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PROFIL FITOKIMIA DAN GC-MS RESIN *DRYOBALANOPS KEITHII*

Phytochemical And Gc-Ms Profile Of Dryobalanops Keithii Resin

Oleh:

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ABSTRAK

Dryobalanops keithii sebagai salah satu jenis dipterokarpa yang ditemukan di Kalimantan Timur, secara tradisional berfungsi sebagai obat luka dalam. Namun resin *Dryobalanops keithii* hingga saat ini belum diketahui kandungan kimianya. Penelitian bertujuan untuk mengetahui kandungan resin *Dryobalanops keithii* menggunakan analisa fitokimia dan GC-MS. Uji fitokimia yang dilakukan antara lain uji alkaloid, flavonoid, saponin, tannin, terpenoid dan steroid. Uji GC-MS menggunakan Shimadzu QP 2010, tipe kolom RTx-5MS. Hasil uji fitokimia dari resin kapur *D.keithii* yang larut dalam etanol 96% mengandung alkaloid dan terpenoid. Hasil uji GC-MS mendeteksi 29 senyawa kimia Senyawa 2-[4-Cyclohexylbutanoylamino]-3-chloro-1,4-naphthoquinone (C₂₀H₂₂ClNO₃) sebanyak 0,99% tergolong alkaloid. α -Pinene, β -pinene, humulene dan γ -gurjunene merupakan contoh senyawa terpenoid.

Kata kunci: Dryobalanops keithii, resin, fitokimia, GC-MS

ABSTRACT

Dryobalanops keithii, one of the dipterocarp species found in East Kalimantan, has traditionally functioned as internal wounds medicine. Unfortunately, the chemical content of *Dryobalanops keithii* resin is unknown. The study aimed to determine the resin content of *Dryobalanops keithii* using phytochemical and GC-MS tests. Phytochemical tests carried out alkaloids, flavonoids, saponins, tannins, terpenoids, and steroids. GC-MS test using Shimadzu QP 2010, column type RTx-5MS. Phytochemical test results of *D. keithii* lime resin, which is soluble in 96% ethanol, contain alkaloids and terpenoids. The results of the GC-MS test detected 29 chemical compounds. Compounds 2-[4-Cyclohexylbutanoylamino]-3-chloro-1,4-naphthoquinone (C₂₀H₂₂ClNO₃) as much as 0,99% were classified as alkaloids. α -Pinene, β -pinene, humulene, and γ -gurjunene are terpenoid compounds.

Keywords: *Dryobalanops keithii*, resin, phytochemical, GC-MS

I. PENDAHULUAN

Pulau Kalimantan memiliki hutan hujan tropis yang didominasi oleh family Dipterocarpaceae (Eni et al., 2018). Jenis Dipterokarpa dapat dimanfaatkan baik kayunya maupun non kayu, seperti resin (Purwaningsih, 2004). Resin pada jenis dipterokarpa dapat dibedakan menjadi *solid resin* dan *oleoresin* (Evans et al., 2003). *Solid resin* berbentuk padatan yang rapuh dengan warna yang spesifik. *Oleoresin* berwujud cair dan memiliki aroma yang khas. Resin/oleoresin didapatkan apabila kulit batang atau kayunya dilukai (Evans et al., 2003; Purwaningsih, 2004).

Senyawa kimia yang dimiliki dari tiap jenis Dipterokarpa berbeda-beda dan khas. Contohnya adalah dari jenis *Shorea javanica* yang salah satu produk unggulannya adalah damar mata kucing dengan senyawa kimia *germacrene*, *vulgarol B* dan lain-lain (Gusti, 2014; Yunanta et al., 2014). Sedangkan produk dari *Dryobalanops aromatica* adalah kamfer dan parfum, serta diketahui memiliki senyawa seperti *borneol*, *caryophyllene*, dan lain-lain (Aswandi & Kholibrina, 2020; Pasaribu et al., 2014).

Genus *Dryobalanops* (Kapur) merupakan salah satu bagian dari family Dipterocarpaceae (Dayanandan et al., 1999) juga memiliki resin. *Dryobalanops keithii* adalah salah satu jenis

jenis kapur yang ditemukan di Kalimantan Timur. Berdasarkan keterangan pengobat tradisional di desa Merabu, kecamatan Kelay, kabupaten Berau, ±5 gram resin kapur ditumbuk hingga halus, diminum dengan air, berfungsi untuk mengobati luka dalam. Resin dari jenis *Dryobalanops keithii* hingga saat ini belum diketahui kandungan kimia dan potensi kegunaannya pada skala yang lebih luas.

Sebagai langkah awal untuk mengetahui kegunaan resin *D. keithii* perlu dilakukan uji fitokimia dan uji GC-MS. Uji fitokimia berfungsi untuk mengetahui jenis metabolit sekunder, yaitu alkaloid, flavonoid, saponin, tannin, terpenoid dan steroid (Rachmawan & Dalimunthe, 2017). Sedangkan untuk menentukan senyawa kimia dalam suatu bahan dapat digunakan uji GC-MS (Darmapatni et al., 2016). Penelitian bertujuan untuk mengetahui kandungan resin dari jenis *Dryobalanops keithii* menggunakan analisa fitokimia dan GC-MS sehingga dapat diketahui potensi kegunaannya pada skala yang lebih luas.

II. METODOLOGI PENELITIAN

1.1. Preparasi contoh uji

Resin *D. keithii* diperoleh dari pohon yang terluka di hutan sekitar desa Merabu, Kecamatan Kelay, Kabupaten Berau. Berdasarkan identifikasi botani *Dryobalanops keithii* sesuai dengan specimen pada <http://specimens.kew.org/herbarium/K000671041>. Resin diperoleh dari 2 pohon, dengan berat 4,05 gram dan 6,21 gram.

10 gram resin yang telah dihaluskan selanjutnya dicampur dengan 20 ml etanol 96% dalam erlenmeyer dan ditutup. Setelah didiamkan selama 48 jam, filtrate disaring dan dipisahkan hingga 10 ml. Setelah

1.2. Pengujian Fitokimia

Pipet 1 ml filtrat pekat ke dalam tabung reaksi, lalu larutkan dalam 14 ml etanol untuk digunakan sebagai bahan uji fitokimia. Uji fitokimia yang dilakukan antara lain uji

alkaloid, flavonoid, saponin, tannin, terpenoid dan steroid.

a. Pengujian Alkaloid (Amta & Jyoti, 2012)

Tambahkan sebanyak 5 ml ekstrak ke dalam 2 ml HCl, lalu tambahkan 1 ml larutan Dragendorff. Warna larutan akan berubah menjadi jingga atau merah yang menandakan ekstrak tersebut mengandung alkaloid.

b. Pengujian Flavonoid (Oeung et al., 2017)

Tambahkan beberapa tetes natrium hidroksida encer (NaOH 1%) ke dalam total 1 ml ekstrak. Ekstrak tampak berwarna kuning bening dan tidak berwarna setelah ditambahkan asam encer (1% HCl), menunjukkan adanya flavonoid.

c. Pengujian Saponin (Surya and Hari, 2017)

Kocok ekstrak ± 10 menit saponin akan terdeteksi bila kocokan ekstrak menghasilkan buih yang stabil, dan saponin tidak akan hilang ketika ditambahkan 1 tetes HCl 2N.

d. Pengujian Tanin (Auwal et al., 2014)

Ketika larutan timbal asetat 1% ditambahkan ke ekstrak dan endapan kuning terbentuk, tanin terdeteksi.

e. Pengujian Terpenoid dan Steroid (Keo et al., 2017)

Tambahkan 10 tetes anhidrida asetat dan 2 tetes asam sulfat pekat ke dalam 1 ml sampel yang dilarutkan dalam aseton. Selain itu, reagen Liebermann-Burchard ditambahkan ke sampel uji, dikocok dan didiamkan selama beberapa menit. Jika terjadi perubahan warna menjadi merah dan ungu, uji terpenoid positif. Jika warna berubah menjadi hijau dan biru, hasil tes menunjukkan adanya steroid.

1.3. Pengujian GC-MS

Untuk memperkirakan kandungan senyawa dalam ekstrak, dilakukan uji GC-MS. GC-MS (Shimadzu QP 2010: Tipe kolom RTx-5MS (Restek Corp.), panjang 30 m, temperatur masuk dan detektor) 250°C, temperatur kerja 50-300°C. Atur kenaikan suhu 50-120°C, pertahankan pada laju

pemanasan 4°C/menit selama 1 menit, lalu suhu 120-300°C, laju pemanasan 6°C/menit, lalu pertahankan itu pada suhu ini selama 5 menit, waktu retensi total (Rt) bisa sampai 60 menit. Gas pembawa: helium, kisaran berat molekul 50-500. Kuantifikasi senyawa yang diprediksi diperoleh dengan membaca area pada grafik GC-MS. Gunakan perpustakaan perangkat lunak Wiley/NIST untuk memperkirakan senyawa yang dihasilkan oleh uji GC-MS.

III. HASIL DAN PEMBAHASAN

Untuk mengetahui kelompok metabolit sekunder dalam resin *D. keithii* larut etanol 96% dilakukan uji fitokimia.

Tabel 1. Hasil uji fitokimia

Table 1. Phytochemical test results

Komponen fitokimia	Hasil uji	Keterangan
alkaloid	+	sedikit
flavonoid	-	Tidak terdeteksi
saponin	-	Tidak terdeteksi
tanin	-	Tidak terdeteksi
terpenoid	+++	banyak
steroid	-	Tidak terdeteksi

Sumber: data primer

Berdasarkan hasil uji fitokimia, resin *D. keithii* larut etanol 96% mengandung sedikit alkaloid dan banyak terpenoid. Sebagai perbandingan, jenis *Dryobalanops oblongifolia* mengandung komponen steroid (Indriani et al., 2020) dan *Dryobalanops aromatica* mengandung terpenoid (Aswandi & Kholibrina, 2020; Pasaribu et al., 2014).

Alkaloid adalah sekelompok bahan kimia alami yang mengandung sebagian besar atom nitrogen basa. Alkaloid diproduksi oleh

berbagai macam organisme, termasuk bakteri, jamur, hewan dan tumbuhan. Senyawa ini memiliki berbagai manfaat seperti anti kanker, anti depresan, anti nosiseptif, anti-inflamasi, anti piretik, anti-platelet, danti-oksidan dan anti-bakteri (Ain et al., 2016; Hartrampf et al., 2018; Khan et al., 2018). Ciri khas alkaloid yaitu memiliki atom N yang berasal dari asam amino dan merupakan bagian dari heterosiklik (Lichman, 2021).

Terpenoid merupakan kelompok metabolit sekunder terbesar, dilihat dari jumlah senyawa dan variasi struktur dasarnya. Terpenoid banyak ditemukan pada tumbuhan tingkat tinggi, namun dari penelitian diketahui bahwa jamur, organisme laut dan serangga juga menghasilkan terpenoid.

Senyawa terpenoid memiliki sifat yang tersusun atas karbon-karbon dalam kelipatan lima. Kebanyakan terpenoid juga diketahui memiliki kerangka karbon yang terdiri dari dua atau lebih unit C₅ yang disebut unit isopren. Disebut satuan isopren karena kerangka karbon C₅ sama dengan senyawa isoprena. Dari beberapa struktur senyawa terpenoid yang telah diidentifikasi, terlihat bahwa unit-unit isoprena ini saling berhubungan secara teratur dimana “kepala” suatu unit bergabung dengan “ekor” unit lainnya dan pengaturan ini dikenal sebagai aturan isopren (Kristanti, Alfinda Novi, et al. 2019).

Untuk mengetahui kandungan senyawa kimia dalam resin *D. keithii* larut etanol 96% dilakukan uji GC-MS. Dugaan senyawa kimia hasil uji GC-MS dapat dilihat pada tabel 1 berikut ini :

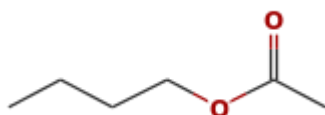
Tabel 2. Hasil uji GC-MS

Table 2. GC-MS test results

No.	Waktu retensi (menit) <i>Retention time (minutes)</i>	Konsentrasi (%) <i>Concentration (%)</i>	Senyawa kimia <i>Chemical compound</i>
1	4.543	1.38	1,6-Heptadien-3-yne

2	5.263	0.06	Acetic acid, butyl ester
3	6.004	0.36	Tyranton -
4	8.253	0.17	α -Pinene
5	8.355	0.01	Rhodium, [1,2-bis(.eta.2-ethenyl)-4-ethenylcyclohexane]di-.mu.-chlorodi-
6	8.509	0.44	α -Pinene
7	9.860	0.04	β -Pinene
8	16.601	2.85	Patchoulane
9	18.306	0.35	1b,5,5,6a-Tetramethyl-octahydro-1-oxa-cyclopropa[a]inden-6-one
10	19.044	0.30	12-Oxabicyclo[9.1.0]dodeca-3,7-diene, 1,5,5,8-tetramethyl-, [1R-(1R*,3E,7E,11R*)]-
11	21.115	0.15	Tetracyclo[6.3.2.0(2,5).0(1,8)]tridecan-9-ol, 4,4-dimethyl-
12	21.526	0.10	Patchoulane
13	22.434	0.27	Patchoulane
14	22.710	0.85	Isocaryophyllene
15	23.231	0.73	Patchoulane
16	23.549	0.10	Humulene
17	24.699	0.04	α -Bulnesene
18	26.718	0.04	Spiro[2.2]pentane-1-carboxylic acid, 2-cyclopropyl-2-methyl-
19	28.612	0.11	Bicyclo[3.3.0]octan-2-one, 7-isopropylidene-
20	32.975	0.07	Palmitic acid
21	44.865	1.04	Phthalic acid
22	45.754	0.08	Bicyclo[2.2.1]heptane-2,5-diol, 1,7,7-trimethyl-, (2-endo,5-exo)-
23	53.000	3.22	Lupeol
24	53.059	14.62	γ -Gurjunene
25	53.385	0.28	Bicyclo[3.3.1]nonan-2-one, 1-methyl-9-(1-methylethylidene)-
26	53.405	0.99	2-[4-Cyclohexylbutanoylamino]-3-chloro-1,4-naphthoquinone
27	54.209	68.90	9,19-Cycloergost-24(28)-en-3-ol, 4,14-dimethyl-, acetate, (3 β .,4. α .,5 α)-
28	54.545	0.67	Phenylacetic acid, 2-(1-adamantyl)ethyl ester
29	54.573	1.77	Betulin

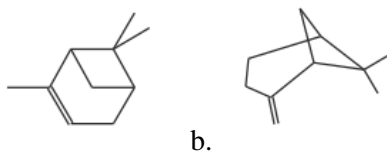
Sumber : Data primer



Gambar (Figure) 1. Acetic acid, butyl ester

Dari hasil uji GC-MS diduga senyawa acetic acid, butyl ester ($C_6H_{12}O_2$) terkandung dalam resin kapur *D.keithii* yang larut dalam

etanol 96% sebanyak 0,06% dengan waktu retensi 5,263 menit. Acetic acid, butyl ester merupakan kelompok ester dari tumbuhan yang berfungsi sebagai antibiotik (Mukhtarovna, 2021), antioksidan (K. Li et al., 2007) dan penyedap rasa (FAO, 2015; Pubchem, 2020). Ester dari tumbuhan aman untuk digunakan sebagai bahan makanan pada manusia.



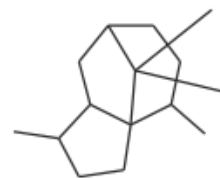
Gambar 2 a. Alpha-pinene, b. Betha-pinene

Berdasarkan tabel hasil uji GC-MS resin kapur *D.keithii* yang larut dalam etanol 96%, diduga mengandung senyawa Alpha-Pinene dan Betha-pinene merupakan senyawa organik dari kelas terpene (Jaoui & Kamens, 2003) dengan rumus molekul $C_{10}H_{16}$ (Gambar 2). Kedua senyawa ini terdapat sebanyak 0,17% dan 0,04% dengan waktu retensi masing-masing senyawa 8,253 menit dan 9,860 menit. Senyawa ini berfungsi sebagai anti-inflamasi dan *inhibitor acetylcholinesterase*, (membantu daya ingat) (Nissen et al., 2010), antimikroba (Silva, Ana Cristina Rivas da et al., 2012; Van Zyl et al., 2006; Wibowo & Komarayati, 2015), anti jamur, pewangi (Salehi et al., 2019).

Senyawa α -pinene dan β -pinene juga memiliki sifat antiseptik yang kuat serta berfungsi sebagai agen antibiotik (Kovač et al., 2015), anti-koagulan (N. Y. Yang et al., 2011), anti-inflammatory (Cho et al., 2017; Kim et al., 2015; X. J. Li et al., 2016; Rufino et al., 2014) serta antitumor (Catanzaro et al., 2012; W. Chen et al., 2015; W. Q. Chen et al., 2014; Kusuhara et al., 2012; Matsuo et al., 2011; Susilo et al., 2020; Y. Wang et al., 2019; Q. Xu et al., 2018; Zhang et al., 2015; Zhao et al., 2018).

Alpha-pinene dan Betha-pinene juga merupakan salah satu dari komponen obat hati dan ginjal (Sybilska et al., 1994). Serta digunakan sebagai anti-bakteri karena efek toksiknya pada membran (Alma et al., 2004). Pada penelitian lain ditemukan memiliki efek penghambat untuk kanker payudara dan leukimia (Zhou et al., 2004) serta alpha-pinene digunakan sebagai antioksidan (Austin et al., 1988; Bouzenna et al., 2017; Karthikeyan et al., 2018, 2019; Türkez & Aydın, 2016) begitu pula dengan betha-pinene (Sharopov et al.,

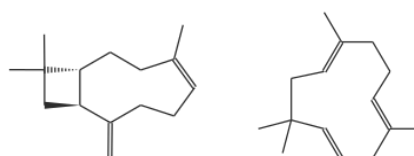
2015). Pemberian senyawa pinenes juga terbukti sangat fleksibel dalam sintesis polimer (Kamigaito & Satoh, 2017; Winnacker & Rieger, 2015). Senyawa ini juga ditemukan dalam minyak esensial rosemary (*Rosmarinus officinalis*) dan pinus (Chahboun et al., 2014; Derwich et al., 2011; Salehi et al., 2019; Wibowo & Komarayati, 2015)



Gambar (Figure) 3. Patchoulane

Patchoulane ($C_{15}H_{26}$) dengan struktur kimia seperti pada gambar 3 juga ditemukan dalam resin kapur *D.keithii* yang larut dalam etanol 96% berdasarkan dari hasil uji GC-MS pada tabel 1. sebanyak 0,10% dengan waktu retensi sebesar 21,526 menit yang di dapat dari hasil GC-MS. Senyawa ini banyak terdapat pada bunga yang mengandung minyak esensial seperti *Cyperus rotundus* dan *Pulicaria vulgaris* (Ameen et al., 2011; Ololade et al., 2014; Zhanzhaxina et al., 2020).

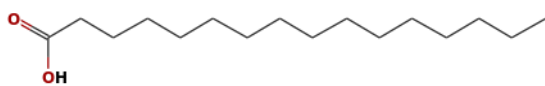
Senyawa Patchoulane yang masih tergolong ke dalam senyawa sesquiterpene ini memiliki sifat sitotoksik dikarenakan kemampuannya dalam menghambat sel kanker ovarium dan sel kanker endometrium (Ahn et al., 2015). Senyawa ini juga dilaporkan sebagai senyawa analgesik dan anti-inflamasi (Chavan et al., 2012). Beberapa penelitian menyebutkan bahwasanya senyawa ini dapat berpotensi menjadi anti-HBV dengan gugus karboksil pada posisi C-4 yang menunjukkan aktivitas melawan sekresi HbsAg dengan cara yang bergantung pada pemberian dosis (H. B. Xu et al., 2015).



Gambar (Figure) 4. A. Isocaryophyllene, b. Humulene

Dalam studi ini, diduga resin kapur *D. keithii* yang larut dalam etanol 96% mengandung senyawa Isocaryophyllene dan Humulene (C₁₅H₂₄) sebanyak 22,710% dan 23,549% dengan waktu retensi masing-masing 0,85 menit dan 0,10 menit. Senyawa ini memiliki struktur kimia seperti gambar 4 ini masih termasuk ke dalam golongan terpena. Kedua senyawa ini biasanya terdapat dalam minyak esensial dari tanaman rempah-rempah seperti oregano (*Origanum vulgare* L.), cinnamon (*Cinnamomum* sp.) dan black pepper (*Piper nigrum* L.) dan minyak atsiri seperti pada *Salvia officinalis*. Senyawa ini dikenal karena penggunaannya penyedap rasa dan kosmetik karena memiliki rasa aromatik yang lemah (Jayaprakasha et al., 2003; Mockute et al., 2001; Orav et al., 2004; Rzepa et al., 2009; Sköld et al., 2006).

Senyawa Isocaryophyllene dan Humulene dapat digunakan sebagai anti-kanker, antibiotik, anti-inflamasi, antioksidan dan digunakan untuk anestesi lokal (Legault & Pichette, 2010). Senyawa humulene dan isocaryophyllene ini juga dapat menghambat pertumbuhan sel MCF-7 masing-masing sebesar 50% dan 69%, dan apabila kedua senyawa ini dikombinasikan maka dapat meningkatkan sitotoksisitas terhadap sel tumor secara in vitro sebesar 75% dan 90% (Legault et al., 2003; Legault & Pichette, 2010; Passos et al., 2007).

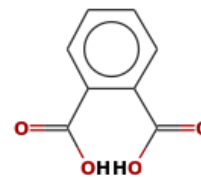


Gambar (Figure) 5. Palmitic-acid

Pada hasil uji GC-MS resin kapur *D. keithii* yang larut dalam etanol 96% terdapat beberapa senyawa dari golongan ester, yaitu Palmitic-acid (PA) (C₁₆H₃₂O₂) yang merupakan asam lemak rantai panjang dengan tulang punggung 16-karbon (gambar 5) sebanyak 32,975% dengan waktu retensi 0,07 menit, Phthalic acid (gambar 6) sebanyak 44,865% dengan waktu retensi 1,04 menit, dan

Phenylacetic acid, 2-(1-adamantyl) ethyl ester 54,545% dengan waktu retensi 0,67 menit.

PA termasuk dalam golongan senyawa ester. Senyawa ini biasanya terdapat pada biji kelapa sawit (Denke & Grundy, 1992) dan memiliki fungsi sebagai antiproliferasi pada sel kanker kolon. PA menunjukkan aktivitas antitumor pada beberapa ekstrak tanaman serta antioksidan (Valente et al., 2012). Palmitic-acid dapat menjadi kandidat obat untuk dematitis atopik dikarenakan senyawa ini mengikat filaggrin pada asam amino LEU D75 (Earlia et al., 2019). PA juga memainkan peran dalam pelemahan ROS (Reactive Oxygen Species), ER stress, apoptosis dan inflamasi yang menyebabkan terjadinya berbagai penyakit, khususnya penyakit degeneratif seperti kanker, diabetes, dll (Nemecz et al., 2019).

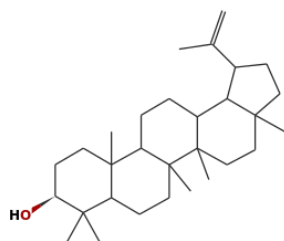


Gambar (Figure) 6. Phthalic acid

Phthalic acid (PTA) dengan gugus kimia seperti gambar 6 (C₈H₆O₄) merupakan oroisomer kristal yang tidak berwarna, dan diduga senyawa ini terdapat dalam resin kapur *D. keithii* yang larut dalam etanol 96%. Senyawa ini digunakan dalam produksi parfum, pewarna dan sintesis beberapa senyawa organik. Senyawa ini terdapat di beberapa tanaman, seperti kemangi (*Ocimum basilicum* L.) (Fattahi et al., 2019), *Capparis decidua* (Pradeep Singh et al., 2011), *Nigella sativa* L. (Aftab et al., 2020) dan *Melastomastrum capitatum* (Ukwubile et al., 2020).

Senyawa PTA ini dapat dijadikan sebagai pendeteksi salah satu antioksidan seperti Glutathione (GSH) dalam suplemen makanan (Detsri & Seeharaj, 2017). Diketahui pula bahwa senyawa ini dapat berfungsi sebagai obat anti kanker (Ukwubile et al.,

2020), antibiotik (Felton, 2019) dan digunakan pula untuk memproduksi produk plastik seperti mainan, botol dan lain-lain (Tan et al., 2017; Yilmaz, 2019).



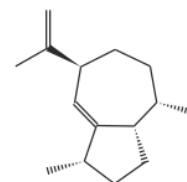
Gambar (Figure) 7. Lupeol

Berdasarkan hasil uji GC-MS resin kapur *D.keithii* yang larut dalam etanol 96% diduga terdapat senyawa Lupeol ($C_{30}H_{50}O$). Senyawa ini merupakan salah satu senyawa dari golongan terpenoid yang memiliki aktivitas biologis seperti anti-virus (Silva et al., 2017), obat ansiolitik (Wal et al., 2019), anti-inflamasi (Fernández et al., 2010; Geetha & Varalakshmi, 2001; Lee et al., 2016; Oliveira-Junior et al., 2019), anti-kanker (Bhattacharyya et al., 2019; He et al., 2018; Jiang et al., 2020; Min et al., 2019; Pitchai et al., 2014; Prasad et al., 2018; Rauth et al., 2016), anti-diabetes (Malik et al., 2019; Pushpanjali et al., 2019; Shreenithi et al., 2019; Soni et al., 2018; Taylor et al., 2012).

Penelitian lain juga menyebutkan beberap fungsi dari lupoeol ini, diantaranya sebagai kardioprotektif (Thilakarathna & Vasantha Rupasinghe, 2012; M. Xu et al., 2020), pelindung kulit (Ciurlea et al., 2010; Payal Singh et al., 2017; Srivastava et al., 2016), anti-protozoa (Das et al., 2017; Goyal et al., 2011; Isah et al., 2018; Shai et al., 2009; Siddique & Saleem, 2011; A. Singh et al., 2020), anti-mikroba (Amoussa et al., 2016; Mutai, 2012; Okusa et al., 2014; Silva et al., 2017; Tolo, F.M., et al., 2010), anti-proliferasi (Siddique & Saleem, 2011), dan sebagai agen nefroprotektif (Devkar et al., 2016; Sinha et al., 2019).

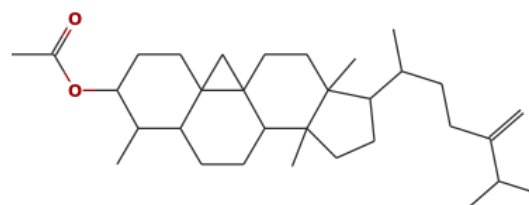
Lupeol ditemukan pada beberapa tanaman buah seperti mangga (Soujanya, 2017), strawberry, anggur, *Hieracium pilosella*

(Gawrońska-Grzywacz & Krzaczek, 2007), *Tamarindus indica* (Imam et al., 2007), *Calendula officinalis* (Kadhim et al., 2019), *Crataeva nurvala* (Sudhahar et al., 2006) dan lain-lain.



Gambar (Figure) 8. gamma-Gurjunene

Hasil uji GC-MS diketahui bahwa dari resin kapur *D.keithii* yang larut dalam etanol 96% diduga terdapat senyawa Gamma-Gurjunene ($C_{15}H_{24}$) sebanyak 53,059% dengan waktu retensi 14,62 menit. Senyawa ini termasuk dalam kelas senyawa organik yang dikenal sebagai seskuiterpenoid. Terdapat dalam *Thymus Kotschyanus* (Karami et al., 2013), rumput teki (*Cyperus rotundus* Linn) (Busman & Kanedi, 2018) serta tanaman lainnya. Seskuiterpenoid digunakan sebagai anti-proliferasi (Ambrož et al., 2017) dan anti-kanker (H. Li et al., 2016; J. Wang et al., 2016; B. Yang et al., 2016).

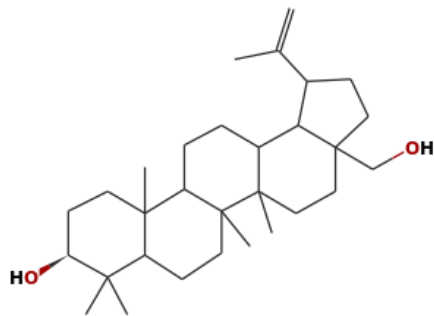


Gambar (Figure) 9. 19-Cycloergost-24(28)-en-3-ol, 4,14-dimethyl-, acetate, (3β,4α,5α)

Hasil uji GC-MS dari resin kapur *D.keithii* yang larut dalam etanol 96% diduga pada waktu rentensi 54,209 menit terdapat senyawa 19-Cycloergost-24(28)-en-3-ol, 4,14-dimethyl-, acetate, (3β,4α,5α) ($C_{32}H_{52}O_2$) dengan konsentrasi 68,90%. 19-Cycloergost-24(28)-en-3-ol, 4,14-dimethyl-, acetate, (3β,4α,5α) termasuk dalam golongan steroid. Senyawa ini memiliki fungsi sebagai anti-mikroba, anti-inflamasi, anti kanker, anti asma, *Hepatoprotective* dan aktivitas diuretik (Chouni et al., 2021; Rajalakshmi & Mohan,

2016). 19-Cycloergost-24(28)-en-3-ol, 4,14-dimethyl-, acetate, (3 β ,4 α ,5 α) dikenal juga dengan nama Cycloeucalenyl acetate atau Cycloeucalenol acetate. Selain tergolong steroid, Cycloeucalenol acetate juga memiliki sifat triterpenoid (Zubair et al., 2017). Cycloeucalenol acetate berfungsi sebagai cardiotonic dengan efek toksisitas yang rendah sehingga dapat digunakan sebagai bahan baku obat di masa datang (Adewusi et al., 2013).

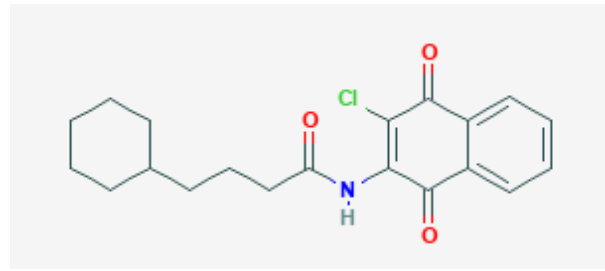
Berdasarkan tabel hasil uji GC-MS diatas diduga dalam resin kapur *D.keithii* yang larut dalam etanol 96% terdapat Phenylacetic acid, 2-(1-adamantyl) ethyl ester yang memiliki gugus ester. Ethyl ester berfungsi sebagai anti-oksidan, hemolitik, Hipokolesterolemia, perasa, nematisida dan aktivitas anti-androgenik (Tyagi & Agarwal, 2017).



Gambar (Figure) 10. Betulin

Senyawa hasil uji GC-MS resin kapur *D.keithii* yang larut dalam etanol 96%, diduga salah satunya adalah senyawa Betulin (C₃₀H₅₀O₂) tergolong dalam terpenoid (gambar 10) yang terdapat pada banyak tanaman.

Senyawa betulin berpotensi sebagai anti-HIV dan anti-inflamasi (Chaniad et al., 2019). Senyawa ini bersifat sitotoksik (Pfarr et al., 2015), berfungsi sebagai anti kanker (Bebenek et al., 2015; Y. Li et al., 2016; Yim et al., 2015; Zehra et al., 2019), anti-leishmanial (Alakurtti et al., 2010), anti oksidan, anti-inflamasi, antivirus, anti-bakteri, anti-neoplastik (Amiri et al., 2020; Rastogi et al., 2015) dan antineurodegeneratif (Tsai et al., 2017).



Hasil uji GC-MS dari resin kapur *D.keithii* yang larut dalam etanol 96% diduga terdapat senyawa 2-[4-Cyclohexylbutanoylamino]-3-chloro-1,4-naphthoquinone (C₂₀H₂₂ClNO₃) sebanyak 0,99% pada waktu retensi 53,405 menit. Senyawa ini termasuk dalam alkaloid karena memiliki gugus N. Senyawa ini juga memiliki gugus fungsional naphthoquinone yang secara umum berfungsi sebagai anti-inflamasi, obat analgesik, anti-bakteri, anti-parasit, antioksidan, leishmanicidal dan anti-tumor (Han et al., 2019; Jentzsch et al., 2020; Rauf et al., 2020; H. Zhao et al., 2020).

IV. KESIMPULAN

Hasil uji fitokimia dari resin kapur *D.keithii* yang larut dalam etanol 96% mengandung alkaloid dan terpenoid. Hasil uji GC-MS mendeteksi 29 senyawa kimia pada waktu retensi 4,543-54,573 menit. Senyawa 2-[4-Cyclohexylbutanoylamino]-3-chloro-1,4-naphthoquinone (C₂₀H₂₂ClNO₃) sebanyak 0,99% pada waktu retensi 53,405 menit tergolong alkaloid. Sedangkan contoh senyawa terpenoid adalah α -Pinene, β -pinene, humulene dan γ -gurjunene. Berbagai senyawa yang terkandung dalam resin *Dryobalanops keithii* berpotensi sebagai antioksidan, anti-inflamasi, antibiotic dan beberapa fungsi biologis lainnya.

DAFTAR PUSTAKA

- Adewusi, E. A., Steenkamp, P., Fouche, G., & Steenkamp, V. (2013). Isolation of cycloeucalenol from *Boopha disticha* and evaluation of its cytotoxicity. *Natural Product Communications*, 8(9), 1213–1216. <https://doi.org/10.1177/1934578x1300800906>
- Aftab, A., Yousaf, Z., Aftab, Z. E. H., Younas, A., Riaz, N., Rashid, M., Shamsheer, H. B., Razzaq, Z., & Javaid, A. (2020). Pharmacological screening and GC-MS

- analysis of vegetative/reproductive parts of *Nigella sativa* L. *Pakistan Journal of Pharmaceutical Sciences*, 33(5), 2103–2111. <https://doi.org/10.36721/PJPS.2020.33.5.REG.2103-2111.1>
- Ahn, J. H., Lee, T. W., Kim, K. H., Byun, H., Ryu, B., Lee, K. T., Jang, D. S., & Choi, J. H. (2015). 6-Acetoxy Cyperene, a Patchoulane-type Sesquiterpene Isolated from *Cyperus rotundus* Rhizomes Induces Caspase-dependent Apoptosis in Human Ovarian Cancer Cells. *Phytotherapy Research*, 29(9), 1330–1338. <https://doi.org/10.1002/ptr.5385>
- Ain, Q. U., Khan, H., Mubarak, M. S., & Pervaiz, A. (2016). Plant alkaloids as antiplatelet agent: Drugs of the future in the light of recent developments. *Frontiers in Pharmacology*, 7(SEP), 1–9. <https://doi.org/10.3389/fphar.2016.00292>
- Alakurtti, S., Heiska, T., Kiriazis, A., Sacerdoti-Sierra, N., Jaffe, C. L., & Yli-Kauhaluoma, J. (2010). Synthesis and anti-leishmanial activity of heterocyclic betulin derivatives. *Bioorganic and Medicinal Chemistry*, 18(4), 1573–1582. <https://doi.org/10.1016/j.bmc.2010.01.003>
- Alma, M. H., Nitz, S., Kollmannsberger, H., Digrak, M., Efe, F. T., & Yilmaz, N. (2004). Chemical composition and antimicrobial activity of the essential oils from the gum of Turkish Pistachio (*Pistacia vera* L.). *Journal of Agricultural and Food Chemistry*, 52(12), 3911–3914. <https://doi.org/10.1021/jf040014e>
- Ambrož, M., Matoušková, P., Skarka, A., Zajdlová, M., Žáková, K., & Skálová, L. (2017). The effects of selected sesquiterpenes from *myrica rubra* essential oil on the efficacy of doxorubicin in sensitive and resistant cancer cell lines. *Molecules*, 22(6). <https://doi.org/10.3390/molecules22061021>
- Ameen, O. M., Usman, L. A., Oladosu, I. A., Olawore, N. O., & Ogunwande, I. A. (2011). Bioactivity of rhizome essential oils from two varieties of *Cyperus articulatus* (L.) grown in Nigeria, using brine shrimp (*Artemia salina*) lethality tests. *Journal of Medicinal Plants Research*, 5(6), 1031–1033.
- Amiri, S., Dastghaib, S., Ahmadi, M., Mehrbod, P., Khadem, F., Behrouj, H., Aghanoori, M. R., Machaj, F., Ghamsari, M., Rosik, J., Hudecki, A., Afkhami, A., Hashemi, M., Los, M. J., Mokarram, P., Madrakian, T., & Ghavami, S. (2020). Betulin and its derivatives as novel compounds with different pharmacological effects. *Biotechnology Advances*, 38. <https://doi.org/10.1016/j.biotechadv.2019.06.008>
- Amoussa, A. M. O., Lagnika, L., Bourjot, M., Vonthron-Senecheau, C., & Sanni, A. (2016). Triterpenoids from *Acacia ataxacantha* DC: Antimicrobial and antioxidant activities. *BMC Complementary and Alternative Medicine*, 16(1), 1–8. <https://doi.org/10.1186/s12906-016-1266-y>
- Aswandi, A., & Kholibrina, C. R. (2020). Potensi Minyak Atsiri Kamfer Sumatera (*Dryobalanops aromatica* Gaertn .) Untuk Bahan Baku Obat Herbal Indonesia merupakan salah satu pemilik ekosistem hutan tropis terluas di untuk modal pembangunan nasional (Biro Pusat Statistik , 2020). Namun , manfaa. *Jurnal Farmasi Udayana*, 171–179.
- Austin, C. A., Shephard, E. A., Pike, S. F., Rabin, B. R., & Phillips, I. R. (1988). The effect of terpenoid compounds on cytochrome P-450 levels in rat liver. *Biochemical Pharmacology*, 37(11), 2223–2229. [https://doi.org/10.1016/0006-2952\(88\)90585-0](https://doi.org/10.1016/0006-2952(88)90585-0)
- Bębenek, E., Chodurek, E., Orchel, A., Dzierzewicz, Z., & Boryczka, S. (2015). Antiproliferative activity of novel acetylenic derivatives of betulin against G-361 human melanoma cells. *Acta Poloniae Pharmaceutica - Drug Research*, 72(4), 699–703.
- Bhattacharyya, S., Mitra, D., Ray, S., Biswas, N., Banerjee, S., Majumder, B., Mustafi, S. M., & Murmu, N. (2019). Reversing effect of Lupeol on vasculogenic mimicry in murine melanoma progression. *Microvascular Research*, 121(October 2018), 52–62. <https://doi.org/10.1016/j.mvr.2018.10.008>
- Bouzenna, H., Hfaiedh, N., Giroux-Metges, M. A., Elfeki, A., & Talarmin, H. (2017). Potential protective effects of alpha-pinene against cytotoxicity caused by aspirin in the IEC-6 cells. *Biomedicine and Pharmacotherapy*, 93, 961–968. <https://doi.org/10.1016/j.biopha.2017.06.031>
- Busman, H., & Kanedi, M. (2018). *CHEMICAL COMPOSITION OF THE ESSENTIAL OILS DISTILLED FROM TUBER OF RUMPUT TEKI (CYPERUS ROTUNDUS LINN .) GROWING IN TANGGAMUS*, *European of Biomedical AND Pharmaceutical sciences*. April.
- Catanzaro, I., Caradonna, F., Barbata, G., Saverini, M., Mauro, M., & Sciandrello, G. (2012).

- Genomic instability induced by α -pinene in Chinese hamster cell line. *Mutagenesis*, 27(4), 463–469.
<https://doi.org/10.1093/mutage/ges005>
- Chahboun, N., Esmail, A., Rhaïem, N., Abed, H., Amiyare, R., Barrahi, M., Berrabeh, M., Oudda, H., & Ouhssine, M. (2014). Extraction and study of the essential oil *Rosmarinus officinalis* cuellie in the region of Taza, Morocco. *Der Pharma Chemica*, 6(3), 367–372.
- Chaniad, P., Sudsai, T., Septama, A. W., Chukaew, A., & Tewtrakul, S. (2019). Evaluation of Anti-HIV-1 integrase and anti-inflammatory activities of compounds from *betula alnoides buch-ham*. *Advances in Pharmacological Sciences*, 2019.
<https://doi.org/10.1155/2019/2573965>
- Chavan, M. J., Wakte, P. S., & Shinde, D. B. (2012). Analgesic and anti-inflammatory activities of the sesquiterpene fraction from *Annona reticulata* L. bark. *Natural Product Research*, 26(16), 1515–1518.
<https://doi.org/10.1080/14786419.2011.564583>
- Chen, W., Liu, Y., Li, M., Mao, J., Zhang, L., Huang, R., Jin, X., & Ye, L. (2015). Antitumor effect of α -pinene on human hepatoma cell lines through inducing G2/M cell cycle arrest. *Journal of Pharmacological Sciences*, 127(3), 332–338.
<https://doi.org/10.1016/j.jphs.2015.01.008>
- Chen, W. Q., Xu, B., Mao, J. W., Wei, F. X., Li, M., Liu, T., Jin, X. B., & Zhang, L. R. (2014). Inhibitory effects of α -pinene on hepatoma carcinoma cell proliferation. In *Asian Pacific Journal of Cancer Prevention* (Vol. 15, Issue 7, pp. 3293–3297).
<https://doi.org/10.7314/APJCP.2014.15.7.3293>
- Cho, K. S., Lim, Y. R., Lee, K., Lee, J., Lee, J. H., & Lee, I. S. (2017). Terpenes from forests and human health. *Toxicological Research*, 33(2), 97–106.
<https://doi.org/10.5487/TR.2017.33.2.097>
- Chouni, A., Pal, A., Gopal, P. K., & Paul, S. (2021). GC-MS analysis and screening of anti-proliferative potential of methanolic extract of *Garcinia cowa* on different cancer cell lines. *Pharmacognosy Journal*, 13(2), 347–361.
<https://doi.org/10.5530/pj.2021.13.45>
- Ciurlea, S., Ionescu, D., Redes, L. C., & Soica, C. (2010). Lupeol, a pentacyclic triterpene that reduces the lesions and irritability on murine skin and is effective on in vitro tumor models. *Journal of Agroalimentary Processes and Technologies*, 16(4), 427–432.
- Darmapatni, K. A. G., Basori, A., & Suaniti, N. made. (2016). Pengembangan Metode Gc- Ms Untuk. *Jurnal Biosains Pascasarjana*, 18(3), 255–270.
- Das, A., Jawed, J. J., Das, M. C., Sandhu, P., De, U. C., Dinda, B., Akhter, Y., & Bhattacharjee, S. (2017). Antileishmanial and immunomodulatory activities of lupeol, a triterpene compound isolated from *Sterculia villosa*. *International Journal of Antimicrobial Agents*, 50(4), 512–522.
<https://doi.org/10.1016/j.ijantimicag.2017.04.022>
- Denke, M. A., & Grundy, S. M. (1992). Comparison of effects of lauric acid and palmitic acid on plasma lipids and lipoproteins. *American Journal of Clinical Nutrition*, 56(5), 895–898.
<https://doi.org/10.1093/ajcn/56.5.895>
- Derwich, E., Benziane, Z., & Chabir, R. (2011). AROMATIC AND MEDICINAL PLANTS OF MOROCCO: CHEMICAL COMPOSITION OF ESSENTIAL OILS OF *Rosmarinus officinalis* AND *Juniperus Phoenicea*. *International Journal of Applied Biology and P*.
- Detsri, E., & Seeharaj, P. (2017). Colorimetric detection of glutathione based on phthalic acid assisted synthesis of silver nanoparticles. *Colloids and Surfaces A: Physicochemical and Engineering Aspects*, 533(August), 125–132.
<https://doi.org/10.1016/j.colsurfa.2017.08.037>
- Devkar, R. A., Chaudhary, S., Adepu, S., Xavier, S. K., Chandrashekar, K. S., & Setty, M. M. (2016). Evaluation of antiurolithiatic and antioxidant potential of *Lepidagathis prostrata*: A Pashanbhed plant. *Pharmaceutical Biology*, 54(7), 1237–1245.
<https://doi.org/10.3109/13880209.2015.1066397>
- Earlia, N., Muslem, Suhendra, R., Amin, M., Prakoeswa, C. R. S., Khairan, & Idroes, R. (2019). GC/MS Analysis of Fatty Acids on Pliek U Oil and Its Pharmacological Study by Molecular Docking to Filaggrin as a Drug Candidate in Atopic Dermatitis Treatment. *Scientific World Journal*, 2019.
<https://doi.org/10.1155/2019/8605743>
- Evans, T. D., Piseth, H., Phaktra, P., & Mary, H. (2003). *A study of resin-tapping and livelihoods in southern Mondulkiri, Cambodia: with implications for*

- conservation and forest management. *January*, 91.
- FAO. (2015). *Online Edition: "Specifications for Flavourings."* 522, 4820.
- Fattahi, B., Arzani, K., Souri, M. K., & Barzegar, M. (2019). Effects of cadmium and lead on seed germination, morphological traits, and essential oil composition of sweet basil (*Ocimum basilicum* L.). *Industrial Crops and Products*, 138(February), 111584. <https://doi.org/10.1016/j.indcrop.2019.111584>
- Felton, C. (2019). *OpenRiver Comparing Beta-Llctamase Inhibition Activity of Phthalic Acid Derivatives*. 2018.
- Fernández, M. A., de las Heras, B., Garcia, M. D., Sáenz, M. T., & Villar, A. (2010). New insights into the mechanism of action of the anti-inflammatory triterpene lupeol. *Journal of Pharmacy and Pharmacology*, 53(11), 1533–1539. <https://doi.org/10.1211/0022357011777909>
- Gawrońska-Grzywacz, M., & Krzaczek, T. (2007). Identification and determination of triterpenoids in *Hieracium pilosella* L. *Journal of Separation Science*, 30(5), 746–750. <https://doi.org/10.1002/jssc.200600253>
- Geetha, T., & Varalakshmi, P. (2001). Anti-inflammatory activity of lupeol and lupeol linoleate in rats. *Journal of Ethnopharmacology*, 76(1), 77–80. [https://doi.org/10.1016/S0378-8741\(01\)00175-1](https://doi.org/10.1016/S0378-8741(01)00175-1)
- Goyal, M., Pareek, A., Nagori, B. P., & Sasmal, D. (2011). *Aerva lanata*: A review on phytochemistry and pharmacological aspects. *Pharmacognosy Reviews*, 5(10), 195–198. <https://doi.org/10.4103/0973-7847.91120>
- Gusti, R. E. P. (2014). Physico chemical properties of purified mata kucing dammar without solvent. *Jurnal Penelitian Hasil Hutan*, 32(3), 167–174.
- Han, Y., Zhang, J., Hu, C. Q., Zhang, X., Ma, B., & Zhang, P. (2019). In silico ADME and toxicity prediction of ceftazidime and its impurities. *Frontiers in Pharmacology*, 10(APR), 1–12. <https://doi.org/10.3389/fphar.2019.00434>
- Hartrampf, N., Winter, N., Pupo, G., Stoltz, B. M., & Trauner, D. (2018). Total Synthesis of the Norhasubanan Alkaloid Stephdiamine. *Journal of the American Chemical Society*, 140(28), 8675–8680. <https://doi.org/10.1021/jacs.8b01918>
- He, W., Li, X., & Xia, S. (2018). Lupeol triterpene exhibits potent antitumor effects in A427 human lung carcinoma cells via mitochondrial mediated apoptosis, ROS generation, loss of mitochondrial membrane potential and downregulation of m-TOR/PI3K/AKT signalling pathway. *Journal of B.U.ON.*, 23(3), 635–640.
- Imam, S., Azhar, I., Hasan, M. M., Ali, M. S., & Ahmed, S. W. (2007). Two triterpenes lupanone and lupeol isolated and identified from *Tamarindus indica* linn. *Pakistan Journal of Pharmaceutical Sciences*, 20(2), 125–127.
- Indriani, I., Aminah, N. S., & Puspaningsih, N. N. T. (2020). Antiplasmodial Activity of Stigmastane Steroids from *Dryobalanops oblongifolia* Stem Bark. *Open Chemistry*, 18(1), 259–264. <https://doi.org/10.1515/chem-2020-0027>
- Isah, M. B., Tajuddeen, N., Umar, M. I., Alhafiz, Z. A., Mohammed, A., & Ibrahim, M. A. (2018). Terpenoids as Emerging Therapeutic Agents: Cellular Targets and Mechanisms of Action against Protozoan Parasites. In *Studies in Natural Products Chemistry* (1st ed., Vol. 59). Elsevier B.V. <https://doi.org/10.1016/B978-0-444-64179-3.00007-4>
- Jaoui, M., & Kamens, R. M. (2003). Gaseous and particulate oxidation products analysis of a mixture of α -pinene + β -pinene/O₃/air in the absence of light and α -pinene + β -pinene/NO_x/air in the presence of natural sunlight. *Journal of Atmospheric Chemistry*, 44(3), 259–297. <https://doi.org/10.1023/A:1022977427523>
- Jayaprakash, G. K., Mohan Rao, L. J., & Sakariah, K. K. (2003). Volatile constituents from *Cinnamomum zeylanicum* fruit stalks and their antioxidant activities. *Journal of Agricultural and Food Chemistry*, 51(15), 4344–4348. <https://doi.org/10.1021/jf034169i>
- Jentzsch, J., Koko, W. S., Al Nasr, I. S., Khan, T. A., Schobert, R., Ersfeld, K., & Biersack, B. (2020). New Antiparasitic Bis-Naphthoquinone Derivatives. *Chemistry and Biodiversity*, 17(2). <https://doi.org/10.1002/cbdv.201900597>
- Jiang, Y., Hong, D., Lou, Z., Tu, X., & Jin, L. (2020). Lupeol inhibits migration and invasion of colorectal cancer cells by suppressing RhoA-ROCK1 signaling pathway. *Naunyn-Schmiedeberg's Archives of Pharmacology*, 393(11), 2185–2196. <https://doi.org/10.1007/s00210-020-01815-3>
- Kadhim, N. A., Al-ani, W. M. K., & Al-joboury, I. S. (2019). Detection of Lupeol in *Calendula*

- Officinalis Grown in Iraq by GC-MS Analysis. *Al Mustansiriyah Journal of Pharmaceutical Sciences*, 19(4), 69–76. <https://doi.org/10.32947/ajps.19.04.0420>
- Kamigaito, M., & Satoh, K. (2017). Sustainable Vinyl Polymers via Controlled Polymerization of Terpenes. *Sustainable Polymers from Biomass*, 55–90. <https://doi.org/10.1002/9783527340200.ch4>
- Karami, E., Ali, A., & Jamshidi, A. H. (2013). Investigation of Chemical Essential Oil Components of Thymus Lancifoliosus in Zagheh Area (In Lorestan Province). *Iranian Journal of Pharmaceutical Research*, 12(Supplement 2), 61. http://ijpr.sbm.ac.ir/?_action=showPDF&article=1379&_ob=130553e1aeab3a2c5a5532d06b48f712&fileName=full_text.pdf%0Ahttp://ovidsp.ovid.com/ovidweb.cgi?T=JS&PAGE=reference&D=emed14&NEWS=N&AN=71267648%0Ahttp://ovidsp.ovid.com/ovidweb.cgi?T=JS&PAGE=reference&
- Karthikeyan, R., Kanimozhi, G., Madahavan, N. R., Agilan, B., Ganesan, M., Prasad, N. R., & Rathinaraj, P. (2019). Alpha-pinene attenuates UVA-induced photoaging through inhibition of matrix metalloproteinases expression in mouse skin. In *Life Sciences* (Vol. 217). Elsevier Inc. <https://doi.org/10.1016/j.lfs.2018.12.003>
- Karthikeyan, R., Kanimozhi, G., Prasad, N. R., Agilan, B., Ganesan, M., & Srithar, G. (2018). Alpha pinene modulates UVA-induced oxidative stress, DNA damage and apoptosis in human skin epidermal keratinocytes. In *Life Sciences* (Vol. 212). Elsevier Inc. <https://doi.org/10.1016/j.lfs.2018.10.004>
- Khan, H., Nabavi, S. M., Sureda, A., Mehterov, N., Gulei, D., Berindan-Neagoe, I., Taniguchi, H., & Atanasov, A. G. (2018). Therapeutic potential of songorine, a diterpenoid alkaloid of the genus Aconitum. *European Journal of Medicinal Chemistry*, 153, 29–33. <https://doi.org/10.1016/j.ejmech.2017.10.065>
- Kim, D. S., Lee, H. J., Jeon, Y. D., Han, Y. H., Kee, J. Y., Kim, H. J., Shin, H. J., Kang, J., Lee, B. S., Kim, S. H., Kim, S. J., Park, S. H., Choi, B. M., Park, S. J., Um, J. Y., & Hong, S. H. (2015). Alpha-Pinene Exhibits Anti-Inflammatory Activity Through the Suppression of MAPKs and the NF- κ B Pathway in Mouse Peritoneal Macrophages. *American Journal of Chinese Medicine*, 43(4), 731–742. <https://doi.org/10.1142/S0192415X15500457>
- Kristanti, Alfinda Novi, Aminah, Nanik Siti, Tanjung, Mulyadi dan Kurniadi Bambang. (2019). Buku Ajar Fitokimia. Universitas Airlangga.
- Kovač, J., Šimunović, K., Wu, Z., Klančnik, A., Bucar, F., Zhang, Q., & Možina, S. S. (2015). Antibiotic resistance modulation and modes of action of (-)- α -Pinene in *Campylobacter jejuni*. *PLoS ONE*, 10(4), 1–14. <https://doi.org/10.1371/journal.pone.0122871>
- Kusuhara, M., Urakami, K., Masuda, Y., Zangiacomì, V., Ishii, H., Tai, S., Maruyama, K., & Yamaguchi, K. (2012). Fragrant environment with α -pinene decreases tumor growth in mice. *Biomedical Research*, 33(1), 57–61. <https://doi.org/10.2220/biomedres.33.57>
- Lee, C., Lee, J. W., Seo, J. Y., Hwang, S. W., Im, J. P., & Kim, J. S. (2016). Lupeol inhibits LPS-induced NF-kappa B signaling in intestinal epithelial cells and macrophages, and attenuates acute and chronic murine colitis. *Life Sciences*, 146, 100–108. <https://doi.org/10.1016/j.lfs.2016.01.001>
- Legault, J., Dahl, W., Debiton, E., Pichette, A., & Madelmont, J. C. (2003). Antitumor activity of balsam fir oil: Production of reactive oxygen species induced by α -humulene as possible mechanism of action. *Planta Medica*, 69(5), 402–407. <https://doi.org/10.1055/s-2003-39695>
- Legault, J., & Pichette, A. (2010). Potentiating effect of β -caryophyllene on anticancer activity of α -humulene, isocaryophyllene and paclitaxel. *Journal of Pharmacy and Pharmacology*, 59(12), 1643–1647. <https://doi.org/10.1211/jpp.59.12.0005>
- Li, H., Li, M., Wang, G., Shao, F., Chen, W., Xia, C., Wang, S., Li, Y., Zhou, G., & Liu, Z. (2016). EM23, a natural sesquiterpene lactone from *Elephantopus mollis*, induces apoptosis in human myeloid leukemia cells through thioredoxin- and reactive oxygen species-mediated signaling pathways. *Frontiers in Pharmacology*, 7(MAR), 1–15. <https://doi.org/10.3389/fphar.2016.00077>
- Li, K., Li, X. M., Ji, N. Y., & Wang, B. G. (2007). Natural bromophenols from the marine red alga *Polysiphonia urceolata* (Rhodomelaceae): Structural elucidation and DPPH radical-scavenging activity. *Bioorganic and Medicinal Chemistry*, 15(21), 6627–6631. <https://doi.org/10.1016/j.bmc.2007.08.023>

- Li, X. J., Yang, Y. J., Li, Y. S., Zhang, W. K., & Tang, H. Bin. (2016). α -Pinene, linalool, and 1-octanol contribute to the topical anti-inflammatory and analgesic activities of frankincense by inhibiting COX-2. *Journal of Ethnopharmacology*, 179, 22–26. <https://doi.org/10.1016/j.jep.2015.12.039>
- Li, Y., Liu, X., Jiang, D., Lin, Y., Wang, Y., Li, Q., Liu, L., & Jin, Y. H. (2016). Betulin induces reactive oxygen species-dependent apoptosis in human gastric cancer SGC7901 cells. *Archives of Pharmacal Research*, 39(9), 1257–1265. <https://doi.org/10.1007/s12272-016-0761-5>
- Lichman, B. R. (2021). The scaffold-forming steps of plant alkaloid biosynthesis. *Natural Product Reports*, 38(1), 103–129. <https://doi.org/10.1039/d0np00031k>
- Malik, A., Jamil, U., Butt, T. T., Waquar, S., Gan, S. H., Shafique, H., & Jafar, T. H. (2019). In silico and in vitro studies of lupeol and iso-orientin as potential antidiabetic agents in a rat model. *Drug Design, Development and Therapy*, 13, 1501–1513. <https://doi.org/10.2147/DDDT.S176698>
- Matsuo, A. L., Figueiredo, C. R., Arruda, D. C., Pereira, F. V., Borin Scutti, J. A., Massaoka, M. H., Travassos, L. R., Sartorelli, P., & Lago, J. H. G. (2011). α -Pinene isolated from *Schinus terebinthifolius* Raddi (Anacardiaceae) induces apoptosis and confers antimetastatic protection in a melanoma model. *Biochemical and Biophysical Research Communications*, 411(2), 449–454. <https://doi.org/10.1016/j.bbrc.2011.06.176>
- Min, T. R., Park, H. J., Ha, K. T., Chi, G. Y., Choi, Y. H., & Park, S. H. (2019). Suppression of EGFR/STAT3 activity by lupeol contributes to the induction of the apoptosis of human non-small cell lung cancer cells. *International Journal of Oncology*, 55(1), 320–330. <https://doi.org/10.3892/ijo.2019.4799>
- Mockute, D., Bernotiene, G., & Judzentiene, A. (2001). The essential oil of *Origanum vulgare* L. ssp. *vulgare* growing wild in Vilnius district (Lithuania). *Phytochemistry*, 57(1), 65–69. [https://doi.org/10.1016/S0031-9422\(00\)00474-X](https://doi.org/10.1016/S0031-9422(00)00474-X)
- Mukhtarovna, N. R. (2021). *Obtaining Butyl Acetate by the Method of Hydrolysis*. April, 4–6.
- Mutai, C. (2012). Effects of Triterpenoids on Herpes Simplex Virus Type1 (Hsv-1) In Vitro. *Medicinal & Aromatic Plants*, 01(S1), 1–4. <https://doi.org/10.4172/scientificreports.140>
- Nemecz, M., Constantin, A., Dumitrescu, M., Alexandru, N., Filippi, A., Tanko, G., & Georgescu, A. (2019). The distinct effects of palmitic and oleic acid on pancreatic beta cell function: The elucidation of associated mechanisms and effector molecules. *Frontiers in Pharmacology*, 9(JAN), 1–16. <https://doi.org/10.3389/fphar.2018.01554>
- Nissen, L., Zatta, A., Stefanini, I., Grandi, S., Sgorbati, B., Biavati, B., & Monti, A. (2010). Characterization and antimicrobial activity of essential oils of industrial hemp varieties (*Cannabis sativa* L.). *Fitoterapia*, 81(5), 413–419. <https://doi.org/10.1016/j.fitote.2009.11.010>
- Okusa, P. N., Stévigny, C., Névrumont, M., Gelbecke, M., Van Antwerpen, P., Braekman, J. C., & Duez, P. (2014). Ferulaldehyde and lupeol as direct and indirect antimicrobial compounds from *Cordia gillettii* (Boraginaceae) root barks. *Natural Product Communications*, 9(5), 619–622. <https://doi.org/10.1177/1934578x1400900506>
- Oliveira-Junior, M. S., Pereira, E. P., de Amorim, V. C. M., Reis, L. T. C., do Nascimento, R. P., da Silva, V. D. A., & Costa, S. L. (2019). Lupeol inhibits LPS-induced neuroinflammation in cerebellar cultures and induces neuroprotection associated to the modulation of astrocyte response and expression of neurotrophic and inflammatory factors. *International Immunopharmacology*, 70(November 2018), 302–312. <https://doi.org/10.1016/j.intimp.2019.02.055>
- Ololade, Z. S., Fakankun, O. A., & Udi, O. U. (2014). *Ocimum basilicum* var. *purpureum* Floral Essential Oil: Phytochemicals, Phenolic Content, Antioxidant, Free Radical Scavenging and Antimicrobial Potentials. *Global Journal of Science Frontier Research: B Chemistry*, 14(7), 31–37.
- Orav, A., Stulova, I., Kailas, T., & Müürisepp, M. (2004). Effect of Storage on the Essential Oil Composition of *Piper nigrum* L. Fruits of Different Ripening States. *Journal of Agricultural and Food Chemistry*, 52(9), 2582–2586. <https://doi.org/10.1021/jf030635s>
- Pasaribu, G., Gusmailina, G., & Komarayati, S. (2014). PEMANFAATAN MINYAK Dryobalanops aromatica SEBAGAI BAHAN PEWANGI ALAMI. *Jurnal Penelitian Hasil Hutan*, 32(3), 235–242.

- <https://doi.org/10.20886/jphh.2014.32.3.235-242>
- Passos, G. F., Fernandes, E. S., da Cunha, F. M., Ferreira, J., Pianowski, L. F., Campos, M. M., & Calixto, J. B. (2007). Anti-inflammatory and anti-allergic properties of the essential oil and active compounds from *Cordia verbenacea*. *Journal of Ethnopharmacology*, *110*(2), 323–333. <https://doi.org/10.1016/j.jep.2006.09.032>
- Pfarr, K., Danciu, C., Arlt, O., Neske, C., Dehelean, C., Pfeilschifter, J. M., & Radeke, H. H. (2015). Simultaneous and dose dependent melanoma cytotoxic and immune stimulatory activity of betulin. *PLoS ONE*, *10*(3), 1–21. <https://doi.org/10.1371/journal.pone.0118802>
- Pitchai, D., Roy, A., & Ignatius, C. (2014). In vitro evaluation of anticancer potentials of lupeol isolated from *Elephantopus scaber* L. on MCF-7 cell line. *Journal of Advanced Pharmaceutical Technology and Research*, *5*(4), 179–184. <https://doi.org/10.4103/2231-4040.143037>
- Prasad, N., Sabarwal, A., Yadav, U. C. S., & Singh, R. P. (2018). Lupeol induces S-phase arrest and mitochondria-mediated apoptosis in cervical cancer cells. *Journal of Biosciences*, *43*(2), 249–261. <https://doi.org/10.1007/s12038-018-9743-8>
- Pubchem. (2020). n-Butyl Acetate. *National Center for Biotechnology Information, August*.
- Purwaningsih. (2004). Ecological distribution of Dipterocarpaceae species in Indonesia. *Biodiversitas, Journal of Biological Diversity*, *5*(2), 89–95. <https://doi.org/10.13057/biodiv/d050210>
- Pushpanjali, G., Vishnupriya, V., Ponnulakshmi, R., Gayathri, R., Madhan, K., Shyamaladevi, B., & Selvaraj, J. (2019). Effect of lupeol on enzymatic and non-enzymatic antioxidants in type-2 diabetic adult male Wistar rats. *Drug Invention Today*, *12*(5), 873–876.
- Rajalakshmi, K., & Mohan, V. R. (2016). Determination of Bioactive Components of *Myxopyrum Serratulum* a.W. Hill (Oleaceae) Stem By Gc-Ms Analysis. *International Research Journal of Pharmacy*, *7*(7), 36–42. <https://doi.org/10.7897/2230-8407.07782>
- Rachmawan, A., & Dalimunthe, C. I. (2017). Prospek Pemanfaatan Metabolit Sekunder Tumbuhan Sebagai Pestisida Nabati Untuk Pengendalian Patogen Pada Tanaman Karet. *Warta Perkaretan*, *36*(1), 15–28. <https://doi.org/10.22302/ppk.wp.v36i1.324>
- Rastogi, S., Pandey, M. M., Kumar, A., & Rawat, S. (2015). Medicinal plants of the genus *Betula*—Traditional uses and a phytochemical–pharmacological review. *Journal of Ethnopharmacology*, *159*(January), 62–83.
- Rauf, A., Abu-Izneid, T., Rashid, U., Alhumaydhi, F. A., Bawazeer, S., Khalil, A. A., Aljohani, A. S. M., Abdallah, E. M., Al-Tawaha, A. R., Mabkhot, Y. N., Shariati, M. A., Plygun, S., Sahab Uddin, M., & Ntsefong, G. N. (2020). Anti-inflammatory, antibacterial, toxicological profile, and in silico studies of dimeric naphthoquinones from *diospyros lotus*. *BioMed Research International*, *2020*. <https://doi.org/10.1155/2020/7942549>
- Rauth, S., Ray, S., Bhattacharyya, S., Mehrotra, D. G., Alam, N., Mondal, G., Nath, P., Roy, A., Biswas, J., & Murmu, N. (2016). Lupeol evokes anticancer effects in oral squamous cell carcinoma by inhibiting oncogenic EGFR pathway. *Molecular and Cellular Biochemistry*, *417*(1–2), 97–110. <https://doi.org/10.1007/s11010-016-2717-y>
- Rufino, A. T., Ribeiro, M., Judas, F., Salgueiro, L., Lopes, M. C., Cavaleiro, C., & Mendes, A. F. (2014). Anti-inflammatory and chondroprotective activity of (+)- α -pinene: Structural and enantiomeric selectivity. *Journal of Natural Products*, *77*(2), 264–269. <https://doi.org/10.1021/np400828x>
- Rzepa, J., Wojtal, Ł., Staszek, D., Grygierczyk, G., Labe, K., Hajnos, M., Kowalska, T., & Waksmundzka-Hajnos, M. (2009). Fingerprint of selected *Salvia* species by HS-GC-MS analysis of their volatile fraction. *Journal of Chromatographic Science*, *47*(7), 575–580. <https://doi.org/10.1093/chromsci/47.7.575>
- Salehi, B., Upadhyay, S., Orhan, I. E., Jugran, A. K., Jayaweera, S. L. D., Dias, D. A., Sharopov, F., Taheri, Y., Martins, N., Baghalpour, N., Cho, W. C., & Sharifi-Rad, J. (2019). Therapeutic potential of α - and β -pinene: A miracle gift of nature. *Biomolecules*, *9*(11), 1–37. <https://doi.org/10.3390/biom9110738>
- Shai, L. J., Bizimenyera, E. S., Bagla, V., MCGAW, L. J., & Eloff, J. N. (2009). *Curtisia dentata* (Cornaceae) leaf extracts and isolated compounds inhibit motility of parasitic and free-living nematodes. *Onderstepoort Journal of Veterinary Research*, *76*(2), 249–256. <https://doi.org/10.4102/ojvr.v76i2.49>
- Sharopov, F. S., Wink, M., & Setzer, W. N. (2015). Radical scavenging and antioxidant

- activities of essential oil components? An experimental and computational investigation. *Natural Product Communications*, 10(1), 153–156. <https://doi.org/10.1177/1934578x1501000135>
- Shreenithi, S., Vishnupriya, V., Ponnulakshmi, R., Gayathri, R., Madhan, K., Shyamaladevi, B., & Selvaraj, J. (2019). In silico and in vivo approach to identify the antidiabetic activity of lupeol. *Drug Invention Today*, 11(5), 1113–1116.
- Siddique, H. R., & Saleem, M. (2011). Beneficial health effects of lupeol triterpene: A review of preclinical studies. *Life Sciences*, 88(7–8), 285–293. <https://doi.org/10.1016/j.lfs.2010.11.020>
- Silva, Ana Cristina Rivas da, P. M. L., Azevedo, M. M. B. de, Costa, D. C. M., Alviano, C. S., & Alviano, D. S. (2012). Biological Activities of α -Pinene and β -Pinene Enantiomers. *Molecules*, 17(6), 6305–6316. <https://doi.org/10.3390/molecules17066305>
- Silva, F. C., Rodrigues, V. G., Duarte, L. P., Lula, I. S., Sinisterra, R. D., Vieira-Filho, S. A., Rodrigues, R. A. L., Kroon, E. G., Oliveira, P. L., Farias, L. M., Magalhães, P. P., & Silva, G. D. F. (2017). Antidiarrheal activity of extracts from *Maytenus Gonoclada* and inhibition of Dengue Virus by lupeol. *Anais Da Academia Brasileira de Ciencias*, 89(3), 1555–1564. <https://doi.org/10.1590/0001-3765201720160046>
- Singh, A., Mukhtar, H. M., Kaur, H., & Kaur, L. (2020). Investigation of antiplasmodial efficacy of lupeol and ursolic acid isolated from *Ficus benjamina* leaves extract. *Natural Product Research*, 34(17), 2514–2517. <https://doi.org/10.1080/14786419.2018.1540476>
- Singh, Payal, Arora, D., & Shukla, Y. (2017). Enhanced chemoprevention by the combined treatment of pterostilbene and lupeol in B[a]P-induced mouse skin tumorigenesis. *Food and Chemical Toxicology*, 99, 182. <https://doi.org/10.1016/j.fct.2016.11.007>
- Singh, Pradeep, Mishra, G., Sangeeta, Srivastava, S., Jha, K. K., & Khosa, R. L. (2011). Traditional uses, phytochemistry and pharmacological properties of *Capparis decidua*: An Overview. *Der Pharmacia Lettre*, 3(2), 71–82.
- Sinha, K., Chowdhury, S., Banerjee, S., Mandal, B., Mandal, M., Majhi, S., Brahmachari, G., Ghosh, J., & Sil, P. C. (2019). Lupeol alters viability of SK-RC-45 (Renal cell carcinoma cell line) by modulating its mitochondrial dynamics. *Heliyon*, 5(8), e02107. <https://doi.org/10.1016/j.heliyon.2019.e02107>
- Sköld, M., Karlberg, A. T., Matura, M., & Börje, A. (2006). The fragrance chemical β -caryophyllene - Air oxidation and skin sensitization. *Food and Chemical Toxicology*, 44(4), 538–545. <https://doi.org/10.1016/j.fct.2005.08.028>
- Soni, L. K., Dobhal, M. P., Arya, D., Bhagour, K., Parasher, P., & Gupta, R. S. (2018). In vitro and in vivo antidiabetic activity of isolated fraction of *Prosopis cineraria* against streptozotocin-induced experimental diabetes: A mechanistic study. *Biomedicine and Pharmacotherapy*, 108(May), 1015–1021. <https://doi.org/10.1016/j.biopha.2018.09.099>
- Soujanya, B. (2017). Quantification of Lupeol in Selected Commercial Coloured Cultivars of Mango (*Mangifera indica* L.) Cultivated in Telangana Region. *International Journal of Pure & Applied Bioscience*, 5(4), 2141–2146. <https://doi.org/10.18782/2320-7051.5730>
- Srivastava, A. K., Mishra, S., Ali, W., & Shukla, Y. (2016). Protective effects of lupeol against mancozeb-induced genotoxicity in cultured human lymphocytes. *Phytomedicine*, 23(7), 714–724. <https://doi.org/10.1016/j.phymed.2016.03.010>
- Sudhahar, V., Kumar, S. A., & Varalakshmi, P. (2006). Role of lupeol and lupeol linoleate on lipemic-oxidative stress in experimental hypercholesterolemia. *Life Sciences*, 78(12), 1329–1335. <https://doi.org/10.1016/j.lfs.2005.07.011>
- Susilo, A., Rumende, C. M., Pitoyo, C. W., Santoso, W. D., Yulianti, M., Herikurniawan, H., Sinto, R., Singh, G., Nainggolan, L., Nelwan, E. J., Chen, L. K., Widhani, A., Wijaya, E., Wicaksana, B., Maksum, M., Annisa, F., Jasirwan, C. O. M., & Yuniastuti, E. (2020). Coronavirus Disease 2019: Tinjauan Literatur Terkini. *Jurnal Penyakit Dalam Indonesia*, 7(1), 45. <https://doi.org/10.7454/jpdi.v7i1.415>
- Sybiliska, D., Kowalczyk, J., Asztemborska, M., Ochocka, R. J., & Lamparczyk, H. (1994). Chromatographic studies of the enantiomeric composition of some therapeutic compositions applied in the treatment of liver and kidney diseases. *Journal of Chromatography A*, 665(1), 67–

73. [https://doi.org/10.1016/0021-9673\(94\)87033-0](https://doi.org/10.1016/0021-9673(94)87033-0)
- Tan, S., Wang, D., Chi, Z., Li, W., & Shan, Y. (2017). Study on the interaction between typical phthalic acid esters (PAEs) and human haemoglobin (hHb) by molecular docking. *Environmental Toxicology and Pharmacology*, 53, 206–211. <https://doi.org/10.1016/j.etap.2017.06.008>
- Taylor, P., Gupta, R., Sharma, A. K., & Sharma, M. C. (2012). Evaluation of antidiabetic and antioxidant potential of lupeol in experimental hyperglycaemia. *Natural Product Research*, 26(October 2012), 1125–1129.
- Thilakarathna, S. H., & Vasantha Rupasinghe, H. P. (2012). Anti-atherosclerotic effects of fruit bioactive compounds: A review of current scientific evidence. *Canadian Journal of Plant Science*, 92(3), 407–419. <https://doi.org/10.4141/CJPS2011-090>
- Tolo, F.M., Rukunga, G.W., Muli, F.W., Ochora, J.M., Irungu, B.N., Mutura, C.N., Wanjiku, C.K., Mungai, G.M., Ngoc Quang, Hashimoto, K., and Asakawa, Y. (2010). The antiviral activity of compounds isolated from Kenyan *Carissa edulis* (Forssk.) Vahl. *Journal of Medicinal Plants Research*, 4(15), 1517–1522. <https://doi.org/10.5897/JMPR09.065>
- Tsai, C. W., Tsai, R. T., Liu, S. P., Chen, C. S., Tsai, M. C., Chien, S. H., Hung, H. S., Lin, S. Z., Shyu, W. C., & Fu, R. H. (2017). Neuroprotective Effects of Betulin in Pharmacological and Transgenic *Caenorhabditis elegans* Models of Parkinson's Disease. *Cell Transplantation*, 26(12), 1903–1918. <https://doi.org/10.1177/0963689717738785>
- Türkez, H., & Aydın, E. (2016). In vitro assessment of cytogenetic and oxidative effects of α -pinene. *Toxicology and Industrial Health*, 32(1), 168–176. <https://doi.org/10.1177/0748233713498456>
- Tyagi, T., & Agarwal, M. (2017). Phytochemical screening and GC-MS analysis of bioactive constituents in the ethanolic extract of *Pistia stratiotes* L. and *Eichhornia crassipes* (Mart.) solms. ~ 195 ~ *Journal of Pharmacognosy and Phytochemistry*, 6(1), 195–206. <https://www.phytojournal.com/archives/?year=2017&vol=6&issue=1&ArticleId=1079>
- Ukwubile, C. A., Ikpefan, E. O., Malgwi, T. S., Bababe, A. B., Odugu, J. A., Angyu, A. N., Otal, O., Bingari, M. S., & Nettey, H. I. (2020). Cytotoxic effects of new bioactive compounds isolated from a Nigerian anticancer plant *Melastomastrum capitatum* Fern. leaf extract. *Scientific African*, 8, e00421. <https://doi.org/10.1016/j.sciaf.2020.e00421>
- Valente, M. J., de Pinho, P. G., Henrique, R., Pereira, J. A., & Carvalho, M. (2012). Further insights into chemical characterization through GC-MS and evaluation for anticancer potential of *Dracaena draco* leaf and fruit extracts. *Food and Chemical Toxicology*, 50(10), 3847–3852. <https://doi.org/10.1016/j.fct.2012.03.050>
- Van Zyl, R. L., Seatlholo, S. T., Van Vuuren, S. F., & Viljoen, A. M. (2006). The biological activities of 20 nature identical essential oil constituents. *Journal of Essential Oil Research*, 18(SPEC. ISS.), 129–133. <https://doi.org/10.1080/10412905.2006.12067134>
- Wal, A., Wal, P., & Tiwari, R. (2019). Neuropharmacological Effects of the Ethanolic Extract of Derivatives of Lupeol in Rats. *Asian Journal of Pharmaceutical and Clinical Research*, 12(1), 222. <https://doi.org/10.22159/ajpcr.2018.v12i1.28533>
- Wang, J., Cui, L., Feng, L., Zhang, Z., Song, J., Liu, D., & Jia, X. (2016). Isoalantolactone inhibits the migration and invasion of human breast cancer MDA-MB-231 cells via suppression of the p38 MAPK/NF- κ B signaling pathway. *Oncology Reports*, 36(3), 1269–1276. <https://doi.org/10.3892/or.2016.4954>
- Wang, Y., Wu, C., Zhang, Q., Shan, Y., Gu, W., & Wang, S. (2019). Design, synthesis and biological evaluation of novel β -pinene-based thiazole derivatives as potential anticancer agents via mitochondrial-mediated apoptosis pathway. *Bioorganic Chemistry*, 84, 468–477. <https://doi.org/10.1016/j.bioorg.2018.12.010>
- Wibowo, S., & Komarayati, S. (2015). SIFAT FISIKO KIMIA MINYAK CUPRESUS (*Cupressus benthamii*) ASAL AEK NAULI , PARAPAT SUMATERA UTARA (Physico-chemical Properties of Cupressus benthamii Oil from Aek Nauli , Parapat North Sumatra). *Jurnal Penelitian Hasil Hutan*, 33(2), 93–103.
- Winnacker, M., & Rieger, B. (2015). Recent Progress in Sustainable Polymers Obtained from Cyclic Terpenes: Synthesis, Properties, and Application Potential. *ChemSusChem*, 8(15), 2455–2471. <https://doi.org/10.1002/cssc.201500421>

- Xu, H. B., Ma, Y. B., Huang, X. Y., Geng, C. A., Wang, H., Zhao, Y., Yang, T. H., Chen, X. L., Yang, C. Y., Zhang, X. M., & Chen, J. J. (2015). Bioactivity-guided isolation of anti-hepatitis B virus active sesquiterpenoids from the traditional Chinese medicine: Rhizomes of *Cyperus rotundus*. *Journal of Ethnopharmacology*, 171, 131–140. <https://doi.org/10.1016/j.jep.2015.05.040>
- Xu, M., Li, X., Song, L., Tao, C., Fang, J., & Tao, L. (2020). Lupeol alleviates coxsackievirus B3-induced viral myocarditis in mice via downregulating toll-like receptor 4. *Journal of International Medical Research*, 48(4). <https://doi.org/10.1177/0300060520910908>
- Xu, Q., Li, M., Yang, M., Yang, J., Xie, J., Lu, X., Wang, F., & Chen, W. (2018). α -pinene regulates miR-221 and induces G2/M phase cell cycle arrest in human hepatocellular carcinoma cells. *Bioscience Reports*, 38(6), 1–11. <https://doi.org/10.1042/BSR20180980>
- Yang, B., Zhao, Y., Lou, C., & Zhao, H. (2016). Eupalinolide O, a novel sesquiterpene lactone from *Eupatorium lindleyanum* DC., induces cell cycle arrest and apoptosis in human MDA-MB-468 breast cancer cells. *Oncology Reports*, 36(5), 2807–2813. <https://doi.org/10.3892/or.2016.5115>
- Yang, N. Y., Zhou, G. S., Tang, Y. P., Yan, H., Guo, S., Liu, P., Duan, J. A., Song, B. S., & He, Z. Q. (2011). Two new α -pinene derivatives from *Angelica sinensis* and their anticoagulative activities. *Fitoterapia*, 82(4), 692–695. <https://doi.org/10.1016/j.fitote.2011.02.007>
- Yilmaz, B. (2019). CHEMICAL CONSTITUENTS OF AJUGA CHAMAEPITYS (L.) SCHREB GROWING IN TURKEY BY GC-MS METHOD Bilal Yilmaz Department of Analytical Chemistry, Faculty of Pharmacy, Ataturk University, Erzurum, Turkey. *International Journal of Pharmacognosy*, 6(3), 108–112. [https://doi.org/10.13040/IJPSR.0975-8232.IJP.6\(3\).108-12](https://doi.org/10.13040/IJPSR.0975-8232.IJP.6(3).108-12)
- Yim, N. H., Jung, Y. P., Kim, A., Kim, T., & Ma, J. Y. (2015). Induction of apoptotic cell death by betulin in multidrug-resistant human renal carcinoma cells. *Oncology Reports*, 34(2), 1058–1064. <https://doi.org/10.3892/or.2015.4045>
- Yunanta, R. R. K., Lukmandaru, G., & Fernandes, A. (2014). Sifat Kimia dari Kayu Shorea Retusa, Shorea Macroptera dan Shorea Macrophylla. *Jurnal Penelitian Dipterokarpa*, 8(1), 15–24. <https://doi.org/10.20886/jped.2014.8.1.15-24>
- Zehra, B., Ahmed, A., Sarwar, R., Khan, A., Farooq, U., Ali, S. A., & Al-Harrasi, A. (2019). Apoptotic and antimetastatic activities of betulin isolated from *Quercus incana* against non-small cell lung cancer cells. *Cancer Management and Research*, 11, 1667–1683. <https://doi.org/10.2147/CMAR.S186956>
- Zhang, Z., S., G., X., L., & X., G. (2015). Synergistic antitumor effect of α -pinene and β -pinene with paclitaxel against non-small-cell lung carcinoma (NSCLC). *Drug Research*, 65(4), 214–218. <http://www.embase.com/search/results?subaction=viewrecord&from=export&id=L603514883%0Ahttp://dx.doi.org/10.1055/s-0034-1377025>
- Zhanzhaxina, A. S., Suleimen, Y. M., Ishmuratova, M. Y., Iskakova, Z. B., Seilkhanov, T. M., Birimzhanova, D. A., & Suleimen, R. N. (2020). Essential oil of *Pulicaria vulgaris* (prostrata) and its biological activity. *Bulletin of the Karaganda University. "Chemistry" Series*, 99(3), 44–50. <https://doi.org/10.31489/2020ch3/44-50>
- Zhao, H., Yang, A., Liu, J., Bao, S., Peng, R., Hu, Y., Yuan, T., Hou, S., Xie, T., Zhang, Q., Chen, X., Wang, X., & Hu, L. (2020). Chartspiroton, a Tetracyclic Spiro-naphthoquinone Derivative from a Medicinal Plant Endophytic Streptomyces. *Organic Letters*, 22(10), 3739–3743. <https://doi.org/10.1021/acs.orglett.0c00696>
- Zhao, Y., Chen, R., Wang, Y., & Yang, Y. (2018). α -Pinene Inhibits Human Prostate Cancer Growth in a Mouse Xenograft Model. *Chemotherapy*, 63(1), 1–7. <https://doi.org/10.1159/000479863>
- Zhou, J. Y., Tang, F. Di, Mao, G. G., & Bian, R. L. (2004). Effect of α -pinene on nuclear translocation of NF- κ B in THP-1 cells. *Acta Pharmacologica Sinica*, 25(4), 480–484.
- Zubair, M. F., Atolani, O., Ibrahim, S. O., Adebisi, O. O., Hamid, A. A., & Sowunmi, R. A. (2017). Chemical constituents and antimicrobial properties of *Phyllanthus amarus* (Schum & Thonn). *Bayero Journal of Pure and Applied Sciences*, 10(1), 238. <https://doi.org/10.4314/bajopas.v10i1.35>

