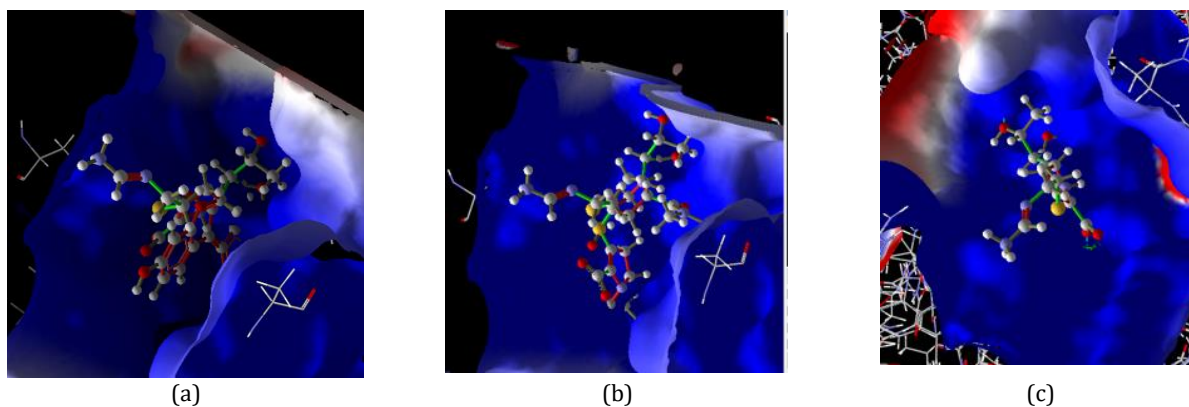


Figure 4. Cavity 1 and Ligan (a) Brazilein; (b) Meropenem; (c) Imipenem

In the interaction of the ligand and the receptor, there is also an interaction of the ligand with some amino acid residues derived from Protease receptor 1LL5. Amino acids involved in the process of interaction of brazilein, meropenem and imipenem compounds with Protease receptors 1LL5 can be seen in Figure 5 and Table 1. The interactions of amino acid receptor residues with

compounds occur through lipophilic / hydrophobic, electronic, and steric receptors. In (Figure 5) and Table 1, it can actually be seen the differences of interactions between each brazilein, meropenem and imipenem compounds with Protease receptor 1LL5 since there are differences in the spatial configuration of the structure of the three compounds.

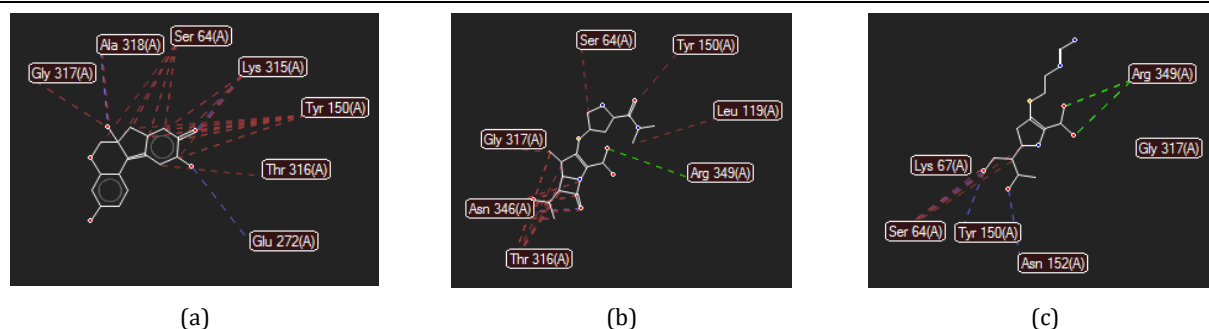


Figure 5. Interactions of Ligand and 1LL5 receptor [A] (H-Bond, electronic, steric) (a) Brazilein; (b) Meropenem; (c) Imipenem

Table 1. Interactions of ligands and 1LL5 receptors [a] (h-bond, electronic, steric)

Ligand	The Bonding of Hydrogen and Amino Acid Residues		Interactions of Electrostatics and Amino Acid Residues		Interactions of Steric and Amino Acid Residues	
	Count	Residues	Count	Residues	Count	Residues
Brazilein	3	Ala 318(A) Lys 315(A) Glu 272(A)	0	-	7	Ala 318(A) Gly 317(A) Ser 64(A) Tyr 150(A) Lys 315(A) Glu 272(A) Thr 316(A)
Meropenem	1	Asn 346(A)	1	Arg 349(A)	6	Gly 317(A) Thr 316(A) Asn 346(A) Ser 64(A) Tyr 150(A) Leu 119(A)
IM2_370[A]	5	Lys 67(A) Ser 64(A) Tyr 150(A) Asn 152(A) Arg 349(A)	1	Arg 349(A)	6	Lys 67(A) Ser 64(A) Tyr 150(A) Asn 152(A) Arg 349(A) Gly 317(A)

The re-docking results of brazilein, meropenem and imipenem compounds with 1LL5 Protease receptors can be seen in Table 2. The binding energy of brazilein with 1LL5 receptors [A] was higher than that of meropenem and imipenem. The results also showed that the rerank score of Brazilein was -77.4202 kcal / mol, -98.2425 for

the rerank score of meropenem, and -85.8187 for the rerank score of imipenem. Based on those rerank scores, brazilein provided higher energy than meropenem and imipenem, so it was less stable in binding to receptors than meropenem and imipenem.

Table 2. Results of re-docking using the molegro virtual docker batchjob program

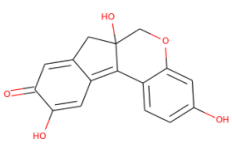
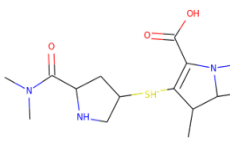
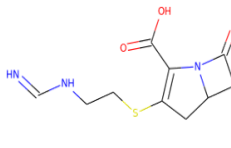
File name	Ligand	Rerank Scores
[02]Brazilein.mvdml	Brazilein	-77.4202
[03]Meropenem.mvdml	Meropenem	-98.2425
[00]IM2_370[A].mvdml	Imipenem	-85.8187

Prediction of physicochemical and pharmacokinetic properties as well as toxicity of compounds (pkCSM)

The in silico prediction results of the physicochemical properties of brazilein, meropenem, and imipenem compounds can be seen in Table 3. Based on a study conducted by Lipinski *et al.* (1997) on 2,245 drugs from the World Drugs Index baseline data argues that the compound would be difficult to absorb and its permeability would be low if the molecular weight is greater than 500, the log coefficient value of octanol /

water (log P) is greater than +5, it has a donor H-bond (HBD), expressed in terms of O-H and N-H groups, greater than 5, or it has an H-acceptor (HBA) bond, expressed by the number of O and N atoms, greater than 10. The analysis above is known as the Lipinski's five law since all values are multiples of five[19]. Table 3 then indicates that brazilein, meropenem, and imipenem have molecular weights less than 500, logP values less than 5, as well as acceptor and donor values less than 10. Hence, it can be concluded that the three compounds are easily absorbed.

Table 3. In silico value predictions of the physicochemical properties of brazilein, meropenem, and imipenem compounds

SMILES Structures			
	Brazilein	Meropenem	Imipenem
BM	284.267	384.478	299.352
LogP	1.624	-1.2263	-0.1760
Torsion	0	5	7
HBA	5	6	5
HBD	3	3	4
PSA (A ²)	119.652	156.137	120.082

Note: BM = Molecular Weight; Log P = logarithm of octanol / water partition coefficients; Torsion = a bond between atoms that can rotate; HBA = Hydrogen Bond Acceptors; HBD = Hydrogen Bond Donors; PSA = Polar Surface Activity.

Afterwards, the in silico prediction results of the pharmacokinetic properties (ADME) and toxicity of

Brazilein, Meropenem, and Imipenem compounds can be seen in Table 4.

Table 4. The in silico predictions of the pharmacokinetic properties (ADME) and toxicity of Brazilein, Meropenem, and Imipenem compounds

ADMET	Brazilein	Meropenem	Imipenem
Intestinal absorption (human) (%)	92.111	33.227	37.266
Skin Permeability (log Kp)	-3.443	-2.735	-2.735
VDss (human) (log L/kg)	0.154	-0.951	-1.283
BBB permeability (log BB)	-0.709	-0.739	-1.139
CYP2D6 substrate (Yes/No)	No	Yes	Yes
CYP2D6 inhibitor (Yes/No)	No	No	No
Total Clearance (log ml/min/kg)	0.264	0.629	0.72
Renal OCT2 substrate (Yes/No)	No	No	No
AMES toxicity (Yes/No)	Yes	No	No
LD50 (mol/kg)	2.024	1.984	1.785

Note: VDSS: Steady State of Volume Distribution, BBB: Blood Brain Barrier, CYP2D6: Cytochrome P2D6, Renal OCT2: Renal Organic Cation Transporter 2

According to Chander *et al.* (2017), a compound can be considered to have a good absorption if its absorption value is > 80%; meanwhile its absorption is considered to be bad if it is <30%. Besides, intestine is considered as the main organ absorbing drugs given orally[20]. Based on Table 4 it can be seen that the intestinal absorption (human) value of brazilein, meropenem, imipenem compounds is more than 80% and not less than 30%. It indicates that the three compounds have good absorption. Moreover, according to Pires *et al.* (2015), a compound is considered to have relatively low skin permeability if it has a log value of Kp > -2.5. Based on Table 4, the values of Skin Permeability (log Kp) of brazilein, meropenem, and imipenem compounds ranged from -2.7 to -3.4, not less than -2.5. It means that all three of these compounds have good skin permeability[18].

Furthermore, volume distribution (VDSS) is a theoretical volume indicating the total dose of a drug distributed evenly to give the same concentration as in blood plasma. The higher the VD value is, the more drugs are distributed in the tissue rather than plasma. According to Pires *et al.* (2015), a compound is considered to have a low Volume Distribution when the Log VD value was <-0.15, and considered to have a high one if the Log VD value was > 0.45. Based on Table 4, the VDss (Steady State of Volume Distribution) value of brazilein compounds was 0.154, -

0.951 for meropenem, and -1.283 for imipenem. Therefore, it can be said that all the derivatives of these compounds can be distributed evenly to give the same concentration as in blood plasma.

The ability of drugs to penetrate the blood brain barrier can actually be considered as an important parameter in reducing side effects and toxicity or in increasing the efficacy of drugs which pharmacological activity is in the brain. Brain-blood permeability measured in vivo in animal models then can be considered as logBB, which is the ratio of logarithmic concentrations in the brain to plasma.

In addition, according to Pires *et al.* (2015), a compound is said to be able to penetrate the blood brain barrier well if it has a Log BB value of > 0.3; meanwhile, it cannot be able to penetrate the blood brain barrier well if the BB log value is <-1. Based on Table 5, the BB log value of the brazilein compound was -0.709, -0.739 for meropenem, and -1.139 for imipenem. It indicates that all the derivatives of these compounds are able to penetrate the blood brain barrier in a moderate manner since the BB log value of each compound was greater than -1.

Besides, it is generally known that most metabolic reactions will involve oxidation processes. Cytochrome P450 is an important detoxification enzyme in the body, and mainly found in the liver. Cytochrome P450 can

oxidize foreign organic compounds, including drugs, as well as facilitate the excretion of these compounds. On the other hand, enzyme inhibitors, such as grapefruit juice, can affect the metabolism of the drug, contraindicating against the cytochrome P450 enzyme. Consequently, it is important to assess the ability of compounds that can inhibit cytochrome P450, represented by cytochrome CYP3A4 isoforms in this study. Besides, based on Table 4, brazilein, meropenem, and imipenem compounds are known not to be able to affect or inhibit the CYP3A4 enzyme. As a result, it can be predicted that these compounds tend to be metabolized by the P450 enzyme. To predict the compound excretion process, furthermore, the Total Clearance (CLTOT) and Renal Organic Cation Transporter 2 (OCT2) substrate were measured. CLTOT is a combination of hepatic clearance (metabolism in the liver and bile) and renal clearance (excretion through the kidneys). This is related to bioavailability, which is important to determine the dose level in achieving steady-state concentration. Based on Table 4, the CLTOT value of brazilein was 0.264, 0.629 for meropenem, and 0.72 for imipenem. Based on these values, the speed of excretion of compounds then can be predicted.

Organic Cation Transporter 2 is a transporter in the kidney which plays an important role in the disposition and clearance of drugs and endogenous compounds. OCT2 substrate also has the potential to cause side interactions if given together with OCT2 inhibitors. Based on Table 4, it then can be seen that the three compounds did not affect the OCT2 substrate. Hence, it can be predicted that the derivative is not an OCT2 substrate.

Subsequently, to determine the toxicity of a compound, Ames Toxicity test was carried out. The Ames Toxicity Test is a widely used method for assessing the mutagenic potential of compounds using bacteria. Positive test results indicate that the compound is mutagenic and can therefore act as a carcinogen. Based on Table 4, it then can be said that brazilein is assumed to cause mutagenic effects. Meanwhile, meropenem and imipenem are assumed not to cause mutagenic effects.

In addition, the results showed that brazilein bond energy is higher than meropenem and target receptors. It means that brazilein has potential as an antimicrobial. But, the in silico molecular docking value of brazilein was lower than that of meropenem and target receptors. Thus, it can be said that the physicochemical and pharmacokinetic properties as well as toxicity of brazilein compounds is assumed to be very well absorbed in the intestine since it can have good skin permeability, can be distributed evenly to provide the same concentration as in blood plasma, can be able to penetrate the blood-brain barrier moderately, can be metabolized by enzyme P450, can have a relatively low toxicity, and can have the greatest cytotoxic activity using the pkCSM online tool.

CONCLUSION

Finally, it can be concluded that Brazilein bond energy is higher than meropenem and target receptors. Based on the binding energy values, moreover, brazilein has potential as an antimicrobial, but its in silico molecular docking value is lower than meropenem and target receptors. It means that the physicochemical and pharmacokinetic properties as well as toxicity of brazilein compounds are assumed to be very well absorbed in the intestine, has good skin permeability, be distributed evenly to provide the same concentration as in blood plasma, be able to penetrate the blood-brain barrier

moderately, be metabolized by enzyme P450, has a relatively low toxicity, and have the greatest cytotoxic activity using the pkCSM online tool.

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AUTHORS CONTRIBUTIONS

Dwi Krihariyani is currently pursuing doctoral education, Eddy Bagus Wasito as promoter, Isnaeni Isnaeni as co-promoter, Siswandono Siswodihardjo, Wiwik Misaco Yuniarti, Entuy Kurniawan as revising it critically for important intellectual content; and final approval of the version to be published.

CONFLICT OF INTERESTS

The authors confirm that this article content has no conflicts of interest

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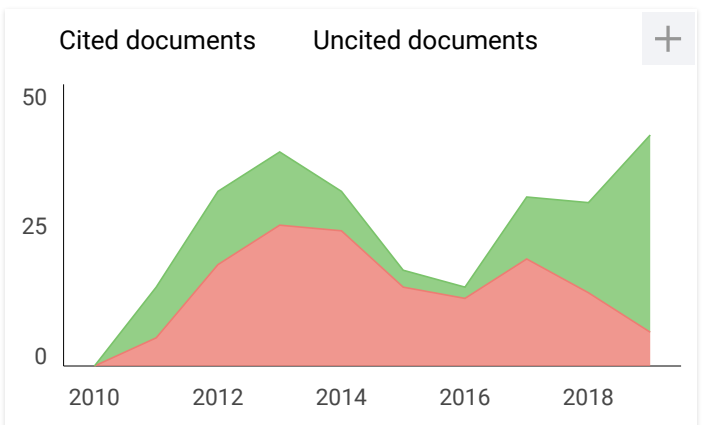
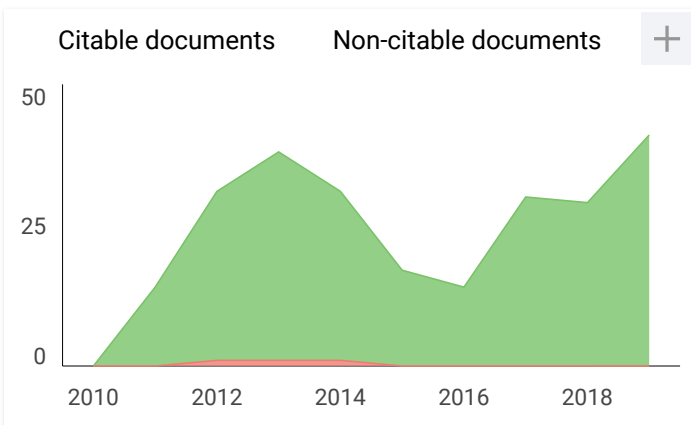
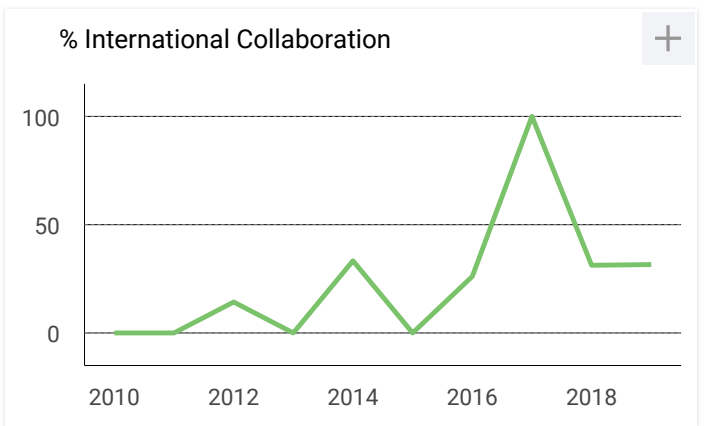
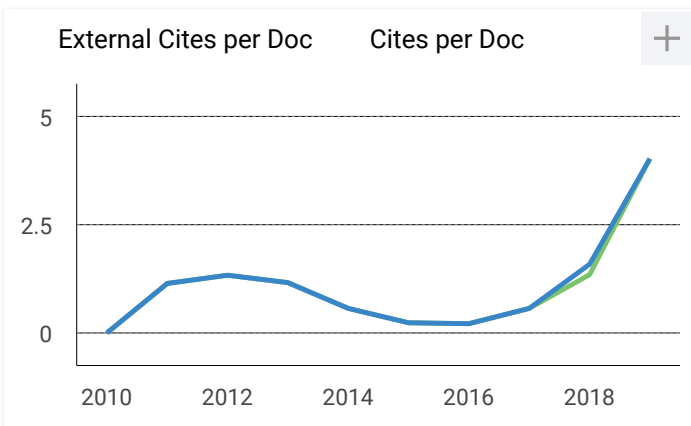
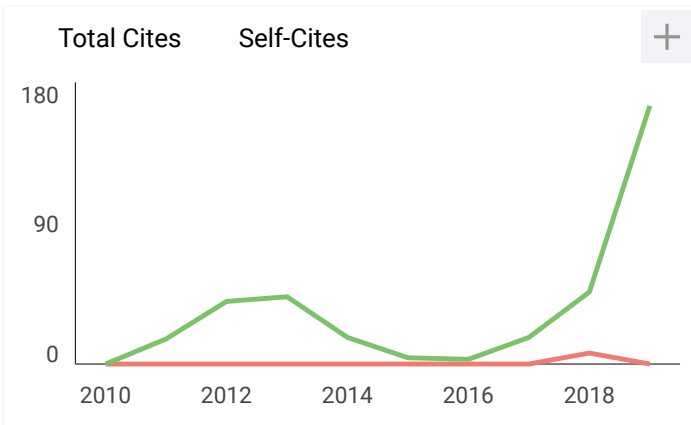
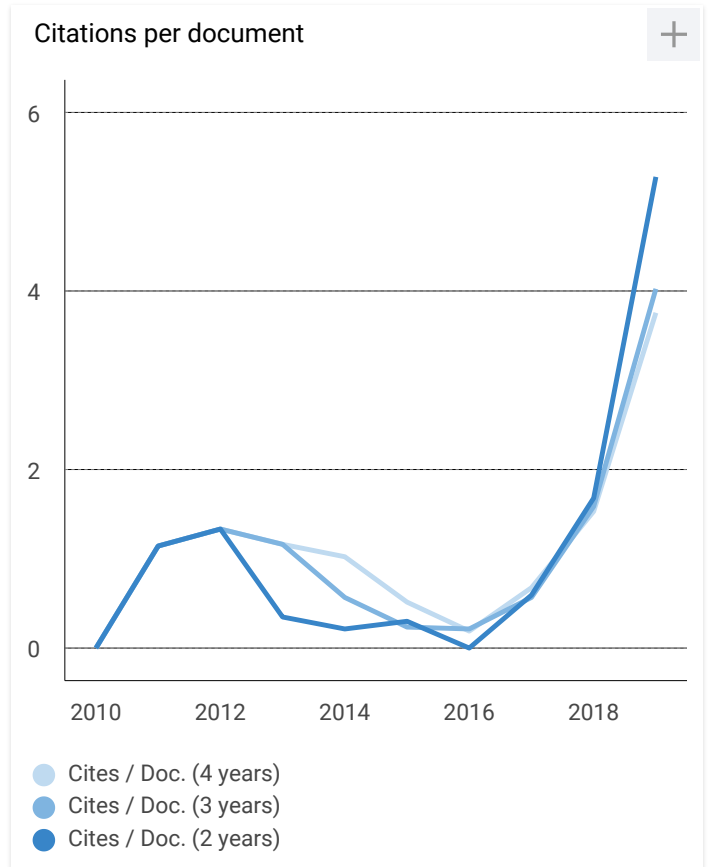
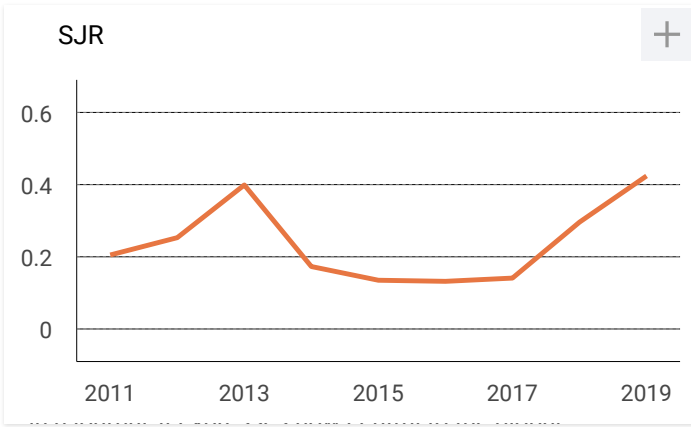
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