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To:
Office of Food Additive Safety (FHS-200)
Center for Food Safety and Applied Nutrition
Food and Drug Administration
5100 Campus Drive
College Park, MD 20740

Subject: GRAS Notice for High-Purity Rebaudioside M

Dear:

In accordance with 21 CFR §170 Subpart E consisting of §170.203 through 170.285, Manus Bio Inc. hereby informs the United States Food and Drug Administration of the conclusion that High-Purity Rebaudioside M, manufactured by Manus Bio Inc., as defined in the enclosed documents, is GRAS under the specified conditions of use as a food ingredient on the basis of scientific procedures, and therefore, is not subject to the premarket approval requirements of the Federal Food, Drug, and Cosmetic Act.

Information supporting the GRAS status of the High-Purity Rebaudioside M, which includes detailed information on the notified substance and a summary of the basis of the safety of High-Purity Rebaudioside M, under the intended conditions of use, also are enclosed for review by the Agency.

I hereby certify that the enclosed electronic files were scanned for viruses prior to submission and are thus certified as being virus-free using Symantec Endpoint Protection.

Should you have any questions or concerns regarding this GRAS Notice, please do not hesitate to contact me at any point during the review process so that we may provide a response in a timely manner.

Sincerely,



Christine Santos, PhD
Chief Technology Officer
Manus Bio Inc.
csantos@manusbio.com

External use permitted

GRAS Notification for high-purity Rebaudioside M produced by enzymatic conversion of stevia leaf extract

Prepared for:

Office of Food Additive Safety (HFS-200)
Center for Food Safety and Applied Nutrition
Food and Drug Administration
5001 Campus Drive
College Park, MD
20740 USA

Date:

November 25, 2020

GRAS Notification for High-Purity Rebaudioside M Produced by Bioconversion of Stevia Leaf Extract

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GRAS Notification for High-Purity Rebaudioside M Produced by Enzymatic Conversion of Stevia Leaf Extract

Part 1.1 §170.225 Signed Statements and Certification

In accordance with 21 CFR §170 Subpart E consisting of §170.203 through 170.285, Manus Bio Inc. hereby informs the United States (U.S.) Food and Drug Administration (FDA) that the high-purity rebaudioside M ($\geq 95\%$ rebaudioside M), manufactured by Manus Bio Inc., and identified as NutraSweet M™ is not subject to the premarket approval requirements of the Federal Food, Drug, and Cosmetic Act based on Manus Bio's view that the notified substance is Generally Recognized as Safe (GRAS) under the conditions of its intended use described in Section 1.3 below. In addition, as a responsible official of Manus Bio, the undersigned hereby certifies that all data and information presented in this notice represents a complete, representative, and balanced submission, and considered all unfavorable as well as favorable information known to Manus Bio. Included is all pertinent information to the evaluation of the safety and GRAS status of high purity rebaudioside M ($\geq 95\%$ rebaudioside M) as a general-purpose sweetener, as described herein.

Signed,



November 25, 2020

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Cambridge, MA 02138
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1.2 Common Name of Notified Substance

Stevia Extract, Stevia Leaf Extract, Stevia Sweetener, Stevia Leaf Sweetener, Steviol glycosides, Rebaudioside M, Reb M, NutraSweet M

1.3 Conditions of Use

Manus Bio intends to market high-purity rebaudioside M produced via enzymatic conversion, as a steviol glycoside preparation comprised of $\geq 95\%$ rebaudioside M, for use as a general purpose sweetener in foods within the U.S.A., in accordance with current Good Manufacturing Practice (cGMP), excluding infant formulas, meat, and poultry products. The U.S. FDA has raised no questions on the use of other high purity rebaudioside M products, including other steviol glycoside preparations, as general-purpose sweetening agents. These have no restrictions on their specific food uses and use-levels. In addition, the use-levels of high-intensity sweeteners are restricted based on the technological properties of the sweetening agent (*i.e.*, sweetness potency). As a result, because the sweetness profile of high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) is comparable to the sweetness profiles of other steviol glycoside preparations, the food uses and use-levels of high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) are likely to reflect those currently permitted in the U.S.

1.4 Basis for GRAS

The U.S. FDA has reviewed the safety of over 50 different steviol glycoside preparations and have consistently raised no objections regarding the GRAS status of steviol glycosides for use as general-purpose sweeteners in food and beverage products. Of note, the U.S. FDA did not raise any objections regarding GRN 667 and GRN 745, in relation to the GRAS status of rebaudioside M produced by enzymatic conversion for use as a general-purpose sweetener in foods (U.S. FDA, 2017, 2018). The rebaudioside M described in GRN 667 and 745 is similar to Manus Bio's high purity rebaudioside M ($\geq 95\%$ rebaudioside M) produced by enzymatic conversion of steviol glycosides. In the Manus Bio process the food ingredient is produced by conversion of stevia leaf extract using UDP-glucosyltransferase enzymes produced by *E. coli* K-12 similar to GRN 745.

Pursuant to Title 21, Section 170.30 of the Code of Federal Regulations (CFR), high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) produced via enzymatic conversion has been determined to be GRAS by Manus Bio for use as a general-purpose sweetener in foods, on the basis of scientific procedures. This GRAS determination is based on information generally available in the public domain pertaining to the safety of steviol glycosides and the production process, as discussed herein. It is also based on consensus among a panel of experts (the GRAS Panel) who are qualified by scientific training and experience to evaluate the safety of high purity rebaudioside M ($\geq 95\%$ rebaudioside M) as a general-purpose sweetener. **[See Appendix A. titled "GRAS Panel Report Concerning the Generally Recognized as Safe (GRAS) Status of high-purity Rebaudioside M ($\geq 95\%$ Rebaudioside M) produced by enzymatic conversion of stevia leaf extract."**

1.5 Availability of Information

The data and information that serve as the basis for this GRAS Notification will be made available to the U.S. FDA for review and copying upon request during business hours at the offices of:

Manus Bio Inc.
1762 Lovers Lane
Augusta, GA. 30901

In addition, Manus Bio will supply additional data and information to the FDA as may be requested to address any questions regarding this notification during or after the Agency's review of the notice.

1.6 Freedom of Information Act, 5 U.S.C. 552

It is Manus Bio's view that all data and information presented in Parts 2 through 7 of this notice do not contain any trade secret, commercial, or financial information that is privileged or confidential, and therefore all data and information presented herein are not exempt from the Freedom of Information Act, 5 U.S.C. 552.

Part 2. §170.230 Identity, Method of Manufacture, Specifications, and Physical or Technical Effect

2.1 Identity

2.1.1 Common or Usual Name

Stevia Extract, Stevia Leaf Extract, Stevia Sweetener, Stevia Leaf Sweetener, Steviol glycosides, Rebaudioside M, Reb M, NutraSweet M

2.1.2 Chemical and Physical Characteristics

Manus Bio's high-purity rebaudioside M ($\geq 95\%$ rebaudioside M; Reb M) is produced by enzymatic conversion of steviol glycosides extracted from the leaves of *Stevia rebaudiana* Bertoni using genetically modified *Escherichia coli* strains derived from *E. coli* K-12. The high purity rebaudioside M ($\geq 95\%$ rebaudioside M) material is a white-to-off-white powder that has a clean taste with no abnormal or off odor and is freely soluble in water. High purity rebaudioside M ($\geq 95\%$ rebaudioside M) is approximately 250 times sweeter than sucrose, which is consistent with the sweetness profile of steviol glycosides (FAO, 2016).

All steviol glycosides share a common steviol backbone and differ only with respect to the type and number of glycoside units they contain (*i.e.*, glucose, xylose, rhamnose, fructose, deoxyglucose, galactose, and/or arabinose) conjugated at positions R₁ and R₂. Due to the common steviol backbone, all steviol glycosides share a similar metabolic fate. The general structure for steviol glycosides is shown in Figure 2.1.2-1.

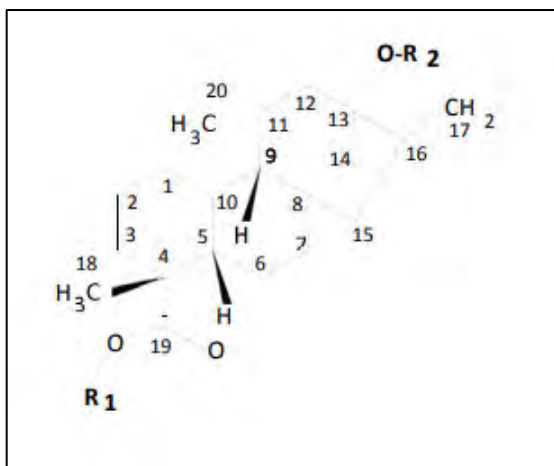


Figure 2.1.2-1 Chemical Structure of Steviol Glycosides

R₁ and R₂ may be a single or multiple glycoside unit, including glucose, xylose, rhamnose, fructose, deoxyglucose, galactose, and/or arabinose.

Rebaudioside M contains 3 glucose units each at R₁ and R₂ (Figure 2.1.2-2). It should be noted that Manus Bio's high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) is a highly purified product that contains $\geq 95\%$ Reb M, which is consistent with the purity criteria for steviol glycosides as established by the Joint FAO/WHO Expert Committee on Food Additives (JECFA) (JECFA, 2017a).

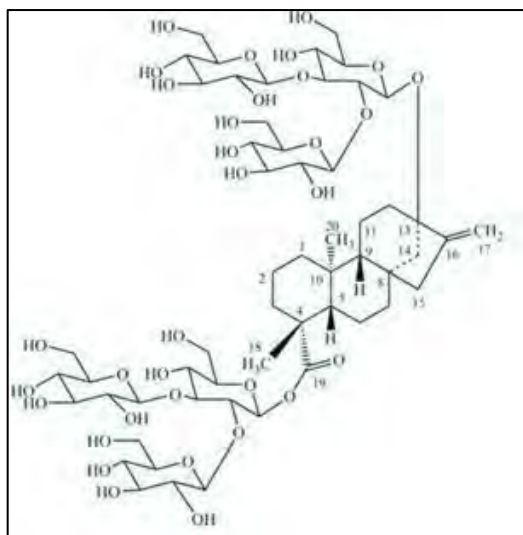


Figure 2.1.2-2 Chemical Structure for Rebaudioside M

2.2 Method of Manufacturing

2.2.1 Raw Materials and Processing Aids

All raw materials, processing aids (Table 2.2.1-1), and purification equipment used to manufacture Manus Bio's rebaudioside M through enzymatic conversion of steviol glycosides are food-grade ingredients, permitted by U.S. regulation, have GRAS status, or have been self-affirmed as safe for use in food for their respective uses. The food-grade ingredients are compliant with the specifications set forth in the Food Chemicals Codex or equivalent international food or pharmacopeia standard (e.g., JECFA, CODEX, United States Pharmacopeia, and European Pharmacopoeia).

The starting material for fermentation is a water/ethanol extract of *S. rebaudiana* sold as a food ingredient by a third-party supplier. This extract is produced in a facility with SQF, GMP, ISO9001, ISO 14001, HACCP, HALAL and Kosher certifications. The manufacturing process for this starting material is described in detail in GRAS Notice (GRN) 275 (U.S. FDA, 2008). In brief, steviol glycosides are extracted from the stevia leaf by a series of crushing, dissolution, solvent extraction, and precipitation steps that are consistent with the methodology outlined in the CTA for steviol glycosides (FAO, 2016). Since the product is specified to contain $\geq 90\%$ total steviol glycosides, it does not meet the purity requirements specified by JECFA (Joint European Committee on Food Additives, 2016; Joint FAO/WHO Expert Committee on Food Additives, 2016); however, following enzymatic conversion and the subsequent purification steps the final product generated is a highly purified steviol glycosides ($\geq 95\%$) preparation which is recognized under numerous GRAS notifications (GRNs 252, 253, 275, 278, 282, 287, 303, 304, 318, 323, 329, 337, 348, 349, 354, 365, 367, 369, 375, 380, 388, 389, 393, 395, 418, 448, 452, 456, 461, 467, 473, 493, 512, 516, 536, 548, 555, 607, 619, 626, 632, 638, 656, 662, 667, 702, 715, 733, 744, 745, 759, 780, 799, 812, 846, 867, 878, 882, 911) These purified glycosides of $\geq 95\%$ purity meet the joint FAO/WHO requirements¹.)

¹ Compliant with the specifications set forth in the Food Chemicals Codex or equivalent international food or pharmacopeia standard (e.g., JECFA, CODEX, United States Pharmacopeia, and European Pharmacopoeia).

Table 2.2.1-1 Materials used in production process

Material	Use	21CFR (U.S.FDA 2018)	Approved use
Glucose Monohydrate C6H14O7	medium nutrient	GRAS	GRAS when used in accordance with cGMP
Ammonium sulfate (NH4)2SO4	medium nutrient	582.1143/184.1443	GRAS when used in accordance with cGMP
Potassium phosphate monobasic KH2PO4	medium nutrient	160.110	GRAS when used in accordance with cGMP
Potassium phosphate dibase, K2HPO4	medium nutrient	160.110	GRAS when used in accordance with cGMP
Citric Acid C6H8O7	medium nutrient	184.1033	pH control agent & processing aid with no limitation other than cGMP
Yeast Extract	medium nutrient	184.1983	GRAS when used in accordance with cGMP
Antifoam	media	173.340	processing when used in accordance with cGMP. Secondary direct food additive, defoaming agent.
Manganese(II) chloride tetrahydrate MnCl2.4H2O	medium nutrient	582.5446/184.1446	GRAS when used in accordance with cGMP
Zinc sulfate monohydrate ZnSO4.H2O	medium nutrient	582.5997/182.8997	GRAS when used in accordance with cGMP
Copper(II) chloride pentahydrate Cl2CuH10O5	medium nutrient	184.1261	GRAS when used in accordance with cGMP
Calcium chloride dihydrate CaCl2.2H2O	medium nutrient	582.1193/582.6193/184.1193	GRAS when used in accordance with cGMP
Peptone	medium nutrient	21CFR 184.1553	GRAS when used in accordance with cGMP
Sodium molybdate dihydrate Na2MoO4.2H2O	medium nutrient	40CFR 180.920	GRAS when used in accordance with cGMP
Iron(II) sulfate heptahydrate FeSO4.7H2O	medium nutrient	184.1315	GRAS when used in accordance with cGMP
Magnesium sulfate Heptahydrate H14MgO11S	medium nutrient	582.5443/184.1443	GRAS when used in accordance with cGMP
Thiamine Hydrochloride C12H18Cl2N4OS	medium nutrient	582.5875/184.1875	Nutrient suppliment with no limitation when used in accordance with cGMP
Stevia Extract	media	170.30	GRAS when used in accordance with cGMP
190 Proof Ethanol CH3CH2OH	Elution Slovent Crystalization	184.1293	GRAS when used in accordance with cGMP
Sucrose C12H22O11	medium nutrient	GRAS	GRAS when used in accordance with cGMP
Sodium Hydroxide NaOH	medium nutrient	184.1205	GRAS when used in accordance with cGMP
Sulfuric Acid H2SO4	medium nutrient	184.1293	GRAS when used in accordance with cGMP
Sodium Benzoate C7H5NaO2	Preservative	184.1733	Used as an antimicrobial agent at levels not to exceed GMP (typically 0.1% in food)
Potassium Sorbate C6H7KO2	Preservative	182.364	GRAS when used in accordance with cGMP
Sodium Carbonate Monohydrate CH2Na2O4	medium nutrient	184.1742	GRAS when used in accordance with cGMP
Glycerin C3H8O3	medium nutrient	182.132	GRAS when used in accordance with cGMP
Potassium Chloride KCl	medium nutrient	184.1622	GRAS when used in accordance with cGMP
Adsorption resin	Purification	173.65	GRAS when used in accordance with cGMP
Ion-exchange resin	Purification	173.25	GRAS when used in accordance with cGMP
Activated carbon	Decolorizing agent	GRAS	used in accordance with cGMP
Microfiltration/ Purification	Ultrafiltration	177.291	used in accordance with cGMP
Boiler chemicals	anticorrosion	173.31	used in accordance with cGMP (non contact)

2.2.2 Production organism

Parental Strain

The parental strain *E. coli* K-12 sub-strain MG1655 Fnr⁻ was obtained from the *E. coli* Genetic Stock Center (CGSC) and is currently listed under the designation CGSC 6300.

Production Strain

The parental strain *E. coli* K-12 was engineered to express enzymes (UDP-glucosyl transferases) for the glycosylation of steviol glycosides and to improve the overall production efficiency of rebaudioside M. In addition, the strain was engineered to increase the supply of uridine diphosphate glucose (UDP-Glu), a precursor required for glycosylation of steviol glycosides through a series of gene deletions and overexpressions. All heterologously overexpressed genes originated from biosafety level 1 organisms that are not associated with any known allergens or toxins, including *Stevia rebaudiana*, *Oryza sativa*, *Glycine max*, and *Bifidobacterium bifidum*.

Engineering Methods

Overexpressed genes were synthesized, codon-optimized for *E. coli*, and introduced into stable, non-essential regions of the genome via standard techniques utilizing homologous recombination with positive selection and counter selectable markers (Datsenko and Wanner, 2000). These regions include but are not limited to *endA*, *recA*, and *araA*. All selection markers used during the engineering process were removed, and no antibiotic resistance markers are present in the final production strain.

Gene deletions were generated using standard techniques utilizing homologous recombination with positive selection and counter selectable markers (Datsenko and Wanner, 2000). All selection markers used during the engineering process were removed, and no antibiotic resistance markers are present in the final production strain. The identity of the final production strain was confirmed by Sanger sequencing of modified regions and by whole genome sequencing.

History of safe use of E. coli K-12

E. coli K-12 is a non-pathogenic and non-toxigenic organism belonging to Biosafety Level 1 according to the National Institutes of Health (NIH, 2016). Additional supporting evidence for the safety of *E. coli* K-12 is cited in other GRAS notices. (Table 2.2.2-1). *E. coli* K-12 has a long history of safe use in the industrial production of specialty chemicals and human drugs (U.S. EPA, 1997). For example, a food enzyme preparation (chymosin) obtained from a genetically modified *E. coli* K-12 strain was affirmed as GRAS by the FDA in 1990 (Flamm, 1991; Olempska-Beer *et al.*, 2006) and has been used safely for cheese production worldwide. *E. coli* also serves as a host for the production of enzymes currently used in a GRAS-approved process for the enzymatic conversion of steviol glycosides (GRN745, 2018).

Table 2.2.2-1 GRAS Notices Using *E. coli* K-12

Various substrains and modified versions of *E. coli* K-12 have been used to produce products deemed Generally Recognized As Safe (GRAS) for food or animal use, including two RebM products (GRAS Notices [GRNs] 745 and 780). These GRNs are described in Table 2.2.2-1. Notably, in the response letter to GRN 659, FDA opined, "We extensively reviewed and determined that *E. coli* K-12 is a non-pathogenic, non-toxicogenic, and safe production strain when used in accordance with good manufacturing practices" (55 FR 10932 at 10934; March 23, 1990). In response to GRN 745 the FDA stated "Based on the information that the petitioner provided, as well as other information available to FDA, we have no questions at this time regarding the petitioner's conclusion that EMSG is GRAS under its intended conditions of use under 21 CFR 170.35. Additionally, *E. coli* K-12 is in the GRAS production of chymosin, which is used to coagulate milk in cheeses and other dairy products (21 CFR §184.1685)

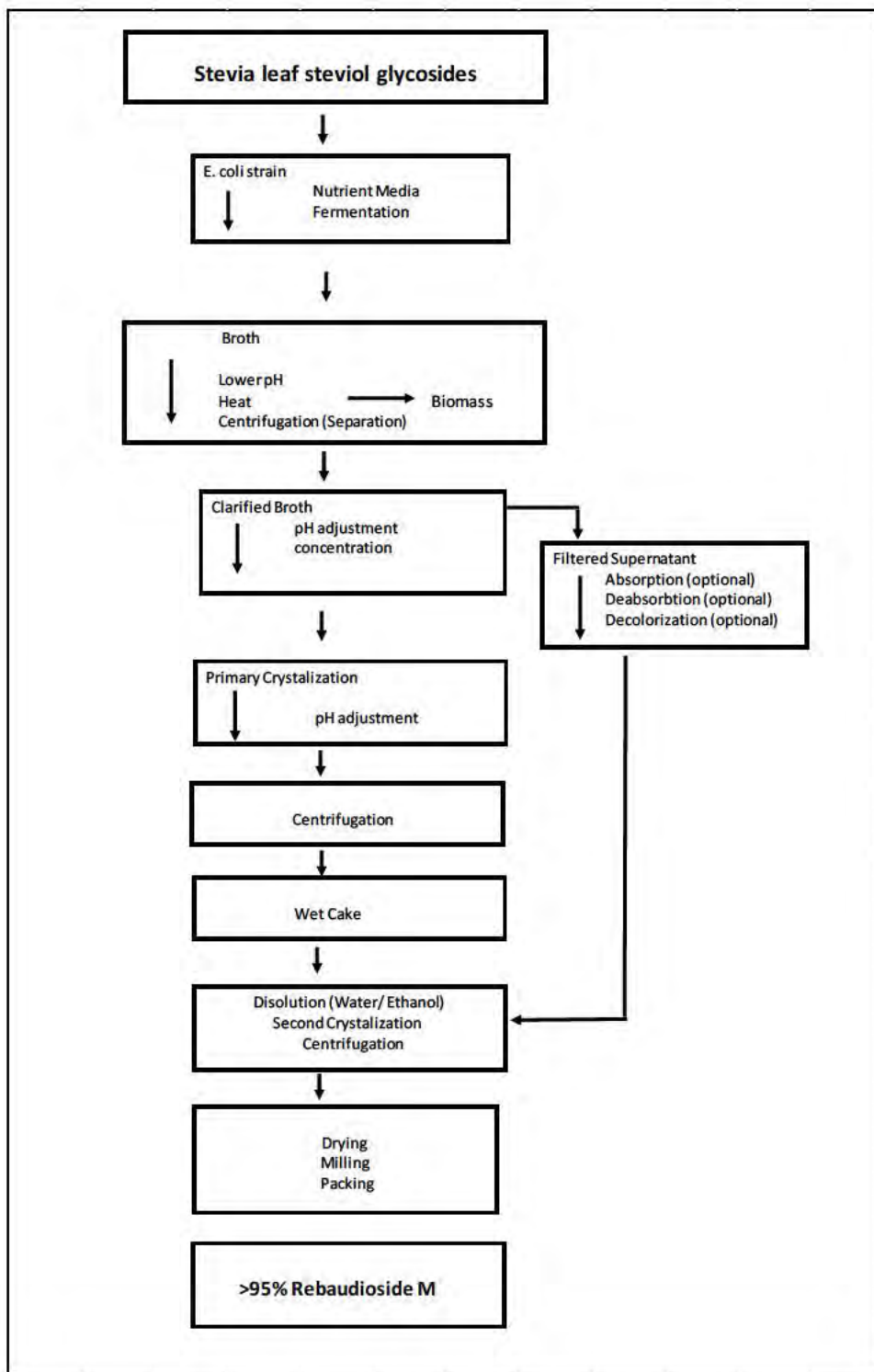
GRN	FDA Response	Product	Methods	GRN	FDA Response	Product	Methods
GRN 624	No questions	D-allulose 3-epimerase enzyme preparation	Substrain W3110 engineered to express D-allulose 3-epimerase from <i>Arthrobacter globiformis</i> M30	GRN 780	No questions	Rebaudioside M	Engineered to express 2 glycosyltransferases and sucrose synthase; isolated enzymes utilized to produce RebM
GRN 659	No questions	Lacto-N-neotetraose	Substrain DH1 underwent 7 modifications to optimize oligosaccharide production	AGRN 16	No questions	L-methionine	Modified (not described)
GRN 735	No questions	2'-fucosyllactose	Substrain W3110 derivative GI724 modified to enhance production of fucosylated oligosaccharides	AGRN 24	No questions	L-methionine	Modified (not described)
GRN 745	No questions	Steviol glycosides consisting of primarily rebaudioside M	Substrain W3110 derivative LE1B109 engineered to express enzymes for stevioside transformations; enzymes isolated to produce RebM	CFR 184.1685	Affirmed as GRAS	Chymosin	nonpathogenic and nontoxicogenic strain of <i>Escherichia coli</i> K-12

GRAS = Generally Recognized As Safe; GRN = GRAS Notice; AGRN = Animal Food GRAS Notice

2.2.3 Manufacturing Process

Manus Bio's high-purity rebaudioside M (≥95% rebaudioside M) is produced by enzymatic conversion of steviol glycosides using an *E. coli* strain derived from *E. coli* K-12. (see Appendix B) The production strains are grown in media containing steviol glycoside extracts prepared from the leaves of *S. rebaudiana* Bertoni in accordance with the methodology outlined in the Chemical and Technical Assessment (CTA) for steviol glycosides (FAO, 2016). *E. coli* K-12 cells mediate the glycosylation of steviol glycosides in the leaf extract to rebaudioside M. The broth is then inactivated and centrifuged to separate the *E. coli* K-12 biomass from the aqueous phase. Rebaudioside M is extracted from the aqueous phase and purified through crystallization and washes. Rebaudioside M crystals are then dried and milled to a final product containing ≥95% Rebaudioside M. The purification processes described are consistent with the methodologies for the manufacture of steviol glycosides as described in the CTA published by FAO/JECFA (FAO, 2016). Manus Bio's ≥95% Rebaudioside M, produced by enzymatic conversion of steviol glycosides, is manufactured in a facility registered as a FDA Food Facility. The plant operates under cGMPs (Good Manufacturing Practices) outlined in the Food Safety Modernization Act 21 CFR 117, including required HACCP and Food Defense Plans, and is subject to audit from regulatory authorities including the US FDA and The State of Georgia Department of Agriculture. Manufacturing shall be certified to a GFSI (Global Food Safety Initiative) compliant audit scheme. An overview of the manufacturing process flow is provided in Figure 2.2.3.1 below.

Figure 2.2.3.1 Manufacturing process of Manus Bio's rebaudioside M produced by enzymatic conversion.



2.3 Product Specifications and Batch Analyses

2.3.1 Product Specifications

Appropriate food-grade specifications have been established for high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) produced via enzymatic conversion based on the specifications for steviol glycosides established by JECFA (2017a) (Table 2.3.1-1). All analytical methods used to measure each specification parameter are internationally- recognized methods (e.g., United States Pharmacopeia [USP], Association of Official Analytical Chemists [AOAC], or JECFA). Total steviol glycoside content is measured using the high-performance liquid chromatography (HPLC) method described in the JECFA specification monograph for steviol glycosides from *S. rebaudiana* Bertoni (JECFA, 2017a,b).

Table 2.3.1-1 Product Specifications for High-Purity Rebaudioside M ($\geq 95\%$ Rebaudioside M) produced via Enzymatic Conversion of Stevia Leaf Extract

Specification Parameter	High-Purity Rebaudioside M ($\geq 95\%$ Rebaudioside M)	Current JECFA Specifications For Steviol Glycosides (JECFA, 2017a)	Method of Analysis
Physical and Chemical Parameters			
Appearance	White to off-white powder	White to light yellow powder	
Total steviol glycosides (anhydrous basis)	$\geq 95\%$	$\geq 95\%$ total steviol glycosides ^a	TN34236 [Monograph 19 (82 nd JECFA Meeting 2016)]
Rebaudioside M	$\geq 95\%$	N/A	TN34240 [Monograph 19 (82 nd JECFA Meeting 2016)]
Loss on drying	$\leq 6\%$	$\leq 6\%$ (105°C, 2h)	TN46040 (CRA E-46)
pH (1% solution)	4.5 to 7.0	4.5 to 7.0	TN60730 (AOAC 981.12)
Residual ethanol	$\leq 5,000$ ppm ($\leq 0.5\%$)	$\leq 0.5\%$	TN64080 (USP 32-NF 27)
Residual methanol	≤ 200 ppm ($\leq 0.02\%$)	$\leq 0.02\%$	TN64080 (USP 32-NF 27)
Total ash	$\leq 1\%$	$\leq 1\%$	TN09560 (AOAC 900.02)
Lead	≤ 1 ppm	≤ 1 ppm	TN44290 (AOAC 993.14)
Arsenic	≤ 1 ppm	≤ 1 ppm	TN44292 (AOAC 993.14)
Cadmium	≤ 1 ppm	NS	TN44291 (AOAC 993.14)
Mercury	≤ 1 ppm	NS	TN44293 (AOAC 993.14)
Microbiological Parameters			
Specification Parameter	High-Purity Rebaudioside M ($\geq 95\%$ Rebaudioside M)	Current JECFA Specifications for Steviol Glycosides (JECFA, 2017a)	Method of Analysis
Total plate count	$< 1,000$ CFU/g	$< 1,000$ CFU/g	TN10560 (CRA Microbiological Methods I-A)
Mold	< 100 CFU/g	< 200 CFU/g	TN47010 (CRA Microbiological Methods II-A-1)
Yeast	< 100 CFU/g	< 200 CFU/g	TN97010 (CRA Microbiological Methods I-A)
Coliforms	< 3 MPN/g	NS	TN10510 (CRA Microbiological Methods IV-B)
<i>Escherichia coli</i>	Not detected	Not detected	TN10512 (CRA Microbiological Methods IV-B)
<i>Salmonella</i>	Negative/25 g	Not detected	TN10547 (CRA Microbiological Methods V-A)

AOAC = Association of Official Analytical Chemists; CFU = colony-forming units; CRA = Corn Refiners Association; JECFA = Joint FAO/WHO Expert Committee on Food Additives; MPN = most probable number; N/A = not applicable; NS = not specified; ppm = parts per million; USP = United States Pharmacopeia.

^A Where steviol glycosides “consists of a mixture of compounds containing a steviol backbone conjugated to any number or combination of the principal sugar moieties in any of the orientations occurring in the leaves of *Stevia rebaudiana* Bertoni including, glucose, rhamnose, xylose, fructose, deoxyglucose, galactose, and arabinose”. (JECFA, 2017a).

2.3.2 Batch Analyses

Analysis of 3 non-consecutive lots of high-purity rebaudioside M (≥95% rebaudioside M) produced by Manus Bio demonstrates that the manufacturing process produces a consistent product which meets the established product specifications. A summary of the batch analyses is presented in Table 2.3.2-2. Product analysis has confirmed that Manus Bio’s steviol glycoside product is comprised of rebaudioside M, other steviol glycosides and water.

(see appendix C for certificates of analysis)

Table 2.3.2-2 Summary of the Product Analysis for 3 Non-Consecutive Lots of High-Purity Rebaudioside M (≥85% Rebaudioside M)

Specification Parameter	Limit	Manufacturing Lot No.		
		MAM060420A	MAM060520E	MAM200716J
Appearance	White-to-off-white powder	Pass	Pass	Pass
Total steviol glycosides (anhydrous basis)	≥95%	>95	>95	>95
Rebaudioside M	≥95%	97.93	97.01	97.2
Loss on drying	≤6%	4.2	3.98	1.74
pH (1% solution)	4.5 to 7.0	Pass	Pass	Pass
Residual ethanol	≤5,000 ppm	Not Detected	Not Detected	Not Detected
Residual methanol	≤200 ppm	Not Detected	Not Detected	Not Detected
Total ash	≤1%	<1%	<1%	<1%
Lead	≤1 ppm	<1 ppm	<1 ppm	<1 ppm
Arsenic	≤1 ppm	<1 ppm	<1 ppm	<1 ppm
Cadmium	≤1 ppm	<1 ppm	<1 ppm	<1 ppm
Mercury	≤1 ppm	<1 ppm	<1 ppm	<1 ppm
Total plate count	<1,000 CFU/g	50 CFU/g	70 CFU/g	500 CFU/g

Table 2.3.2-2 Summary of the Product Analysis for 3 Non-Consecutive Lots of High-Purity Rebaudioside M (≥95% Rebaudioside M)

Specification Parameter	Limit	Manufacturing Lot No.		
		MAM060420A	MAM060520E	MAM200716J
Mold	<100 CFU/g	<10 CFU/g	<10 CFU/g	<10 CFU/g
Yeast	<100 CFU/g	<10 CFU/g	<10 CFU/g	<50 CFU/g
Coliforms	<3 MPN/g	<3 MPN/g	<3 MPN/g	<3 MPN/g
<i>Escherichia coli</i>	Not detected	Not detected	Not detected	Not detected
<i>Salmonella</i>	Negative/25 g	Negative	Negative	Negative

CFU = colony-forming units; MPN = most probable number; NA = not applicable; ppm = parts per million.

2.3.3 Residual Protein Analysis

To confirm the removal of residual protein from the final product, 3 non-consecutive batches of the high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) produced by enzymatic conversion of steviol glycosides (Lot Numbers MAM060420A, MAM060520E, MAM200716J) were analyzed using the bicinchoninic acid (BCA) assay method. Samples were reconstituted to a concentration of 1 g/L and measured using a 96 well plate format BCA assay on a Tecan Infinite M1000 Pro plate reader. The limit of detection for the assay was 22.5 ppm on a w/v basis. The results of the analysis indicated that the levels of protein within the high-purity rebaudioside M material were below the limit of detection, which provides further evidence that the downstream processing successfully removed any residual proteins from the final product (for residual protein analysis see appendix D).

Table 2.3.4.1 Protein Analysis



2.3.4 Residual DNA Analysis

To confirm the absence of residual DNA in rebaudioside M, 3 non-consecutive lots of final product (Lot NO's MAM060520E, MAM200716J, MAM060420A) were assayed by polymerase chain reaction (PCR). Primer design and PCR conditions were designed to amplify a specific fragment of one of the genes inserted in the production strain. Extracted genomic DNA containing the selected gene was used as a positive control. Rebaudioside M samples were prepared in aqueous solution and assayed by PCR at a final concentration of 100 mg/L. Samples were assayed with or without positive control DNA at a final concentration of 1 ng/ μ L. A standard curve was prepared from 1 ng/ μ L to 0.1fg/ μ L to determine the limit of detection of the assay (320 fg/ μ L). All samples with a positive control spike showed an expected PCR product, whereas all the Reb M samples without a genomic DNA spike showed no product. The gel images were analyzed using ImageJ for quantitative analysis. Background subtracted mean grey values from each of the

samples were fit to values from the standard curve and found to be below the limit of detection of the assay. These results confirm the absence of residual genomic DNA in the Rebaudioside M final product above the analytical limit of detection of 320 fg/μL. (For residual DNA analysis see appendix E).

Table 2.3.4-2 **Absence of residual DNA**

Sample	Background Subtracted Mean Grey Value	Calculated Concentration (ng/μL)
1.MAM060420A	-0.877	below LOD
2.MAM060520E	-0.771	below LOD
3.MAM200716J	-0.431	below LOD

*LOD = limit of detection

2.3.5 Pesticide Residue

A combined LC & MS / GC & MS screen using QuEChERS method for pesticide residues was conducted on the starting stevia leaf material (≥90% total steviol glycosides; Lot # 202003051 and Lot #201901171) demonstrated the absence of residues of commonly used pesticides in the product. (see Appendix F for pesticide residue data).

2.4 Stability Data

A number of scientific and authoritative bodies, including JECFA, the European Food Safety Authority (EFSA), and Food Standards Australia/New Zealand (FSANZ), have reviewed the stability of steviol glycosides. The stability of steviol glycosides also are discussed in several published studies (Chang and Cook, 1983; Kroyer, 1999; Oehme *et al.*, 2017). At the 68th meeting, JECFA evaluated the stability of steviol glycosides under conditions mimicking their use in foods and noted that steviol glycosides do not undergo browning or caramelization when heated and are stable under elevated temperatures (JECFA, 2007). In addition, steviol glycosides (approximately 90 to 94% purity) are stable for at least 180 days when stored at temperatures up to 24°C and pH 2.0 to 4.0. However, at elevated temperatures (80°C), steviol glycoside solutions maintained in water and pH 4.0 and 3.0 for 8 hours showed 4 and 8% decomposition, respectively. At temperatures of 100°C, higher rates of decomposition were observed, with 10 and 40% decomposed at pH 4.0 and 3.0, respectively. These results indicate that the stability of steviol glycosides is pH and temperature dependent. Based on the available evidence, JECFA concluded that steviol glycosides are thermally and hydrolytically stable for use in foods and acidic beverages under normal processing and storage conditions. (JECFA, 2007)

The U.S. FDA has reviewed the stability of high-purity rebaudioside M preparations in previous GRAS notices (GRN812, GRN759, GRN745, GRN744, GRN512, GRN882, GRN780 and GRN 846) (U.S. FDA, 2018, 2018, 2014, 2019, 2018, 2019). There exists a number of studies on the stability of steviol glycosides, including stevioside, rebaudioside A, and rebaudioside M, under different storage conditions (*e.g.*, in different forms, such as powder and solution, in acidic conditions, and various temperatures) and in the publicly-available scientific literature (Wood *et al.*, 1955; Chang and Cook, 1983; Kinghorn, 2002; Cargill, 2008; Merisant, 2008; Chaturvedula *et al.*, 2013; Prakash *et al.*, 2014). These studies are discussed in detail in GRN512, GRN667, GRN780, and GRN846 and are incorporated by reference in this notice. Ultimately, the results of these stability studies suggest that the stability of steviol glycosides is pH- and temperature-dependent, which are consistent with the conclusions of JECFA (2007). More recently, a study evaluating the structural

stability of 3 commercial batches each of the dried stevia leaves, the first aqueous infusion of the ground stevia, and a high-purity stevia leaf extract ($\geq 95\%$ steviol glycosides) confirmed that the processing steps do not chemically alter or modify the steviol glycoside content (Oehme *et al.*, 2017).

In addition to the stability studies within the scientific literature, storage stability studies on rebaudioside M were discussed in GRN 512 and GRN 667. GRN 512 presented the results of a stability test on 1 batch of Rebpure™ RM95, which contains $\geq 95\%$ rebaudioside M (U.S. FDA, 2014). In this study, a sample of Rebpure™ RM95 was stored at $25 \pm 5^\circ\text{C}$ and relative humidity of $60 \pm 5\%$ for up to 8 weeks. The results of the study demonstrate that the rebaudioside M and the total steviol glycoside content remained $\geq 95\%$ over the course of the 8-week study period. GRN 667 presented the results of an accelerated storage stability study on rebaudioside M ($\geq 95\%$ rebaudioside M) when stored at $40 \pm 2^\circ\text{C}$ and relative humidity of $75 \pm 5\%$ for up to 6 months (U.S. FDA, 2016). Over the course of the accelerated stability study, rebaudioside M was observed to be stable in that the rebaudioside M content did not change over the 6-month period and remained $\geq 95\%$ rebaudioside M.

The results of these storage stability studies are consistent with the results reviewed by JECFA (2007) in that the stability of steviol glycosides, including rebaudioside M, are thermally stable under normal storage conditions.

Furthermore, while the rebaudioside M content of Manus Bio’s high-purity rebaudioside M and that of the preparations described in GRN 512, GRN 667, GRN 745, GRN 759, GRN 780, GRN 799, GRN 812, GRN 846, & GRN 882 are similar in all rebaudioside M preparations (*i.e.*, $\geq 85\%$ to $\geq 95\%$ total steviol glycosides), Rebaudioside M is expected to exhibit similar chemical stability to other closely related steviol glycosides (*e.g.* stevioside & rebaudioside A) based on their chemical structure similarity. Therefore, it is anticipated that the results of the stability studies on the rebaudioside M preparations described in GRN 512, GRN 667, be GRN 745, GRN 759, GRN 780, GRN 799, GRN 812, GRN 846, & GRN 882 and the results of the stability studies available in the publicly-available scientific literature, can be extended to support the stability of Manus Bio’s high-purity rebaudioside M ($\geq 95\%$ rebaudioside M).

Manus Bio is currently conducting an accelerated stability study on 1 batch of high purity rebaudioside M ($\geq 95\%$ rebaudioside M) produced by enzymatic conversion of steviol glycosides (Lot No. MAM0605520E). In this study, samples of approximately 25 g of high purity rebaudioside M ($\geq 95\%$ rebaudioside M) are stored at 50°C in polypropylene jars with screw top lids and heat sealed in order to mimic the commercial packaging. Total steviol glycosides and Rebaudioside M content have been measured by HPLC at baseline and at 1, and 3 months. The moisture content was measured by gravimetric moisture balance.

Preliminary results at 3 months indicate no significant changes in Rebaudioside M or steviol glycosides content (Table 2.4-1). The accelerated stability study is currently on-going.

Table 2.4-1 Results of an Accelerated Stability Study on 1 Batch of High-Purity Rebaudioside M ($\geq 95\%$ Rebaudioside M) (Lot No MAM 060520E) (Study Currently On-going)

Parameter	Month			
	0 (baseline)	1	3	6
Total steviol glycosides (%) (dry basis)	98.49	98.49	97.27	TBD
Rebaudioside M (%) (dry basis)	97.01	97.01	96.79	TBD

TBD = to be determined.

Part 3. §170.235 Dietary Exposure

3.1 Intended Use of High-Purity Rebaudioside M (≥95% Rebaudioside M) and Levels of Use in Foods

High purity rebaudioside M (≥95% rebaudioside M) is intended for use as a general-purpose sweetener in accordance with cGMP, excluding infant formulas, meat and poultry products. High purity rebaudioside M (≥95% rebaudioside M) has a sweetness intensity of approximately 250 times that of sucrose. To date, the U.S. FDA raised no questions regarding the use of other high-intensity sweeteners, including other steviol glycoside preparations, as general-purpose sweeteners that have no restrictions on their specific food uses and use-levels. In addition, the use-levels of high-intensity sweeteners are restricted based on the technological properties of the sweetening agent (*i.e.*, sweetness potency). Therefore, considering that steviol glycosides, including the ingredient that is the subject of this GRAS notice, are characterized by a sweetness profile that is, for the most part, comparable to other high-intensity sweeteners, the uses and use-levels of high-purity rebaudioside M (≥95% rebaudioside M) produced by enzymatic conversion of steviol glycosides will likely reflect those currently permitted for other high-intensity sweeteners in the U.S.

3.2 Estimated Consumption of High-Purity Rebaudioside M (≥95% Rebaudioside M) Based upon Intended Food Uses

3.2.1 History of Consumption of Steviol Glycosides

Stevia rebaudiana and the individual steviol glycosides derived from the plant have been consumed as sweeteners in various foods and beverages since the late 1800s (Geuns, 2003). According to Blumenthal (1995) and Geuns (2003), the native peoples of Brazil and Paraguay have consumed the *S. rebaudiana* plant for hundreds of years as a food ingredient and as a tea. Similarly, *S. rebaudiana* became a popular herbal tea ingredient in the U.S. in the 1980s (Blumenthal, 1995; Ferlow, 2005). Stevioside, the first isolated steviol glycoside from the *S. rebaudiana* leaf, has been consumed in Japan for more than 30 years (Geuns, 2003; Ferlow, 2005). Approximately 160,000 metric tons of stevioside, as sucrose equivalents, were reportedly consumed in Asia in 1995; in 1999, this level increased to approximately 200,000 metric tons as sucrose equivalents (International Sugar Organization, 2001).

3.2.2 Estimated Consumption of High-Purity Rebaudioside M (≥95% Rebaudioside M) from Proposed Food Uses

The dietary consumption of various steviol glycoside preparations has been estimated using a post-market surveillance approach as outlined in a number of GRAS notices for steviol glycosides submitted to the U.S. FDA (*e.g.*, GRN 667, 715, 733, 745, 759, 780, 799, 812, 846, and 882). Generally, this approach uses the data from Renwick (2008) in which dietary exposure to Rebaudioside A was estimated based on the available post-market surveillance data for other high-intensity sweeteners, and by assuming full replacement of the currently approved high-intensity sweeteners with the new sweetener [*i.e.*, high-purity rebaudioside M (≥95% rebaudioside M)]. While conservative, this approach yields intake estimates that are realistic as they reflect actual post-market intakes of high-intensity sweeteners. Renwick (2008) estimated the average and high-end dietary intakes of Rebaudioside A as sucrose equivalents in various population groups, such as non-diabetic and diabetic adults and children, and adjusted the values accordingly using the sweetness intensity of Rebaudioside A relative to sucrose (approximately 200).

This post-market surveillance approach can be used to estimate the dietary intakes of high purity rebaudioside M ($\geq 95\%$ rebaudioside M) (Table 3.2.2-1) in similar population groups. Manus Bio determined that high purity rebaudioside M (Nutrasweet M™) ($\geq 95\%$ rebaudioside M) is approximately 250 times sweeter than sucrose. This is based on the results of a sweetness potency test conducted with an expert taste panel at Brison, an independent third-party sensory company.

The estimated intake values for high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) were calculated based on the sweetness potency and the molecular weight of Rebaudioside M.

Table 3.2.2-1 Estimated Consumption High-Purity Rebaudioside M ($\geq 95\%$ Rebaudioside M) Using the Intense Sweetener Intake Assessment Methodology described by Renwick (2008)

Population Group	Intakes of Intense Sweeteners (expressed as sucrose equivalents) (mg/kg bw/day)		Consumption Estimates			
			High-Purity Rebaudioside M ($\geq 95\%$ Rebaudioside M) ^a (mg/kg bw/day)		High-Purity Rebaudioside M ($\geq 95\%$ Rebaudioside M) ^{a,b} (mg/kg bw/day) as Steviol	
	Average Consumer	High Consumer	Average Consumer	High Consumer	Average Consumer	High Consumer
Non-diabetic adults	255	675	1.02	2.7	0.26	0.68
Diabetic adults	280	897	1.12	3.59	0.28	0.9
Non-diabetic children	425	990	1.7	3.96	0.43	0.99
Diabetic children	672	908	2.69	3.63	0.67	0.91

bw = body weight.

^a Approximately 250 times as sweet as sucrose.

^b Calculated based on the molecular weights of steviol (318.45 g/mol) and Reb M (1,291.3 g/mol) [steviol conversion factor of 0.25].

For non-diabetic adults, average and high-end intakes of high purity rebaudioside M ($\geq 95\%$ rebaudioside M) of up to 0.26 and 0.68 mg/kg body weight/day expressed as steviol equivalents, respectively, were calculated. For diabetic adults, average and high-end intakes were slightly higher at up to 0.28 and 0.90 mg/kg body weight/day. Average and high-end exposures to high purity rebaudioside M ($\geq 95\%$ rebaudioside M), expressed as steviol equivalents, in non-diabetic children were calculated to be up to 0.43 and 0.99 mg/kg body weight/day, respectively. Although average intakes of high purity rebaudioside M ($\geq 95\%$ rebaudioside M), expressed as steviol equivalents, were estimated to be higher at up to 0.67 mg/kg body weight/day in diabetic children compared to values for non-diabetic children, high-end values in diabetic children (0.91 mg/kg body weight/day) were lower than high-end values in non-diabetic children. The predicted intakes of high-purity rebaudioside M ($\geq 95\%$ rebaudioside M), expressed as steviol equivalents, are all below the current acceptable daily intake (ADI) defined by the JECFA for steviol glycosides (FAO, 2016) of 0 to 4 mg/kg body weight/day as steviol.

In 2008, JECFA considered various intake models for the estimation of dietary exposure to steviol glycosides, including the intake analysis conducted by Renwick (2008) as part of their evaluation of the safety of steviol glycosides. Although higher intake estimates than those presented by Renwick (2008) were identified using other methodologies, including ones considering replacement of all sweeteners used in or as food (up to approximately 6 mg/kg body weight/day, expressed as steviol equivalents), JECFA noted that such replacement estimates were highly conservative and that actual exposures to steviol glycosides (expressed

as steviol equivalents) would be 20 to 30% of these values (1 to 2 mg/kg body weight/day, expressed as steviol equivalents). JECFA also noted that the post-market surveillance approach further confirmed the lower intake estimate range.

Part 4. §170.240 Self-Limiting Levels of Use

The use of high purity rebaudioside M ($\geq 95\%$ rebaudioside M) is largely limited by the desired sweetness intended for a particular food or beverage product. Therefore, the use of high purity rebaudioside M ($\geq 95\%$ rebaudioside M) as a general-purpose sweetener in foods is self-limiting based on its organoleptic properties.

Part 5. §170.245 Experience Based on Common Use in Food Before 1958

Not applicable as high purity rebaudioside M ($\geq 95\%$ rebaudioside M) was not used in food before 1958.

Part 6. §170.250 Narrative and Safety Information

The safety of steviol glycosides has been extensively reviewed by the U.S. FDA in a number of GRAS notices. The Agency raised no objections to 57 GRAS notices describing the GRAS status of major individual steviol glycosides, including stevioside, rebaudiosides A, C, D, E, I, M, and X/M, mixtures of steviol glycosides, and glucosylated and enzyme-modified steviol glycosides (GRNs 252, 253, 275, 278, 282, 287, 303, 304, 318, 323, 329, 337, 348, 349, 354, 365, 367, 369, 375, 380, 388, 389, 393, 395, 418, 448, 452, 456, 461, 467, 473, 493, 512, 516, 536, 548, 555, 607, 619, 626, 632, 638, 656, 662, 667, 702, 715, 733, 744, 745, 759, 780, 799, 812, 823, 846, 867, 878, 882, 911).

In addition to the U.S. FDA, the safety of steviol glycosides has been reviewed by several scientific bodies and regulatory agencies, including JECFA, European Commission's Scientific Committee on Food (SCF), EFSA, FSANZ, and Health Canada. The existing safety database on steviol glycosides includes an extensive evaluation of the metabolism and pharmacokinetics of steviol glycosides in rodents and humans, and a standard battery of toxicological tests, including acute toxicity, short- and long-term toxicity and carcinogenicity, reproductive and developmental toxicity, *in vitro* and *in vivo* mutagenicity and genotoxicity, as well as several human studies.

Much of the early studies investigating the safety of steviol glycosides were conducted with stevioside, the predominant steviol glycoside in *S. rebaudiana* leaves (Aze *et al.*, 1991; Toyoda *et al.*, 1997). Since then, additional toxicity testing has been conducted on rebaudioside A and D (Curry and Roberts, 2008; Curry *et al.*, 2008; Nikiforov and Eapen, 2008; Williams and Burdock, 2009). Due to the common metabolic fate of steviol glycosides, the scientific bodies and regulatory agencies described above have extended their safety opinions to include all steviol glycosides, rather than individual steviol glycosides (JECFA, 2017a,b).

Thus, considering that the existing safety database on steviol glycosides has been extensively reviewed by the U.S. FDA, the pertinent generally available data and information used to support the safety of steviol glycosides, including major individual steviol glycosides and other steviol glycoside mixtures/preparations, is incorporated by reference to information cited within prior GRAS notifications.

Updated searches of the scientific literature were conducted through November 2020 to identify new data and information relevant to the safety of steviol glycosides that have been published since the U.S. FDA's last review. Given the shared metabolic fate of steviol glycosides, the safety of high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) produced by enzymatic conversion of steviol glycosides can be supported based on the existing safety database for steviol glycosides, the safety conclusions for steviol glycosides by JECFA and other scientific and regulatory authorities/bodies, and the safety of the production strains. At the time of preparation of this GRAS notice, GRN 882 was the most recent steviol glycoside GRAS notice to receive a "no questions" letter from the U.S. FDA which summarized literature prior to March 2019.

6.1 Absorption, Distribution, Metabolism, and Elimination

An extensive database exists outlining the metabolic fate (absorption, distribution, metabolism, and elimination) of steviol glycosides. The available data and information on the metabolic fate of individual steviol glycosides as discussed in detail in several GRAS notices (e.g., GRN 619, 626, 667) is incorporated by reference in this dossier. Briefly, steviol glycosides are not hydrolyzed in the upper gastrointestinal tract due to the presence of β -glycosidic bonds; the unchanged steviol glycosides enter the colon and are subject to microbial degradation by the gut microflora, resulting in the release of the aglycone steviol (Wingard *et al.*, 1980; Hutapea *et al.*, 1997; Gardana *et al.*, 2003; Koyama *et al.*, 2003a,b; Geuns *et al.*, 2003, 2007; Renwick and Tarka, 2008; Nikiforov *et al.*, 2013; Purkayastha *et al.*, 2016). Specifically, Purkayastha *et al.* (2016) conducted incubation studies with pooled human fecal homogenates collected from adult males and females using rebaudiosides A, B, C, D, E, F, M, steviolbioside and dulcoside A, and showed that all glycosides were converted to steviol over a 24–48 h incubation period. These studies demonstrated that the number and location of sugar units attached to the steviol backbone do not significantly affect the microbial hydrolysis. Likewise, this study demonstrated that steviol glycosides, irrespective of the type of sugar moiety (e.g., glucose, rhamnose, xylose) attached to the steviol backbone, were metabolized to steviol at generally similar hydrolysis rates. Incubation tests were also conducted in fecal homogenates collected from adult and pediatric populations with steviol glycoside from stevia leaf extracts and steviol glycosides produced by enzymatic conversion of Reb A to larger molecules by attaching glucose units via β - or α -glycosidic bonds. The steviol glycosides produced by extraction from stevia leaf or enzymatic conversion of stevia leaf extract were also reported to share a similar metabolic fate in the fecal homogenates from adults and children supporting a similar metabolic fate of steviol glycosides in all ages (Purkayastha 2020). Steviol glycosides are hydrolyzed sequentially, in which one sugar moiety is removed at a time, with the degradation rates dependent on the structural complexity of each steviol glycoside (Wingard *et al.*, 1980; Koyama *et al.*, 2003b). Despite the differences in chemical structure, however, the rates of hydrolysis of different steviol glycosides to steviol are relatively similar, especially during the first 24 hours of incubation in *in vitro* metabolic studies with human fecal homogenates (Purkayastha *et al.*, 2014, 2015, and 2016). Following microbial degradation, the steviol metabolite is absorbed systemically into the portal vein and distributed to the liver, spleen, adrenal glands, fat, and blood (Nakayama *et al.*, 1986; Sung, 2002 [unpublished]; Koyama *et al.*, 2003b; Wang *et al.*, 2004; Roberts and Renwick, 2008). Steviol is conjugated to glucuronic acid to form steviol glucuronide in the liver. The steviol glucuronide metabolite and any unconjugated steviol or unhydrolyzed fraction of the administered glycosides are excreted primarily in the urine, and, to a lesser extent, feces in humans (Wingard *et al.*, 1980; Nakayama *et al.*, 1986; Kraemer and Maurer, 1994; Sung, 2002 [unpublished]; Geuns and Pietta, 2004 [unpublished]; Simonetti *et al.*, 2004; Geuns *et al.*, 2006, 2007; Roberts and Renwick, 2008; Wheeler *et al.*, 2008).

In summary, due to the common molecular structure for steviol glycosides, consisting of a steviol backbone conjugated to different numbers and types of sugar moieties, all individual steviol glycosides share a common metabolic fate, as described above. Therefore, the safety database that has been established for individual steviol glycosides (*e.g.*, stevioside, Rebaudioside A, Rebaudioside D) can be extrapolated to support the safe use of purified steviol glycosides in general, regardless of the steviol glycoside distribution of the preparation, including high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) produced by enzymatic conversion of steviol glycosides.

6.2 Summary of Safety Evaluations on Steviol Glycosides by Scientific and Regulatory Authorities/Bodies

The safety of steviol glycosides was reviewed by JECFA at their 51st, 63rd, 68th, 69th, 82nd, 86th and 87th meetings in 1998, 2004, 2007, 2008, and 2016, 2018 and 2019 respectively. In addition, the safety of steviol glycosides has been reviewed by FSANZ, the European Commission's SCF, the EFSA, and Health Canada (SCF, 1985, 1999; FSANZ, 2008; EFSA, 2010, 2015; Health Canada, 2012). These scientific bodies and regulatory agencies have unanimously concluded that consumption of steviol glycosides is not a safety concern and have established an ADI of 0 to 4 mg/kg body weight, expressed as steviol equivalents. Subsequent to these evaluations, EFSA concluded that "*extending the current specifications to include [two additional steviol glycosides], rebaudiosides D and M, as alternatives to Reb A in the predominant components of steviol glycosides would not be of safety concern*" (EFSA, 2015), while EFSA, JECFA, FSANZ, and Health Canada recently expanded the definition of steviol glycosides to include all individual steviol glycosides present in the *S. rebaudiana* Bertoni leaf (EFSA, 2020; FSANZ, 2017; Health Canada, 2017; JECFA, 2017a,b). In addition to these safety evaluations, the U.S. FDA has reviewed the safety of 57 different steviol glycoside preparations and has consistently raised no objections regarding the GRAS status of steviol glycosides. The FAO JECFA Monographs 23 defines enzymatic conversion as a process in which steviol glycosides that have been extracted from the leaves of *Stevia rebaudiana* Bertoni undergo enzymatic conversion of major steviol glycosides to minor ones. Annex 3 JECFA (2019) monograph 23 defines Enzyme-modified steviol glycosides as a mixture of compounds containing a steviol backbone conjugated to any number or combination of the principal sugar moieties (glucose, rhamnose, xylose, fructose, arabinose, galactose and deoxyglucose) in any of the orientations occurring in the leaves of *Stevia rebaudiana* Bertoni. The product is obtained from the enzymatic treatment of purified steviol glycosides extracted from the leaves of *Stevia rebaudiana* Bertoni. The purified leaf extract is treated with enzymes produced by non-toxicogenic nonpathogenic strains of *Pichia pastoris* and *Escherichia coli* that have been genetically modified with genes from multiple donor organisms to produce glucosyltransferase (EC 2.4.1.17) and sucrose synthase (EC 2.4.1.13). The resulting material is heated and filtered to denature and remove the enzymes. The raw product is concentrated using resin adsorption/desorption or solid/liquid filtration, followed by purification and preparation of the product of commerce using processes that may include decolonization, crystallization, and spray drying. This manufacturing technique maximizes the production of specific steviol glycosides that are not naturally present in high concentrations in the leaf extract, primarily rebaudioside M and rebaudioside D with minor amounts of other steviol glycosides. (87th JECFA (2019) and published in FAO Monographs 23 (2019)).

In these evaluations, the safety data and information that were reviewed by these scientific bodies and regulatory agencies were generally available in the published scientific literature. In a 2-year study in rats, a no-observed-adverse-effect level (NOAEL) of 970 mg/kg body weight/day, equivalent to 383 mg/kg body weight/day as steviol, was determined (Toyoda *et al.*, 1997). The results of the study by Toyoda *et al.* (1997)

was the basis for the established ADI of 0 to 4 mg/kg body weight, expressed as steviol equivalents, for steviol glycosides following application of a safety factor of 100 (JECFA, 2006; FSANZ, 2008; EFSA, 2010; Health Canada, 2012).

6.3 Additional Safety Data for Steviol Glycosides

The safety of steviol glycosides has been extensively reviewed in a number of GRAS notifications submitted to the U.S. FDA, as outlined above, which are incorporated by reference in this notice. The safety of steviol glycosides was most recently evaluated by the U.S. FDA in its evaluation of GRN 882 for purified steviol glycosides, which included a comprehensive search of the scientific literature to capture publications relevant to the safety of steviol glycosides up to March 2019. In order to identify new data related to the safety of steviol glycosides following the U.S. FDA review of GRN 882, a comprehensive search of the scientific literature was conducted from March 2019 to November 2020. The search was limited to articles with full texts within peer-reviewed scientific journals. The literature search was completed using ProQuest and included searches of the following databases for pertinent literature on the safety of steviol glycosides: BIOSIS® Toxicology, BIOSIS Previews®, CAB ABSTRACTS, Embase®, Lancet Titles, MEDLINE®, New England Journal of Medicine, NTIS: National Technical Information Service, Registry of Toxic Effects of Chemical Substances (RTECS®), ToxFile®, TOXLINE. A number of studies reporting results from non-standardized toxicological tests with steviol glycosides for assessing human safety were identified in the literature search. Upon close inspection of the study titles it was determined that a recent study conducted by Purkayastha et al (2020) provided additional information regarding metabolism in children and so was included within the notification, the remainder were considered to be of limited relevance with respect to the overall safety assessment of steviol glycosides as determined by International regulatory Authorities, including the U.S. FDA. Overall, the additional studies identified in the updated search of the scientific literature did not call into question the safety of steviol glycosides. (see Appendix G)

6.4 Safety of the Production Strains

The production strains are derived from *E. coli* K-12. The genome of *E. coli* K-12 has been sequenced and confirms the absence of antibiotic resistance genes and other sequences of concern (Blattner *et al.*, 1997; Hayashi *et al.*, 2006; NCBI, 2018). Furthermore, the *E. coli* parental strain is a member of the well-defined family *Enterobacteriaceae*. All overexpressed genes originated from biosafety level 1 organisms that are not associated with any known allergens or toxins. The manufacturing process includes steps for sterilization and purification procedures to remove DNA and proteins. Analysis of 3 non-consecutive batches of high purity rebaudioside M (≥95% rebaudioside M) demonstrates the successful removal of DNA, and any proteins from the final product. JECFA monograph 23 has also indicated that *E. coli* is an acceptable microorganism to produce the enzymes used in the enzymatic conversion process.

6.4.1 History of Use and the Production Strain

E. coli K-12 has been in use as a laboratory organism for over 50 years and it constitutes one of the most extensively characterized microorganisms (Bachmann, 1972; Jensen, 1993). Along with its use in laboratory research, *E. coli* K-12 has a long history of safe use in the food and pharmaceutical industries. Chymosin, a food enzyme preparation used in the production of cheese, derived from a genetically modified *E. coli* K-12 strain was affirmed as GRAS by the U.S. FDA in 1990 (Flamm, 1991; Olempska-Beer *et al.*, 2006). In addition,

FDA review of GRN 745 concerning the production of Rebaudioside M using a strain of *E. coli* K-12 received “no questions” from the Agency regarding its GRAS status for use in the production of Steviol Glycosides with a high Rebaudioside M content. (U.S. FDA, 2018).

6.4.2 Pathogenicity/Toxicogenicity of the Parental Strain

As discussed in GRN 745, *E. coli* K-12 is not considered a human or animal pathogen and has been classified as Biosafety Level 1 according to the NIH Guidelines (NIH, 2016). *E. coli* K-12 is often used as a reference organism when investigating the virulence factors of pathogenic *E. coli* strains as it is non-pathogenic (Blanc-Potard *et al.*, 2002; Kaper *et al.*, 2004). This species and its derivatives are unable to colonize the mammalian gastrointestinal tract, and do not produce toxins such as Shiga toxin, and are unable to persist in the soil and water (Bogosian *et al.*, 1996; U.S. EPA, 1997). As previously described, the parental strain does not carry any introduced antibiotic resistance genes and the complete genome of this strain has been sequenced, confirming the absence of any toxigenic potential (Blattner *et al.*, 1997; Hayashi *et al.*, 2006).

6.4.3 Potential Allergenicity of the Enzymes

The potential allergenicity of all non-native enzymes was investigated using an *in silico* approach. A sequence homology search was conducted according to the approach outlined by the Food and Agriculture Organization of the United Nations (FAO)/World Health Organization (WHO) (2001) and the WHO/FAO (2009) using the Allergen Online Database Version 20 (available at <http://www.allergenonline.org>; updated February 2020) maintained by the Food Allergy Research and Resource Program (FARRP) of the University of Nebraska (FARRP, 2020). This was done to confirm that the enzymes do not contain amino acid sequences similar to other known allergens that might produce an allergic response. The database contains a comprehensive list of putative allergenic proteins developed via a peer reviewed process for the purpose of evaluating food safety.

No matches were identified from searching with the full amino acid sequence for each enzyme. According to the FARRP guidelines, an identity threshold of greater than 50% or an E-score lower than 1×10^{-7} suggest cross-reactivity with the known allergen to be a possibility.

A second homology search was conducted according to the approach outlined by the FAO/WHO (2001) and the WHO/FAO (2009). In accordance with this guideline, the Allergen Online database was searched using a sliding window of 80-amino acid sequences (segments 1-80, 2-81, 3-82, etc.) derived from the full-length amino acid sequence for each enzyme. The 80-amino acid alignment search was conducted using default settings (E value cutoff = 1 and maximum alignments of 20). Significant homology is defined as an identity match of greater than 35% (Codex Alimentarius, 2009). Using this search strategy, again no matches were identified and the level of protein in the final product NutraSweet M™ is reported as below the limit of detection.

6.5 GRAS Panel Evaluation

Manus Bio has concluded that high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) produced by enzymatic conversion of steviol glycosides, meeting appropriate food-grade specifications and manufactured consistent with cGMP, is GRAS for use as an ingredient in various food products, as described in Part 1.3, on the basis of scientific procedures. Manus Bio's high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) produced by enzymatic conversion of steviol glycosides is substantially equivalent to other steviol glycoside products currently on the U.S. market, including those produced by enzymatic conversion using *E. coli* K-12

as the host micro-organism for producing the enzymes used in the enzymatic conversion process as well as those extracted from the leaves of *S. rebaudiana*.

The GRAS status of high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) produced by enzymatic conversion of steviol glycosides is based on conclusions of scientific bodies and regulatory authorities regarding steviol glycoside safety, data generally available in the public domain pertaining to the safety of steviol glycosides, and a unanimous opinion among a panel of experts ('the GRAS Panel'), who are qualified by scientific training and experience to evaluate the safety of food ingredients. The GRAS Panel consisted of the following qualified scientific experts: Ashley Roberts Ph.D. President AR Toxicology Inc., Jose -Avalos, Ph.D. Princeton University Princeton, NJ., and Stanley M. Tarka, Jr., Ph.D. (The Tarka Group Inc., and The Pennsylvania State University, College of Medicine).

The GRAS panel, convened by Manus Bio, independently and critically evaluated all data and information presented herein, and concluded that high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) produced by enzymatic conversion of steviol glycosides is GRAS for use as a general purpose sweetener, as described in Section 1.3, based on scientific procedures. A summary of data and information reviewed by the GRAS Panel and evaluation of such data as it pertains to the proposed GRAS uses of high-purity rebaudioside M ($\geq 95\%$ rebaudioside M), are presented in Appendix A.

6.6 Conclusions

Based on the data and information presented herein, Manus Bio has concluded high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) produced by enzymatic conversion of steviol glycosides, meeting appropriate food-grade specifications, and manufactured according to cGMP, is safe for use as a general purpose sweetener as presented in Section 1.3. Manus Bio also has further concluded that pivotal data and information relevant to the safety of high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) produced by enzymatic conversion of steviol glycosides are publicly available and therefore the intended uses of high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) can be concluded to be GRAS on the basis of scientific procedures.

Part 7. §170.255 List of Supporting Data and Information

- Aze Y, Toyoda K, Imaida K, Hayashi S, Imazawa T, Hayashi Y et al. (1991). Subchronic oral toxicity study of stevioside in F344 rats. *Eisei Shikenjo Hokoku [Bull Natl Inst Hyg Sci]* (109):48-54 [Japanese].
- Bachmann BJ (1972). Pedigrees of some mutant strains of *Escherichia coli* K-12. *Bacteriol Rev* 36(4):525-557.
- Blanc-Potard A-B, Tinsley C, Scaletsky I, Le Bouguenec C, Guignot J, Servin AL, et al. (2002). Representational difference analysis between Afa/Dr diffusely adhering *Escherichia coli* and nonpathogenic *E. coli* K-12. *Infect Immun* 70(10):5503-5511. DOI:10.1128/IAI.70.10.5503-5511.2002.
- Blattner FR, Plunkett G III, Bloch CA, Perna NT, Burland V, Riley M, et al. (1997). The complete genome sequence of *Escherichia coli* K-12. *Science* 277(5331):1453-1462 [plus supplementary data]. DOI:10.1126/science.277.5331.1453.
- Blumenthal M (1995). FDA lifts import alert on stevia. Herb can be imported only as dietary supplement; future use as a sweetener is still unclear. *HerbalGram* (35 - Fall):17-18.
- Bogosian G, Sammons LE, Morris PJJ, O'Neil JP, Heitkamp MA, Weber DB (1996). Death of the *Escherichia coli* K-12 strain W3110 in soil and water. *Appl Environ Microbiol* 62(11):4114-4120.
- Chang SS, Cook JM (1983). Stability studies of stevioside and rebaudioside A in carbonated beverages. *J Agric Food Chem* 31(2):409-412. DOI:10.1021/jf00116a056.
- Curry LL, Roberts A (2008). Subchronic toxicity of rebaudioside A. *Food Chem Toxicol* 46(Suppl. 7):S11-S20.
- Curry LL, Roberts A, Brown N (2008). Rebaudioside A: two-generation reproductive toxicity study in rats. *Food Chem Toxicol* 46(Suppl. 7):S21-S30.
- Datsenko, KA, BL Wanner 2000. One-step inactivation of chromosomal genes in *Escherichia coli* K-12 using PCR products. *Proc. Natl. Acad. Sci. U.S.A.* 97(12):6640-5.
- EFSA (2010). EFSA Panel on Food Additives and Nutrient Sources scientific opinion on safety of steviol glycosides for the proposed uses as a food additive. (Question number: EFSA-Q-2007-071; EFSA-Q-2008-387; EFSA-Q-2008-401, adopted on 10 March 2010 by European Food Safety Authority). *EFSA J* 78(4):1537. [85 pp]. DOI:10.2903/j.efsa.2010.1537. Available at: <https://www.efsa.europa.eu/en/scdocs/scdoc/1537.htm>.
- EFSA (2015). Scientific opinion on the safety of the proposed amendment of the specifications for steviol glycosides (E 960) as a food additive. (EFSA Panel on Food Additives and Nutrient Sources Added to Food/ANS) (Question no EFSA-Q-2014-00002, adopted on 17 November 2015 by European Food Safety Authority). *EFSA J* 13(12):4316 [29 pp.]. DOI: 10.2903/j.efsa.2015.4316. Available at: <https://www.efsa.europa.eu/en/efsajournal/pub/4316>.
- EFSA (2020). Safety of a proposed amendment of the specifications for steviol glycosides (E 960) as a food additive: to expand the list of steviol glycosides to all those identified in the leaves of *Stevia Rebaudiana* Bertoni. (EFSA Panel on Food Additives and Flavourings (FAF)) (EFSA-Q-2019-00063, adopted on 24 March 2020 by European Food Safety Authority). Available at: <https://doi.org/10.2903/j.efsa.2020.6106>.

- FAO/WHO (2001). *Evaluation of Allergenicity of Genetically Modified Foods*. Report of a Joint FAO/WHO Expert Consultation on Allergenicity of Foods Derived from Biotechnology 22-25 January 2001. Rome, Italy: Food and Agriculture Organization of the United Nations (FAO), World Health Organization (WHO). Available at: <http://www.who.int/foodsafety/publications/gmo-allergenicity/en/>.
- FAO (2016). *Steviol Glycosides: Chemical and Technical Assessment*. (82nd JECFA - Chemical and Technical Assessment/CTA). Rome, Italy: Food and Agriculture Organization of the United Nations (FAO). Available at: <http://www.fao.org/3/a-br566e.pdf> [Last accessed Oct. 4, 2017].
- FARRP (2018). *AllergenOnline Version 18A: Home of the FARRP Allergen Protein Database*. Lincoln (NE): University of Nebraska-Lincoln, Food Allergy Research and Resource Program (FARRP). Available at: <http://www.allergenonline.org/> [Released: February 1, 2018].
- FCC (2016). Steviol glycosides. In: *Food Chemicals Codex, 10th edition*. Rockville (MD): United States Pharmacopeial Convention (USP), pp. 1234-1238.
- Ferlow K (2005). Stevia - The sweetest substance on Earth. *NutraCos* 4(2, Suppl.):10-11. Flamm
- EL (1991). How FDA approved chymosin: a case history. *Bio/Technology* 9 (4):349-351.
- FSANZ (2008). *Final Assessment Report: Application A540 Steviol Glycosides as Intense Sweeteners*. Canberra, Australia: Food Standards Australia New Zealand (FSANZ). Available at: http://www.foodstandards.gov.au/code/applications/documents/FAR_A540_Steviol_glycosides.pdf
- FSANZ (2017). Schedule 3: Identity and purity. In: *Australia New Zealand Food Standards Code*. Canberra, Australia / Wellington, New Zealand: Food Standards Australia New Zealand (FSANZ). Available online: <https://www.legislation.gov.au/Details/F2017C00542>.
- Gardana C, Simonetti P, Canzi E, Zanchi R, Pietta P (2003). Metabolism of stevioside and rebaudioside A from *Stevia rebaudiana* extracts by human microflora. *J Agric Food Chem* 51(22):6618-6622. DOI:10.1021/jf0303619.
- Geuns JMC (2003). Stevioside. *Phytochemistry* 64(5):913-921. DOI:10.1016/S0031-9422(03)00426-6.
- Geuns JMC, Pietta, P (2004) [unpublished]. Stevioside metabolism by human volunteers. Report from Segrate (MI), Italy: Laboratory Functional Biology, Kuleuven, Leuven Belgium and ITB-CNR. Submitted to WHO by Belgium: Federal Ministry of Social Affairs, Public Health and the Environment. Cited In: JECFA, 2006.
- Geuns JM, Augustijns P, Mols R, Buyse JG, Driessen B (2003). Metabolism of stevioside in pigs and intestinal absorption characteristics of stevioside, rebaudioside A and steviol. *Food Chem Toxicol* 41(11):1599-1607. DOI:10.1016/S0278-6915(03)00191-1.
- Geuns JMC, Buyse J, Vankeirsbilck A, Temme EHM, Compennolle F, Toppet S (2006). Identification of steviol glucuronide in human urine. *J Agric Food Chem* 54(7):2794-2798. DOI:10.1021/jf052693e.
- Geuns JMC, Buyse J, Vankeirsbilck A, Temme EHM (2007). Metabolism of stevioside by healthy subjects. *Exp Biol Med* 232(1):164-173.
- Hayashi K, Morooka N, Yamamoto Y, Fujita K, Isono K, Choi S, et al. (2006). Highly accurate genome sequences

of *Escherichia coli* K-12 strains MG1655 and W3110. *Mol Syst Biol* 2(1):22006.0007 [5pp].
DOI:10.1038/msb4100049.

Health Canada (2012). *Notice of Modification to the Lists of Permitted Food Additives to Enable the Use of Steviol Glycosides as a Table-Top Sweetener and as a Sweetener in Certain Food Categories*. Ottawa (ON): Health Canada, Bureau of Chemical Safety. Available at: <http://www.hc-sc.gc.ca/fn-an/consult/nom-adm-0002/index-eng.php> [Date Modified: 2012-11-30]

Health Canada (2017). *Notice of Modification to the List of Permitted Sweeteners to Enable the Use of Steviol Glycosides from *Stevia rebaudiana* Bertoni as a Sweetener*. (Reference Number: NOM/ADM-0102). Ottawa (ON): Health Canada, Bureau of Chemical Safety, Food Directorate, Health Products and Food Branch. Available at: <https://www.canada.ca/en/health-canada/services/food-nutrition/legislation-guidelines/acts-regulations/modification-list-permitted-sweeteners-steviol-glycosides.html> [Date Modified: 2017-08-29].

Hutapea AM, Toskulkao C, Buddhasukh D, Wilairat P, Glinsukon T (1997). Digestion of stevioside, a natural sweetener, by various digestive enzymes. *J Clin Biochem Nutr* 23(3):177-186.
DOI:10.3164/jcbs.23.177.

International Sugar Organization (2001). Developments in the high-intensity sweeteners markets. *Zuckerindustrie* 126(12):970.

JECFA (2006). Steviol glycosides. In: *Safety Evaluation of Certain Food Additives*. Sixty-third Meeting of the Joint FAO/WHO Expert Committee on Food Additives, June 8-17, 2004, Geneva, Switz. (WHO Food Additives Series, no 54). Geneva, Switz.: World Health Organization (WHO), International Programme on Chemical Safety (IPCS), pp. 117-144, 638. Available at: http://whqlibdoc.who.int/publications/2006/9241660546_eng.pdf.

JECFA (2007). Steviol glycosides. In: *68th JECFA - Chemical and Technical Assessment (CTA)*. [Sixty-eighth meeting held June 17-26, 2008]. (Prepared by Harriet Wallin and revised by Paul M. Kuznesof Ph.D). Geneva, Switz.: Joint FAO/WHO Expert Committee on Food Additives (JECFA). Available at: http://www.fao.org/fileadmin/templates/agns/pdf/jecfa/cta/68/Steviol_glycosides.pdf.

JECFA (2017a). Steviol glycosides from *Stevia rebaudiana* Bertoni [New specifications prepared at the 84th JECFA, 2017), Superseding tentative specifications prepared at the 82nd JECFA (2016)]. In: *Compendium of Food Additive Specifications*. 84th Meeting, Rome, 6-15 June 2017 (FAO JECFA Monographs 20). Rome, Italy: Food and Agriculture Organization of the United Nations (FAO) / Geneva, Switz.: World Health Organization (WHO), pp. 50-69. Available at: <http://www.fao.org/documents/card/en/c/4b06cdda-3e70-4c80-b7e5-56034601836b/>.

JECFA (2017b). Stevia glycosides (addendum). In: *Safety Evaluation of Certain Food Additives*. Eighty-second Meeting of the Joint FAO/WHO Expert Committee on Food Additives (JECFA), June 7-16, 2016. (WHO Food Additives Series, no 73). Geneva, Switz.: World Health Organization / Rome, Italy: Food and Agriculture Organization of the United Nations (FAO), pp. 181-218, 492. Available at: <http://apps.who.int/iris/bitstream/10665/258934/1/9789241660730-eng.pdf?ua=1>.

JECFA (2019). Stevia glycosides. In: *Joint FAO/WHO Expert Committee on Food Additives 87th Meeting: FAO JECFA Monographs 23 (framework for) Steviol Glycosides, June 4-13th, 2019 Rome Italy*: Food and Agriculture Organization of the United Nations (FAO) / Geneva, Switz.: World Health Organization (WHO). Available at: <http://www.fao.org/documents/card/en/c/ca7513en>.

- Jensen KF (1993). The *Escherichia coli* K-12 "wild types" W3110 and MG1655 have an *rph* frameshift mutation that leads to pyrimidine starvation due to low *pyrE* expression levels. *J Bacteriol* 175(11):3401-3407.
- Kaper JB, Nataro JP, Mobley HLT (2004). Pathogenic *Escherichia coli*. *Nat Rev Microbiol* 2(2):123-140. DOI:10.1038/nrmicro818.
- Koyama E, Sakai N, Ohori Y, Kitazawa K, Izawa O, Kakegawa K, et al. (2003a). Absorption and metabolism of glycosidic sweeteners of stevia mixture and their aglycone, steviol, in rats and humans. *Food Chem Toxicol* 41(6):875-883. DOI:10.1016/S0278-6915(03)00039-5.
- Koyama E, Kitazawa K, Ohori Y, Izawa O, Kakegawa K, Fujino A, et al. (2003b). In vitro metabolism of the glycosidic sweeteners, stevia mixture and enzymatically modified stevia in human intestinal microflora. *Food Chem Toxicol* 41(3):359-374. DOI:10.1016/S0278-6915(02)00235-1.
- Kraemer T, Maurer HH (1994). On the metabolism of the sweetener stevioside in humans. *Eur J Pharm Sci* 2(1/2):103 [abstract FC12]. DOI:10.1016/0928-0987(94)90121-X.
- Kroyer GT (1999). The low calorie sweetener stevioside: stability and interaction with food ingredients. *Lebensm Wiss Technol* 32(8):509-512.
- Nakayama K, Kasahara D, Yamamoto F (1986). Absorption, distribution, metabolism and excretion of stevioside in rats. *Shokuhin Eiseigaku Zasshi* 27(1):1-8. DOI:10.3358/shokueishi.27.1.
- NCBI (2018). *Escherichia coli* str. K-12 substr. W3110 DNA, complete genome (GenBank: AP009048.1). In: *NCBI Nucleotide Database*. Bethesda (MD): National Center for Biotechnology Information (NCBI), U.S. National Library of Medicine (NLM). Available at: <https://www.ncbi.nlm.nih.gov/nuccore/AP009048.1> [Last accessed: March 14, 2018].
- NIH (2016). *NIH Guidelines for Research Involving Recombinant or Synthetic Nucleic Acid Molecules*. (NIH Guidelines). Bethesda (MD): National Institutes of Health (NIH), Office of Science Policy. Available at: https://osp.od.nih.gov/wp-content/uploads/2013/06/NIH_Guidelines.pdf [Apr. 2016].
- Nikiforov AI, Eapen AK (2008). A 90-day oral (dietary) toxicity study of rebaudioside A in Sprague-Dawley rats. *Int J Toxicol* 27(1):65-80.
- Nikiforov AI, Rihner MO, Eapen AK, Thomas JA (2013). Metabolism and toxicity studies supporting the safety of rebaudioside D. *Int J Toxicol* 32(4):261-273. DOI:10.1177/1091581813492828.
- Oehme A, Wüst M, Wölwer-Rieck U (2017). Steviol glycosides are not altered during commercial extraction and purification processes. *Int J Food Sci Technol* 52(10):2156-2162. DOI:10.1111/ijfs.13494.
- Olempska-Beer ZS, Merker RI, Ditto MD, Dinovi MJ (2006). Food-processing enzymes from recombinant microorganisms-a review. *Regul Toxicol Pharmacol* 45(2):144-158. DOI:10.1016/j.yrtph.2006.05.001.
- Pariza MW, Johnson EA (2001). Evaluating the safety of microbial enzyme preparations used in food processing: update for a new century. *Regul Toxicol Pharmacol* 33(2):173-186. DOI:10.1006/rtph.2001.1466.
- Purkayastha S, Pugh G, Lynch B, Roberts A, Kwok D, Tarka SM (2014). *In vitro* metabolism of rebaudioside B, D, and M under anaerobic conditions: comparison with rebaudioside A. *Regul Toxicol Pharmacol*

68(2):259-268. DOI:10.1016/j.yrtph.2013.12.004.

Purkayastha S, Bhusari S, Pugh G, Jr., Teng X, Kwok D, Tarka SM (2015). *In vitro* metabolism of rebaudioside E under anaerobic conditions: comparison with rebaudioside A. *Regul Toxicol Pharmacol* 72(3):646- 657. DOI:10.1016/j.yrtph.2015.05.019.

Purkayastha S, Markosayan A, Prakash I, Bhusari S, Pugh G, Lynch B, et al. (2016). Steviol glycosides in purified stevia leaf extract sharing the same metabolic fate. *Regul Toxicol Pharmacol* 77:125-133. DOI:10.1016/j.yrtph.2016.02.015.

Purkayastha S, Kwok D (2020). Metabolic fate in adult and pediatric population of steviol glycosides produced from stevia leaf extract by different production technologies. *Regul Toxicol Pharmacol* 116 104727. <https://doi.org/10.1016/j.yrtph.2020.104727>.

Renwick AG (2008). The use of a sweetener substitution method to predict dietary exposures for the intense sweetener rebaudioside A. *Food Chem Toxicol* 46(Suppl. 7):S61-S69. DOI:10.1016/j.fct.2008.05.009.

Renwick AG, Tarka SM (2008). Microbial hydrolysis of steviol glycosides. *Food Chem Toxicol* 46(Suppl. 7):S70-S74. DOI:10.1016/j.fct.2008.05.008.

Roberts A, Renwick AG (2008). Comparative toxicokinetics and metabolism of rebaudioside A, stevioside, and steviol in rats. *Food Chem Toxicol* 46(Suppl. 7):S31-S39. DOI:10.1016/j.fct.2008.05.006.

SCF (1985). Reports of the Scientific Committee for Food concerning sweeteners (Opinion expressed by the SCF on 14 September 1984) [Stevioside]. In: *Food Science and Techniques*. (Reports of the Scientific Committee for Food, 16th series). Brussels, Belgium: Commission of the European Communities (EEC), Scientific Committee for Food (SCF), 2, 4, 7, 15, 20. Available at: https://ec.europa.eu/food/sites/food/files/safety/docs/sci-com_scf_reports_16.pdf.

SCF (1999). *Opinion on Stevioside as a Sweetener (Adopted on 17 June 1999)*. (CS/ADD/EDUL/167 Final). Brussels, Belgium: European Commission, Health & Consumer Protection Directorate-General, Scientific Committee on Food (SCF). Available at: https://ec.europa.eu/food/sites/food/files/safety/docs/sci-com_scf_out34_en.pdf.

Simonetti P, Gardana C, Bramati L, Pietta PG (2004). Bioavailability of stevioside from *Stevia rebaudiana* in humans: preliminary report. In: Geuns JMC, Buyse J, editors. *Safety of Stevioside: Proceedings of the First Symposium Sponsored by KULeuven, April 16, 2004, Leuven, Belgium*. Heverlee, Belgium: Euprint ed., pp. 51-62.

Sung LH (2002) [unpublished]. Report on pharmacokinetic (PK) studies of T100 sunstevia 95% stevioside in rats. Report from Singapore: Sunlabel Pte Ltd. Submitted to WHO by the Ministry of Health and Welfare, Japan. Cited In: JECFA, 2006.

Toyoda K, Matsui H, Shoda T, Uneyama C, Takada K, Takahashi M (1997). Assessment of the carcinogenicity of stevioside in F344 rats. *Food Chem Toxicol* 35(6):597-603. DOI:10.1016/S0278-6915(97)00023-9.

U.S. EPA (1997). *Escherichia Coli K-12 Final Risk Assessment: Attachment I--Final Risk Assessment of Escherichia Coli K-12 Derivatives*. Washington (DC): U.S. Environmental Protection Agency (U.S. EPA), Biotechnology Program under the Toxic Substances Control Act (TSCA). Available at: <http://www2.epa.gov/sites/production/files/2015-09/documents/fra004.pdf> [Last updated on September 27, 2012].

- U.S. FDA (2008). *Generally Recognized as Safe (GRAS) Notification for the Use of Stevia Rebaudiana (Bertoni) Bertoni*. (Submitted as U.S. FDA, 2009 - GRN 275). Prepared by Bethesda (MD): Spherix Incorporated for Fort Washington (PA): McNeil Nutritionals, LLC for submission to College Park (MD): U.S. Food and Drug Administration (U.S. FDA), Center for Food Safety and Applied Nutrition (CFSAN), Office of Food Additive Safety. Available at: <http://www.accessdata.fda.gov/scripts/fdcc/index.cfm?set=GRASNotices&id=275>.
- U.S. FDA (2016). *Agency Response Letter GRAS Notice No. GRN 000624 [D-allulose 3-epimerase from Arthrobacter globiformis M30 produced in Escherichia coli, Itami City, Hyogo, Japan: Matsutani Chemical Industry Co. Ltd.]*. College Park (MD): U.S. Food and Drug Administration (U.S. FDA), Center for Food Safety & Applied Nutrition (CFSAN), Office of Food Additive Safety. Available at: <http://www.accessdata.fda.gov/scripts/fdcc/index.cfm?set=GRASNotices&id=624> [Aug. 18, 2016].
- U.S. FDA (2017). *Agency Response Letter GRAS Notice No. GRN 000667 [Rebaudioside M, Rancho Santa Margarita (CA): Blue California]*. College Park (MD): U.S. Food and Drug Administration (U.S. FDA), Center for Food Safety & Applied Nutrition (CFSAN), Office of Food Additive Safety. Available at: <http://www.accessdata.fda.gov/scripts/fdcc/index.cfm?set=GRASNotices&id=667> [Feb. 17, 2017 - FDA response - no questions].
- U.S. FDA (2018). *Agency Response Letter GRAS Notice No. GRN 000745 [Rebaudioside M, 915 Harger Road, Suite 250 Oak Brook, Illinois PureCircle Limited]*. College Park (MD): U.S. Food and Drug Administration (U.S. FDA), Center for Food Safety & Applied Nutrition (CFSAN), Office of Food Additive Safety. Available at: https://www.cfsanappsexternal.fda.gov/scripts/fdcc/?set=GRASNotices&id=745&sort=GRN_No&order=D ESC&startrow=1&type=basic&search=745 [Apr. 20, 2018 - FDA response - no questions].
- Wang LZ, Goh BC, Fan L, Lee HS (2004). Sensitive high-performance liquid chromatography/mass spectrometry method for determination of steviol in rat plasma. *Rapid Commun Mass Spectrom* 18(1):83-86. DOI:10.1002/rcm.1285.
- Wheeler A, Boileau AC, Winkler PC, Compton JC, Prakash I, Jiang X, et al. (2008). Pharmacokinetics of rebaudioside A and stevioside after single oral doses in healthy men. *Food Chem Toxicol* 46(Suppl. 7):S54-S60. DOI:10.1016/j.fct.2008.04.041.
- WHO/FAO (2009). *Foods Derived from Modern Biotechnology, 2nd edition*. Geneva, Switz.: World Health Organization (WHO) / Rome, Italy: Food and Agriculture Organization of the United Nations (FAO). Available at: <http://www.fao.org/docrep/011/a1554e/a1554e00.htm>.
- Williams LD, Burdock GA (2009). Genotoxicity studies on a high-purity rebaudioside A preparation. *Food Chem Toxicol* 47(8):1831-1836. DOI:10.1016/j.fct.2009.04.046.
- Wingard RE, Brown JP, Enderlin FE, Dale JA, Hale RL, Seitz CT (1980). Intestinal degradation and absorption of the glycosidic sweeteners stevioside and rebaudioside A. *Experientia* 36(5):519-520. DOI:10.1007/BF01965774.

APPENDIX A

GRAS Panel Consensus Report

APPENDIX A:

GRAS Panel Report Concerning the Generally Recognized as Safe (GRAS) Status of high-purity Rebaudioside M ($\geq 95\%$ Rebaudioside M) produced by enzymatic conversion of stevia leaf extract.

November 10, 2020

INTRODUCTION

Manus Bio intends to market high purity rebaudioside M ($\geq 95\%$ rebaudioside M) and identified as NutraSweet M™, and produced via enzymatic conversion for use as a general purpose sweetener in the United States (U.S.). Steviol glycosides have historically been obtained by hot water extraction from the leaves of *Stevia rebaudiana* Bertoni and solvent purification. More than 40 different steviol glycosides have been identified in the leaf extracts to date. Manus Bio has developed an alternative manufacturing process to produce rebaudioside M. Manus Bio's high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) is produced through enzymatic conversion of steviol glycosides ($\geq 90\%$ total steviol glycosides) extracted from *S. rebaudiana* Bertoni using UDP- glucosyltransferase enzymes derived from genetically modified *Escherichia coli* strains derived from *E. coli* K-12. High-purity rebaudioside M is comprised of $\geq 95\%$ rebaudioside M and $\geq 95\%$ total steviol glycosides, which meets or exceeds the $\geq 95\%$ steviol glycoside purity criteria established by the Joint FAO/WHO Expert Committee on Food Additives (JECFA) and the Food Chemicals Codex.

At the request of Manus Bio, an Expert Panel of independent scientists (the GRAS Panel), qualified by their relevant national and international experience and scientific training to evaluate the safety of food ingredients, was specially convened to conduct a critical and comprehensive evaluation of the available pertinent data and information, and to determine whether, under the conditions of intended use as a sweetening agent, high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) would be Generally Recognized as Safe (GRAS), based on scientific procedures. The GRAS Panel consisted of the below-signed qualified scientific experts: Ashley Roberts Ph.D., President AR Toxicology Inc.; Jose -Avalos, Ph.D. Princeton University, Princeton, NJ., and Stanley M. Tarka Jr., Ph.D. (The Tarka Group Inc., and The Pennsylvania State University, College of Medicine). For purposes of the GRAS Panel's evaluation, "safe" or "safety" means there is a reasonable certainty in the minds of competent scientists that the substance is not harmful under the intended conditions of use, as defined by the U.S. Food and Drug Administration (FDA) in 21 CFR 170.3(i) (U.S. FDA, 2017).

The GRAS Panel independently and collectively evaluated a dossier titled "*Documentation Supporting NutraSweet M as Generally Recognized as Safe (GRAS) for Use as a General Purpose Sweetener*", which included a comprehensive summary of scientific information on high-purity rebaudioside M ($\geq 95\%$ rebaudioside M). This dossier was prepared with information available in the public domain and also included details pertaining to the manufacturing method, product specifications and supporting batch analyses, intended uses and use-levels in food and beverages, consumption estimates for all intended uses, and a summary of the scientific literature pertaining to the safety of steviol glycosides. The GRAS Panel also evaluated other information deemed appropriate or necessary.

Following their independent and critical evaluation of such data and information, the GRAS Panel convened on November 25th 2020 *via* teleconference and unanimously concluded that the intended use described herein for high-purity rebaudioside M ($\geq 95\%$ rebaudioside M), meeting appropriate food-grade specifications as described in the supporting dossier and manufactured according to current Good Manufacturing Practice (cGMP), is safe, suitable, and GRAS based on scientific procedures. A summary of the basis of the GRAS Panel's conclusion is presented below.

CHEMISTRY AND MANUFACTURING

The subject of this GRAS evaluation is high-purity rebaudioside M ($\geq 95\%$ rebaudioside M), which is a highly purified product comprised of $\geq 95\%$ rebaudioside M and $\geq 95\%$ total steviol glycosides, consistent with the purity criteria for steviol glycosides as established by JECFA (JECFA, 2017a,b). The remaining 5% of high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) includes additional steviol glycosides and water. Due to the common steviol backbone, all steviol glycosides share a common metabolic fate in which they are hydrolyzed to steviol in the lower gastrointestinal tract, which is then absorbed into the body, conjugated with glucuronic acid, and eliminated through the urine in humans.

High-purity rebaudioside M ($\geq 95\%$ rebaudioside M) is manufactured using raw materials, processing aids, and equipment that are food-grade ingredients¹, permitted by U.S. regulation, have GRAS status, or have been self-affirmed as safe for use in food for their respective uses. The high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) product is manufactured by an enzymatic conversion process using MG1655 derived from modified *E. coli* strains derived from *E. coli* K-12. High-purity rebaudioside M ($\geq 95\%$ rebaudioside M) is manufactured in a multi-stage process; in the first step steviol glycosides are extracted from the stevia leaf by a series of crushing, dissolution, extraction, and precipitation steps that are consistent with the methodology outlined in the CTA for steviol glycosides (FAO, 2016). The steviol glycoside mixture contains $\geq 90\%$ total steviol glycosides. In the second stage, the production strains are introduced to the media containing the glycosides, and nutrients and undergo enzymatic conversion. In the final stage the resulting steviol glycoside mixture is purified in a series of filtration and washing steps to yield a final product containing $\geq 95\%$ rebaudioside M and $\geq 95\%$ total steviol glycosides.

Appropriate food-grade product specifications have been established for high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) based on the steviol glycosides specification established by JECFA (JECFA, 2017a,b). The steviol glycoside content of high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) is measured using the high-performance liquid chromatography (HPLC) method described in the JECFA specification monograph for steviol glycosides from *Stevia rebaudiana* Bertoni (JECFA, 2017a,b). Review by the GRAS Panel of the analytical data for 3 non-consecutive lots of high purity rebaudioside M ($\geq 95\%$ rebaudioside M) confirms that the final product is produced in compliance with the established product specifications. In addition, residual protein in the final product was confirmed to be below the detection level in 3 non-consecutive lots of high purity rebaudioside M ($\geq 95\%$ rebaudioside M) *via* the bicinchoninic acid (BCA) assay. The absence of residual DNA in the final product for these same lots was confirmed to be below the detection limit *via* polymerase chain reaction (PCR).

¹ Compliant with the specifications set forth in the Food Chemicals Codex or equivalent international food or pharmacopeia standard (*e.g.*, JECFA, CODEX, United States Pharmacopeia, European Pharmacopeia).

Absence of residues of commonly used pesticides in the final product was confirmed in Lot 202003051 and Lot 202008081 of the starting material ($\geq 90\%$ total steviol glycosides).

JECFA has concluded that steviol glycosides are thermally and hydrolytically stable for use in foods and acidic beverages under normal processing and storage conditions (JECFA, 2007). The stability study on high-purity Manus Bio rebaudioside M ($\geq 95\%$ rebaudioside M) is currently on-going; however, the 3-month results indicate that the total steviol glycoside and rebaudioside M contents are stable when kept in commercial packaging at 50°C .

INTENDED FOOD USES AND ESTIMATED INTAKE

High-purity rebaudioside M ($\geq 95\%$ rebaudioside M) is intended for use as a general-purpose sweetener that will be added to various food and beverage products that are consistent with the current uses of other high-intensity sweeteners on the U.S. market. The estimated intakes of high purity rebaudioside M ($\geq 95\%$ rebaudioside M) were calculated using a post-market surveillance approach as described by Renwick (2008). The estimated intakes were calculated by adjusting the post-market surveillance data for other high-intensity sweeteners using the sweetness intensity of high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) relative to sucrose (*i.e.*, approximately 250). The results are shown in Table 1. For non-diabetic adults, average and high-end intakes of high purity rebaudioside M ($\geq 95\%$ rebaudioside M) of up to 0.26 and 0.68 mg/kg body weight/day expressed as steviol equivalents, respectively, were calculated. For diabetic adults, average and high-end intakes were slightly higher at up to 0.28 and 0.90 mg/kg body weight/day. Average and high-end exposures to high purity rebaudioside M ($\geq 95\%$ rebaudioside M), expressed as steviol equivalents, in non-diabetic children were calculated to be up to 0.43 and 0.99 mg/kg body weight/day, respectively. Although average intakes of high-purity rebaudioside M ($\geq 95\%$ rebaudioside M), expressed as steviol equivalents were estimated to be higher at up to 0.67 mg/kg body weight/day in diabetic children compared to values for non-diabetic children, high-end values in diabetic children (0.91 mg/kg body weight/day) were lower than high-end values in non-diabetic children. The predicted intakes of high-purity rebaudioside M ($\geq 95\%$ rebaudioside M), expressed as steviol equivalents, are all below the current acceptable daily intake (ADI) defined by the JECFA for steviol glycosides (FAO, 2016) of 0 to 4 mg/kg body weight/day as steviol. Recently, JECFA re-assessed the dietary exposure to steviol glycosides using different intake models, including the approach described by Renwick (2008), and noted that the replacement estimates were highly conservative and that actual exposures to steviol glycosides, expressed as steviol equivalents, would range from 0.4 to 7.2 mg/kg body weight/day (FAO, 2016). JECFA made note that this method overestimates dietary exposure (FAO, 2016).

Table 1 Estimated Consumption High-Purity Rebaudioside M ($\geq 95\%$ Rebaudioside M) Produced by Enzymatic Conversion of Steviol Glycosides Using the Intense Sweetener Intake Assessment Methodology described by Renwick (2008)

Population Group	Intakes of Intense Sweeteners (expressed as sucrose equivalents) (mg/kg bw/day)		Consumption Estimates			
	Average Consumer	High Consumer	High-Purity Rebaudioside M (≥95% Rebaudioside M) ^a (mg/kg bw/day)		High-Purity Rebaudioside M (≥95% Rebaudioside M) ^{a,b} (mg/kg bw/day) as Steviol	
Non-diabetic adults	255	675	1.02	2.7	0.26	0.68
Diabetic adults	280	897	1.12	3.59	0.28	0.9
Non-diabetic children	425	990	1.7	3.96	0.43	0.99
Diabetic children	672	908	2.69	3.63	0.67	0.91

bw = body weight

^a Approximately 250 times as sweet as sucrose.

^b Calculated based on the molecular weights of steviol (318.45 g/mol) and rebaudioside M (1,291.3 g/mol) [steviol conversion factor of 0.25].

INFORMATION TO ESTABLISH SAFETY

The GRAS Panel reviewed the available data supporting the safety of individual steviol glycosides to evaluate the safety of high-purity rebaudioside M (≥95% rebaudioside M). The available data included a discussion on the metabolic fate of steviol glycosides, a summary of the extensive conclusions on the safety of steviol glycosides by global scientific and regulatory authorities/bodies, including the U.S. FDA, and other data that was deemed pivotal in determining the safety of high-purity rebaudioside M (≥95% rebaudioside M) produced by Manus.

Steviol glycosides are not hydrolyzed in the upper gastrointestinal tract due to the presence of β-glycosidic bonds; the unchanged steviol glycosides enter the colon and are subject to microbial degradation by the gut microflora, resulting in the release of the aglycone steviol (Wingard *et al.*, 1980; Hutapea *et al.*, 1997; Gardana *et al.*, 2003; Koyama *et al.*, 2003a,b; Geuns *et al.*, 2003, 2007; Renwick and Tarka, 2008; Nikiforov *et al.*, 2013; Purkayastha *et al.*, 2016). Specifically, Purkayastha *et al.* (2016) conducted incubation studies with pooled human fecal homogenates collected from adult males and females using rebaudiosides A, B, C, D, E, F, M, steviolbioside and dulcoside A, and showed that all glycosides were converted to steviol over a 24–48 h incubation period. These studies demonstrated that the number and location of sugar units attached to the steviol backbone do not significantly affect the microbial hydrolysis. Likewise, this study demonstrated that steviol glycosides, irrespective of the type of sugar moiety (e.g., glucose, rhamnose, xylose) attached to the steviol backbone, were metabolized to steviol at generally similar hydrolysis rates. Incubation tests were also conducted in fecal homogenates collected from adult and pediatric populations with steviol glycoside from stevia leaf extracts and steviol glycosides produced by enzymatic conversion of Reb A to larger molecules by attaching glucose units via β- or α-glycosidic bonds. The steviol glycosides produced by extraction from stevia leaf or enzymatic conversion of stevia leaf extract were also reported to share a similar metabolic fate in the fecal homogenates from adults and children supporting a similar metabolic fate of steviol glycosides in all ages (Purkayastha 2020). Steviol glycosides are hydrolyzed sequentially, in which 1 sugar moiety is removed at a time, with the degradation rates dependent on the structural complexity of each steviol glycoside (Wingard *et al.*, 1980; Koyama *et al.*, 2003b). Despite the differences in chemical structure, however, the rate of hydrolysis of different steviol glycosides to steviol are relatively similar, especially during the first 24 hours of incubation in *in vitro* metabolic studies with human fecal homogenates (Purkayastha *et al.*, 2014, 2015, 2016). Following microbial degradation, the steviol metabolite is absorbed systemically into the portal vein and distributed to the

liver, spleen, adrenal glands, fat, and blood (Nakayama *et al.*, 1986; Sung, 2002 [unpublished]; Koyama *et al.*, 2003b; Wang *et al.*, 2004; Roberts and Renwick, 2008). Steviol is conjugated to glucuronic acid to form steviol glucuronide in the liver. The steviol glucuronide metabolite and any unconjugated steviol or unhydrolyzed fraction of the administered glycosides are excreted primarily in the urine, and, to a lesser extent, feces in humans (Wingard *et al.*, 1980; Nakayama *et al.*, 1986; Kraemer and Maurer, 1994; Sung, 2002 [unpublished]; Geuns and Pietta, 2004 [unpublished]; Simonetti *et al.*, 2004; Geuns *et al.*, 2006, 2007; Roberts and Renwick, 2008; Wheeler *et al.*, 2008). Thus, due to the shared metabolic fate of steviol glycosides, the safety database that has been established for individual steviol glycosides (*e.g.*, stevioside, rebaudioside A, rebaudioside D), can be extrapolated to support the safety of other purified steviol glycosides in general, regardless of the steviol glycoside distribution of the preparation, including high purity rebaudioside M ($\geq 95\%$ rebaudioside M) produced by enzymatic conversion of steviol glycosides.

The safety of steviol glycosides have been extensively reviewed by various scientific and regulatory authorities/bodies, such as JECFA, U.S. FDA, Food Standards Australia New Zealand (FSANZ), the European Commission's Scientific Committee on Food (SCF), the European Food Safety Authority (EFSA), and Health Canada (SCF, 1985, 1999; FSANZ, 2008; EFSA, 2010, 2015; Health Canada, 2012b; JECFA, 2006, 2017a,b). Based on safety data and information that were generally available in the published scientific literature, these scientific bodies and regulatory agencies have unanimously concluded that steviol glycosides is of no safety concern and have established an ADI of 0 to 4 mg/kg body weight, expressed as steviol equivalents, based on the results of a 2-year study in rats (Toyoda *et al.*, 1997). The no-observed-adverse-effect level (NOAEL) of 970 mg/kg body weight/day, equivalent to 383 mg/kg body weight/day as steviol, determined from the results of the study by Toyoda *et al.* (1997) became the basis for the established ADI following application of a safety factor of 100 (JECFA, 2009; FSANZ, 2008; EFSA, 2010; Health Canada, 2012b). Subsequent to these safety evaluations, the EFSA concluded that "*extending the current specifications to include [two additional steviol glycosides], rebaudiosides D and M, as alternatives to Reb A in the predominant components of steviol glycosides would not be of safety concern*". (EFSA, 2015) EFSA, JECFA, FSANZ, and Health Canada recently expanded the definition of steviol glycosides to include all individual steviol glycosides present in the *S. rebaudiana* Bertoni leaf. (EFSA, 2020; FSANZ, 2017b; Health Canada, 2017; JECFA, 2017) The FAO JECFA Monographs 23 defines enzymatic conversion as a process in which steviol glycosides that have been extracted from the leaves of *Stevia rebaudiana* Bertoni undergo enzymatic conversion of major steviol glycosides to minor ones. Annex 3 JECFA (2019) monograph 23 defines Enzyme-modified steviol glycosides as a mixture of compounds containing a steviol backbone conjugated to any number or combination of the principal sugar moieties (glucose, rhamnose, xylose, fructose, arabinose, galactose and deoxyglucose) in any of the orientations occurring in the leaves of *Stevia rebaudiana* Bertoni. The product is obtained from the enzymatic treatment of purified steviol glycosides extracted from the leaves of *Stevia rebaudiana* Bertoni. The purified leaf extract is treated with enzymes produced by non-toxicogenic nonpathogenic strains of *Pichia pastoris* and *Escherichia coli* that have been genetically modified with genes from multiple donor organisms to produce glucosyltransferase (EC 2.4.1.17) and sucrose synthase (EC 2.4.1.13). The resulting material is heated and filtered to denature and remove the enzymes. The raw product is concentrated using resin adsorption/desorption or solid/liquid filtration, followed by purification and preparation of the product of commerce using processes that may include decolonization, crystallization, and spray drying. This manufacturing technique maximizes the production of specific steviol glycosides that are not naturally present in high concentrations in the leaf extract, primarily rebaudioside M and rebaudioside D with minor amounts of other steviol glycosides. (87th JECFA (2019) and published in FAO Monographs 23 (2019)).

The U.S. FDA has reviewed the safety of over 50 different steviol glycoside preparations and have consistently raised no objections regarding the GRAS status of steviol glycosides for use as general-purpose sweeteners in

food and beverage products. Of note, the U.S. FDA did not raise any objections regarding GRN 667 and GRN 745, in relation to which describes the GRAS status of rebaudioside M produced by an enzymatic bioconversion process for use as a general-purpose sweetener in foods (U.S. FDA, 2017, 2018). The rebaudioside M described in GRN 667 and 745 is similar to Manus Bio's high purity rebaudioside M ($\geq 95\%$ rebaudioside M) produced by enzymatic conversion of steviol glycosides. In the Manus Bio process the food ingredient is produced by conversion of stevia leaf extract using UDP-glucosyltransferase enzymes produced by *E. coli* K-12 similar to GRN 745.

A comprehensive search of the scientific literature was conducted through November 2020 to capture publications relevant to the safety of steviol glycosides that became available following the U.S. FDA review of GRN 882². The search identified studies investigating the genotoxicity, antidiabetic, and immune effects of steviol glycosides. Upon review, the GRAS Panel noted that the results of these studies also support the safety conclusions on steviol glycosides as established by a number of scientific and regulatory authorities/bodies (e.g., JECFA, FSANZ, U.S. FDA, EFSA, and Health Canada).

The parental strain, *E. coli* K-12, from which the production strains are derived, has an extensive history of use as a laboratory organism and is considered one of the most extensively characterized microorganisms (Bachmann, 1972; Jensen, 1993). *E. coli* K-12 has a long history of safe use in the food and pharmaceutical industries. For example, a genetically modified strain of *E. coli* K-12 producing chymosin was affirmed as GRAS by the U.S. FDA in 1990 (Flamm, 1991; Olempska-Beer *et al.*, 2006). *E. coli* also serves as a host for the production of enzymes currently used in a GRAS-approved process for the enzymatic conversion of steviol glycosides. (GRN745). Further, *E. coli* K-12 and its derivatives are unable to colonize the mammalian gastrointestinal tract, and do not produce toxins and are unable to persist in the soil and water (Bogosian *et al.*, 1996; U.S. EPA, 1997). The parental strain does not carry any introduced antibiotic resistance genes and the complete genome of this strain has been sequenced, confirming the absence of any toxigenic potential (Blattner *et al.*, 1997; Hayashi *et al.*, 2006). The identity of the final production strain was also confirmed by both Sanger sequencing of engineered regions and whole genome sequencing.

The scientific evidence reviewed by the GRAS Panel demonstrates that under the conditions of intended use, Manus Bio's high purity rebaudioside M ($\geq 95\%$ rebaudioside M) would not produce any adverse health effects.

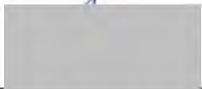
² At the time of the GRAS Panel's evaluation, GRN 882 was the most recent GRAS notice reviewed by the U.S. FDA to receive a "no questions" letter.

CONCLUSION

We, the members of the GRAS Panel, have, independently and collectively, critically evaluated the data and information summarized above and conclude that Manus Bio's high-purity rebaudioside M ($\geq 95\%$ rebaudioside M), meeting appropriate food-grade specifications and produced in accordance with current Good Manufacturing Practice (cGMP), is safe for use as a general-purpose sweetener in foods and beverages.

We further unanimously conclude that the proposed use of Manus Bio's high-purity rebaudioside M ($\geq 95\%$ rebaudioside M) meeting appropriate food-grade specifications, as presented in the supporting dossier and produced consistent with cGMP is Generally Recognized as Safe (GRAS) under its intended conditions of use as a general-purpose sweetener in food and beverages based on scientific procedures.

It is our opinion that other qualified experts would concur with these conclusions.


Ashley Roberts Ph.D.
President – AR Toxicology Inc.

8 December 2020

Date


Jose -Avalos, Ph.D.
Princeton University
Princeton, NJ.

12/03/2020

Date


Stanley M. Tarka, Jr., Ph.D.
Fellow of ATS
The Tarka Group Inc.
The Pennsylvania State University,
College of Medicine

09 December 2020

Date

APPENDIX B: Manufacturing Process Details

The process starts by the stevia leaf containing glycosides being sterilized and then dissolved with the media in a seed preparation. The seed is grown for 12 hours at 36-38 C. Media is prepared in the main tank and the seed is transferred where the conversion takes place over 36 hours while feeding the glycoside solution. The solution is heated to 70-75C and pH adjusted to 2.5-3.5 for 2 hours. The next step is separation of the biomass through a disc stacked centrifuge. The clarified broth containing Rebaudioside M is captured after passing through a 0.2-1.0 micron filter while the biomass is disposed of and not reprocessed. The clarified broth has the pH increased to 5.5 and is cooled to 10C in a primary crystallization step. The material is then centrifuged yielding a wet cake or crude rebaudioside M. The crude wet cake is dissolved in water/ ethanol, pH increased with sodium carbonate to 10.5-11.5, passed through a carbon column followed by a 0.2 to 0.45 micron filter, and heated to 70C. In the next step the material is pH adjusted to 5.5, cooled down to 10C in a secondary crystallization process. Crystals of rebaudioside are collected through centrifugation, they are washed, vacuum dried at 90-110C for 6 hours. Once dry, the material is milled and packaged for commercial sale. |

APPENDIX C: Three nonconsecutive lot analysis



1752 Lowers Lane,
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Ph.: (1) 706-503-5600

Certificate of Analysis

Specification Parameter	High-Purity Rebaudioside M (≥95% Rebaudioside M)	Manufacturing Lot No MAM060420A Result	Method of Analysis
Physical and Chemical Parameters			
Appearance	White to off-white powder	Pass	
Total steviol glycosides (anhydrous basis)	≥95%	>95	TN34236 [Monograph 23 (87, JECFA Meeting 2019)]
Rebaudioside M	≥95%	97.93	TN34240 [Monograph 23 (87, JECFA Meeting 2019)]
Loss on drying	≤6%	4.2	TN46040 (CRA E-46)
pH (1% solution)	4.5 to 7.0	Pass	TN60730 (AOAC 981.12)
Residual ethanol	≤5,000 ppm	Not Detected	TN64080 (USP 32-NF 27)
Residual methanol	≤200 ppm	Not Detected	TN64080 (USP 32-NF 27)
Total ash	≤1%	<1%	TN09560 (AOAC 900.02)
Lead	≤1 ppm	<1 ppm	TN44290 (AOAC 993.14)
Arsenic	≤1 ppm	<1 ppm	TN44292 (AOAC 993.14)
Cadmium	≤1 ppm	<1 ppm	TN44291 (AOAC 993.14)
Mercury	≤1 ppm	<1 ppm	TN44293 (AOAC 993.14)
Microbial Parameters			
Specification Parameter	High-Purity Rebaudioside M (≥95% Rebaudioside M)	Manufacturing Lot No Result	Method of Analysis
Total plate count	<1,000 CFU/g	50 CFU/g	TN10547 (CRA Microbiological Methods V-A)
Mold	<100 CFU/g	<10 CFU/g	TN10560 (CRA Microbiological Methods I-B)
Yeast	<100 CFU/g	<10 CFU/g	TN47010 (CRA Microbiological Methods II-A-1)
Coliforms	<3 MPN/g	<3 MPN/g	TN97010 (CRA Microbiological Methods I-A)
<i>Escherichia coli</i>	Not detected	Not detected	TN10510 (CRA Microbiological Methods IV-B)
<i>Salmonella</i>	Negative/25 g	Negative	TN10512 (CRA Microbiological Methods IV-B)



K. E. Furlong, VP Quality, 04 Nov 20





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Certificate of Analysis

Specification Parameter	High-Purity Rebaudioside M (≥95% Rebaudioside M)	Manufacturing Lot No. MAM060520E Result	Method of Analysis
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Physical and Chemical Parameters

Appearance	White to off-white powder	Pass	
Total steviol glycosides (anhydrous basis)	≥95%	>95	TN34236 [Monograph 23 (87 th JECFA Meeting 2019)]
Rebaudioside M	≥95%	97.01	TN34240 [Monograph 23 (87 th JECFA Meeting 2019)]
Loss on drying	≤6%	3.98	TN46040 (CRA E-46)
pH (1% solution)	4.5 to 7.0	Pass	TN60730 (AOAC 981.12)
Residual ethanol	≤5,000 ppm	Not Detected	TN64080 (USP 32-NF 27)
Residual methanol	≤200 ppm	Not Detected	TN64080 (USP 32-NF 27)
Total ash	≤1%	<1%	TN09560 (AOAC 900.02)
Lead	≤1 ppm	<1 ppm	TN44290 (AOAC 995.14)
Arsenic	≤1 ppm	<1 ppm	TN44292 (AOAC 995.14)
Cadmium	≤1 ppm	<1 ppm	TN44291 (AOAC 995.14)
Mercury	≤1 ppm	<1 ppm	TN44293 (AOAC 995.14)

Microbial Parameters

Specification Parameter	High-Purity Rebaudioside M (≥95% Rebaudioside M)	Manufacturing Lot No. Result	Method of Analysis
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Total plate count	<1,000 CFU/g	70 CFU/g	TN10547 (CRA Microbiological Methods V-A)
Mold	<100 CFU/g	<10 CFU/g	TN10560 (CRA Microbiological Methods I-A)
Yeast	<100 CFU/g	<10 CFU/g	TN47010 (CRA Microbiological Methods II-A-1)
Coliforms	<3 MPN/g	<3 MPN/g	TN97010 (CRA Microbiological Methods I-A)
<i>Escherichia coli</i>	Not detected	Not detected	TN10510 (CRA Microbiological Methods IV-B)
<i>Salmonella</i>	Negative/25 g	Negative	TN10512 (CRA Microbiological Methods IV-B)

K. E. Furlong, VP Quality, 04 Nov 20





1762 Evers Lane,
Augusta GA
Ph: (1) 706-383-5600

Certificate of Analysis

Specification Parameter	High Purity Rebaudioside M (≥95% Rebaudioside M)	Manufacturing Lot No MAM200716J Result	Method of Analysis
-------------------------	--------------------------------------------------	----------------------------------------------	--------------------

Physical and Chemical Parameters

Appearance	White to off-white powder	Pass	
Total steroid glycosides (anhydrous basis)	≥95%	>95	TN34236 (Monograph 23 (87) ⁴ , JECFA Meeting 2019)
Rebaudioside M	≥95%	97.3	TN34240 (Monograph 23 (87) ⁴ , JECFA Meeting 2019)
Loss on drying	≤6%	1.74	TN46040 (CRA E-46)
pH (1% solution)	4.5 to 7.0	Pass	TN60730 (AOAC 981.12)
Residual ethanol	≤3,000 ppm	Not Detected	TN64080 (USP 32-NF 27)
Residual methanol	≤200 ppm	Not Detected	TN64080 (USP 32-NF 27)
Total ash	≤1%	<1%	TN09560 (AOAC 900.02)
Lead	≤1 ppm	<1 ppm	TN44290 (AOAC 993.14)
Arsenic	≤1 ppm	<1 ppm	TN44292 (AOAC 993.14)
Cadmium	≤1 ppm	<1 ppm	TN44291 (AOAC 993.14)
Mercury	≤1 ppm	<1 ppm	TN44293 (AOAC 993.14)

Microbial Parameters

Specification Parameter	High Purity Rebaudioside M (≥95% Rebaudioside M)	Manufacturing Lot No Result	Method of Analysis
-------------------------	--------------------------------------------------	--------------------------------	--------------------

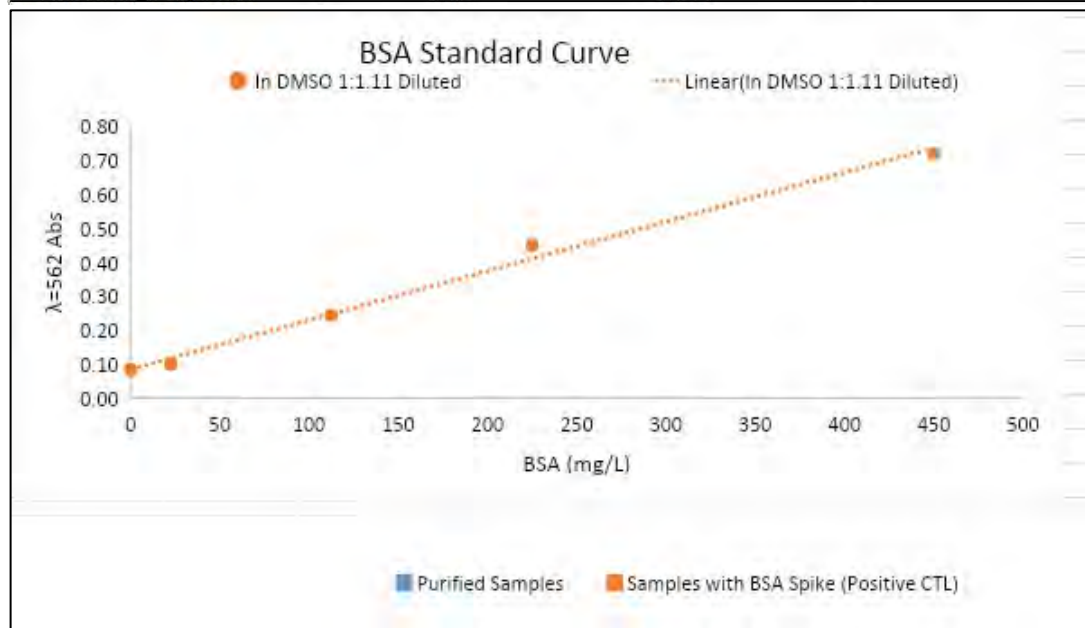
Total plate count	<1,000 CFU/g	500 CFU/g	TN10547 (CRA Microbiological Methods V-A)
Mold	<100 CFU/g	<10 CFU/g	TN10560 (CRA Microbiological Methods I-A)
Yeast	<100 CFU/g	<10 CFU/g	TN47010 (CRA Microbiological Methods II-A-1)
Coliforms	<3 MPN/g	<3 MPN/g	TN97010 (CRA Microbiological Methods I-A)
Escherichia coli	Not detected	Not detected	TN10510 (CRA Microbiological Methods IV-B)
Salmonella	Negative/25 g	Negative	TN10512 (CRA Microbiological Methods IV-B)

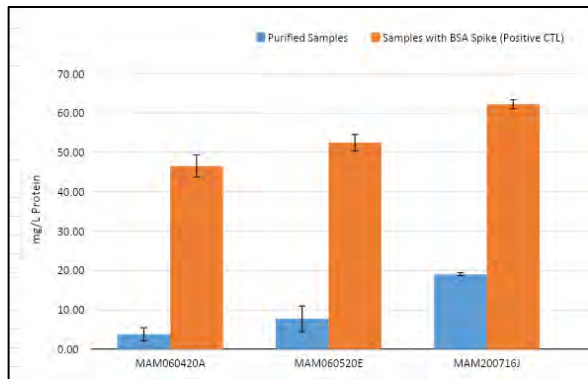
K. E. Furlong, VP Quality, 4 Nov 20



APPENDIX D: Residual Protein analysis

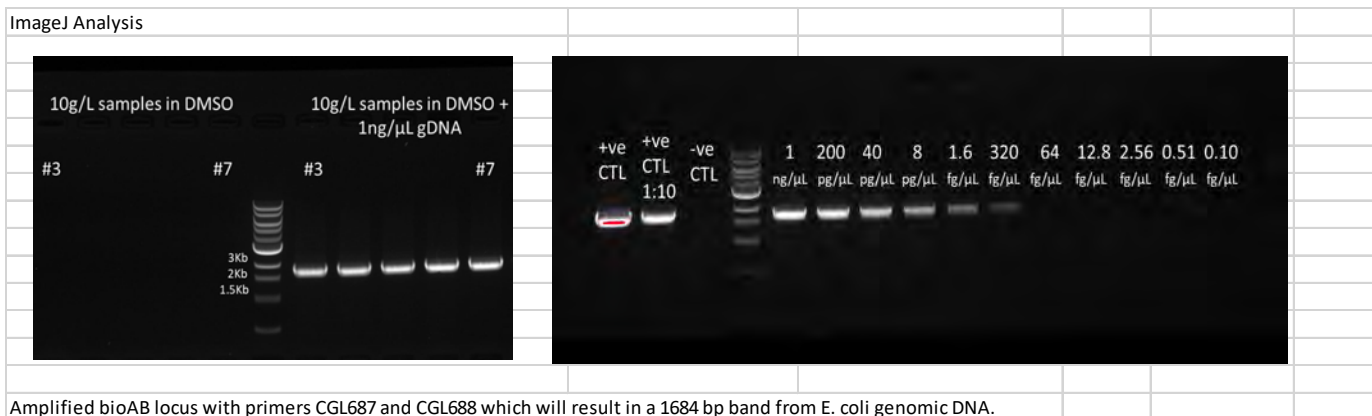
BCA Assay Protocol:	https://www.thermofisher.com/document-connect/document-connect.html?url=https%3A%2F%2Fassets.thermofisher.com%2Fassets%2Fmanuals%2FMAN0011430_Pierce_BCA_Protein_Asy_UG.pdf&title=VXNlciBHdWlkZTogUGllcmNlIEJDOSB0cm90ZWluIEFzc2F5IEtpdA==				
	Standard Curve				
Standard Concentrations (mg/L):	1800	1350	900	675	
Response (Abs λ 562):	2.31	1.80	1.41	1.05	
S-Assets%2FASSETS%2FManuals%2FMAN0011430_Pierce_BCA_Protein_Asy_UG.pdf&title=VXNlciBHdWlkZTogUGllcmNlIEJDOSB0cm90ZWluIEFzc2F5IEtpdA==					
	450	225	113	23	0
	0.72	0.45	0.24	0.10	0.08
Sample Name	Raw (Abs λ 562)	Fit Concentration (mg/L)	Average (mg/L)	Standard Deviation (mg/L)	CTL Difference (mg/L)
MAM060420A	0.09	4.93	3.73	1.70	
MAM060420A	0.09	2.53			
MAM060520E	0.09	5.40	7.70	3.25	
MAM060520E	0.10	10.00			
MAM200716J	0.11	19.27	19.00	0.38	
MAM200716J	0.11	18.73			
MAM060420A_40mg/L BSA	0.15	44.53	46.53	2.83	42.80
MAM060420A_40mg/L BSA	0.16	48.53			
MAM060520E_40mg/L BSA	0.17	53.93	52.47	2.07	44.77
MAM060520E_40mg/L BSA	0.16	51.00			
MAM200716J_40mg/L BSA	0.18	63.00	62.20	1.13	43.20
MAM200716J_40mg/L BSA	0.18	61.40			





APPENDIX E: Residual DNA Analysis

GEL 1



Amplified bioAB locus with primers CGL687 and CGL688 which will result in a 1684 bp band from *E. coli* genomic DNA.

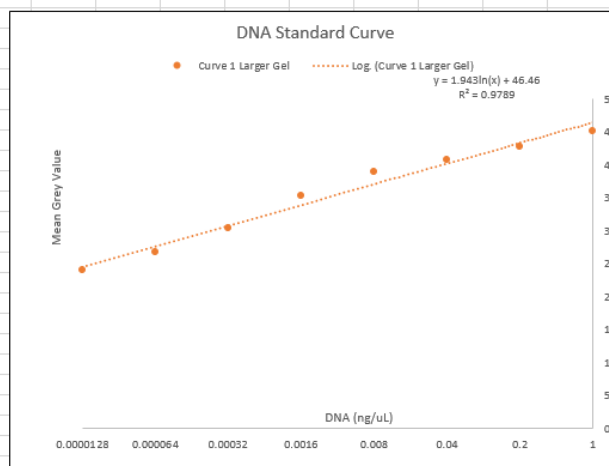
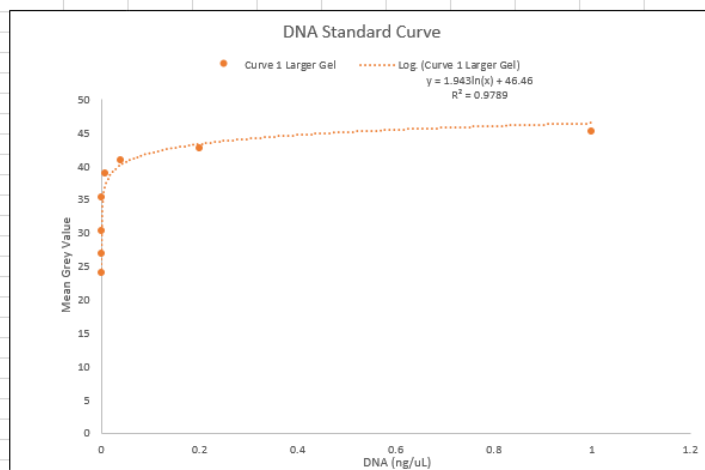
ImageJ Quant					
Concentration (ng/μL)	Mean Grey Value Product	Mean Grey Value Background	Bg Sub	Ln Conc	Fit
1	80.307	17.417	62.89	0	0.292046
0.2	74.422	17.55	56.872	-1.609437912	0.132059
0.04	64.741	16.889	47.852	-3.218875825	0.040193
0.008	51.938	16.088	35.85	-4.828313737	0.008255
0.0016	37.525	14.701	22.824	-6.43775165	0.001481
0.00032	25.574	14.059	11.515	-8.047189562	0.000333
0.000064	13.152	13.381	-0.229	-9.656627475	7.08E-05
0.0000128	12.839	13.093	-0.254	-11.26606539	7.06E-05
0.00000256	12.159	12.519	-0.36	-12.8755033	6.96E-05
0.000000512	12.184	11.547	0.637	-14.48494121	7.94E-05
1.024E-07	10.293	10.717	-0.424	-16.09437912	6.9E-05
Limit of detection					
	-9.524833171				
	7.30159E-05				
	73 fg/μL				
Sample					
Raw	Background	Bg Sub	Fit		
3.MAM200716J	11.289	11.72	-0.431	6.89814E-05	
3.MAM200716J + 1ng/μL DNA	69.686	15.667	54.019		

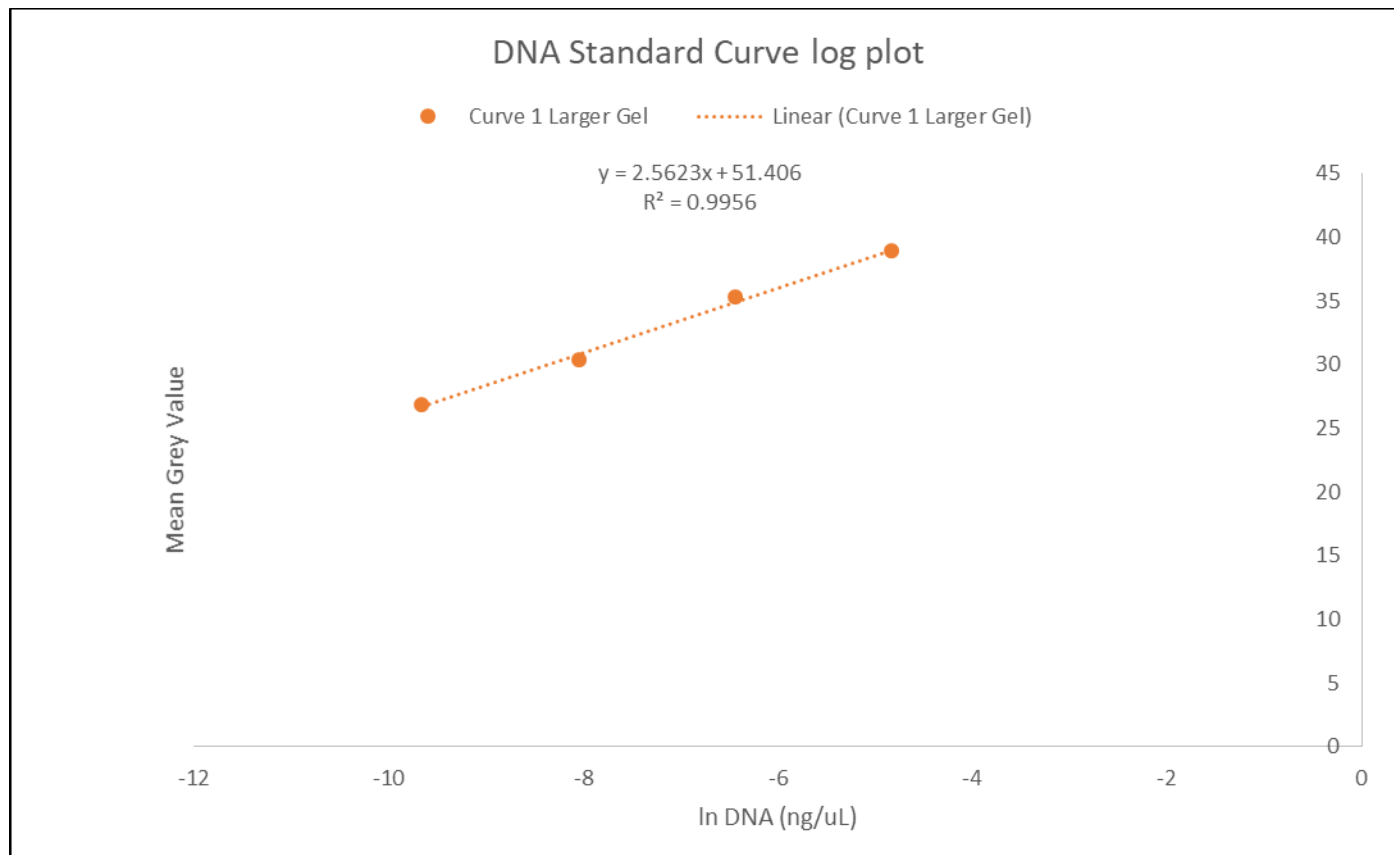
Amplified bioAB locus with primers CGL687 and CGL688 which will result in a 1684 bp band from E. coli genomic DNA.

ImageJ Quant				
Concentration (ng/μL)	Mean Grey Value Product	Mean Grey Value Background	Bg Sub	Ln Conc
1	67.148	21.967	45.181	0
0.2	65.117	22.383	42.734	-1.609437912
0.04	62.636	21.839	40.797	-3.218875825
0.008	60.004	21.07	38.934	-4.828313737
0.0016	55.714	20.432	35.282	-6.43775165
0.00032	49.768	19.425	30.343	-8.047189562
0.000064	44.852	18.018	26.834	-9.656627475
0.0000128	40.358	16.343	24.015	-11.26606539

Limit of detection (extrapolated)	
	-20.0624439
	1.93638E-09
	1.93 atto g/μL

Sample	Raw	Background	Bg Sub	Fit	
1.MAM060420A		14.327	15.204	-0.877	6.5041E-05
2.MAM060520E		15.985	16.756	-0.771	6.59566E-05
1.MAM060420A + 1ng/μL DNA		67.35	21.35	46	
2.MAM060520E + 1ng/μL DNA		67.912	22.374	45.538	






Summary Table

DNA Analysis - Summary.


Sample	Background Subtracted Mean Grey Value	Calculated Concentration (ng/μL)
1.MAM060420A	-0.877	below LOD
2.MAM060520E	-0.771	below LOD
3.MAM200716J	-0.431	below LOD

*LOD = limit of detection

APPENDIX F: Pesticide Residue

REPORT NUMBER 20-282-4059		 <p>Midwest Laboratories 13611 B Street • Omaha, Nebraska 68144-3693 • (402) 334-7770 www.midwestlabs.com</p>		PAGE 1/10		
REPORT DATE Oct 08, 2020	SEND TO 30263			ISSUE DATE Oct 08, 2020		
RECEIVED DATE Oct 02, 2020						
<p>MANUS BIO INC MANUS BIO INC 1762 LOVERS LANE AUGUSTA GA 30903-1869</p>		<p>REPORT OF ANALYSIS For: (30263) MANUS BIO INC Other (provide in comments or see FAQ) Pesticide</p>				
Analysis	Level Found As Received	Units	Reporting Limit	Method	Analyst- Date	Verified- Date
Sample ID: IOT 202008081	Lab Number: 8814029	Date Sampled: 2020-10-01				
alpha-BHC (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
beta-BHC (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
gamma-BHC (Lindane) (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
delta-BHC (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Heptachlor (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Aldrin (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Heptachlor epoxide (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
alpha Chlordane (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
gamma Chlordane (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
pp-DDE (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Endrin (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Dieldrin (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Endosulfan I (alpha) (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Endosulfan II (beta) (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
pp-DDD (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Endosulfan sulfate (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
pp-DDT (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Endrin ketone (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Methoxychlor (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08

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<p>MANUS BIO INC MANUS BIO INC 1762 LOVERS LANE AUGUSTA GA 30903-1869</p>		<p>REPORT OF ANALYSIS For: (30263) MANUS BIO INC Other (provide in comments or see FAQ) Pesticide</p>				
Analysis	Level Found As Received	Units	Reporting Limit	Method	Analyst- Date	Verified- Date
Sample ID: IOT 202008081	Lab Number: 8814029 (con't)					
Dichlorvos (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Mevinphos-1 (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Mevinphos-2 (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Demeton-S (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Ethoprop (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Naled (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Sulfotep (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Dimethoate (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Phorate (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Demeton-O (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Diazinon (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Disulfoton (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Methyl parathion (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Ronnel (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Malathion (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Fenitrothion (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Parathion ethyl (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
TEPP (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08
Trichlorfonal (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	ajj2-2020/10/08


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**MANUS BIO INC
MANUS BIO INC
1762 LOVERS LANE
AUGUSTA GA 30903-1869**

REPORT OF ANALYSIS
For: (30263) MANUS BIO INC
Other (provide in comments or see FAQ)
Pesticide

Analysis	Level Found As Received	Reporting			Analyst- Date	Verified- Date
		Units	Limit	Method		
Sample ID: IOT 202008081 Lab Number: 8814029 (con't)						
Merphos (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Tetrachlorvinphos (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Tokuthion (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Tribufos (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Fensulfotthion (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Sulprofos (Bolstar) (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
EPN (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Azinphos-methyl (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Coumaphos (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
EPTC (Eptam) (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Butylate (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Propachlor (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Disopropylatrazine (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Ethalfuralin (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Desethyl atrazine (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Trifluralin (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Benfluraline (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Prometon (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Simazine (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08


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REPORT OF ANALYSIS
For: (30263) MANUS BIO INC
Other (provide in comments or see FAQ)
Pesticide

Analysis	Level Found As Received	Reporting			Analyst- Date	Verified- Date
		Units	Limit	Method		
Sample ID: IOT 202008081 Lab Number: 8814029 (con't)						
Atrazine (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Propazine (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Terbufos (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Fonofos (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Tnallate (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Dimethenamid (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Acetochlor (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Metribuzin (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Alachlor (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Ametryn (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Prometryn (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Bromacil (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Metolachlor (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Chlorpyrifos (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Cyanazine (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Pendimethalin (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Isoprophos (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Butachlor (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Oxadiazon (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08


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REPORT OF ANALYSIS
For: (30263) MANUS BIO INC
██████████
Other (provide in comments or see FAQ)
Pesticide

Analysis	Level Found As Received	Units	Reporting Limit	Method	Analyst- Date	Verified- Date
Sample ID: IOT 202008081	Lab Number: 8814029 (con't)					
Hexazinone (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Sample ID: Lot 202003051	Lab Number: 8814030 Date Sampled: 2020-10-01					
alpha-BHC (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
beta-BHC (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
gamma-BHC (Lindane) (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
delta-BHC (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Heptachlor (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Aldrin (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Heptachlor epoxide (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
alpha Chlordane (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
gamma Chlordane (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
pp-DDE (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Endrin (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Dieldrin (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Endosulfan I (alpha) (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Endosulfan II (beta) (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
pp-DDD (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Endosulfan sulfate (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
pp-DDT (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08


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REPORT OF ANALYSIS
For: (30263) MANUS BIO INC
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Other (provide in comments or see FAQ)
Pesticide

Analysis	Level Found As Received	Units	Reporting Limit	Method	Analyst- Date	Verified- Date
Sample ID: Lot 202003051	Lab Number: 8814030 (con't)					
Endrin ketone (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Methoxychlor (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Dichlorvos (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Mevinphos-1 (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Mevinphos-2 (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Demeton-S (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Ethioprop (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Naled (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Sulfotep (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Dimethoate (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Phorate (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Demeton-O (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Diazinon (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Disulfoton (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Methyl parathion (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Ronnel (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Malathion (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Fenthion (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08
Parathion ethyl (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bms3-2020/10/08	ahj2-2020/10/08


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REPORT OF ANALYSIS
For: (30263) MANUS BIO INC
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Other (provide in comments or see FAQ)
Pesticide

Analysis	Level Found As Received	Units	Reporting		Analyst- Date	Verified- Date
			Limit	Method		
Sample ID: Lot 202003051 Lab Number: 8814030 (cont)						
TEPP (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Trichloronat (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Merphos (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Tetrachlorvinphos (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Toluthion (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Tribufos (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Fensulfothion (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Sulprofos (Bolstar) (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
EPN (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Azinphos-methyl (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Coumaphos (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
EPTC (Eptam) (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Butylate (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Propachlor (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Deisopropylatrazine (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Ethalfuralin (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Desethyl atrazine (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Trifluralin (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Benfluraline (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08


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For: (30263) MANUS BIO INC
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Pesticide

Analysis	Level Found As Received	Units	Reporting		Analyst- Date	Verified- Date
			Limit	Method		
Sample ID: Lot 202003051 Lab Number: 8814030 (cont)						
Prometon (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Simazine (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Atrazine (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Propazine (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Terbufos (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Fonofos (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Triallate (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Dimethenamid (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Acetochlor (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Meflufuzin (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Alachlor (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Ametryn (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Prometryn (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Bromacil (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Metolachlor (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Chlorpyrifos (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Cyanazine (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Pendimethalin (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08
Isophenphos (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	brm3-2020/10/08	akj2-2020/10/08

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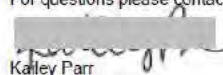
For: (30263) MANUS BIO INC

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Pesticide

Analysis	Level Found	Units	Reporting		Analyst- Date	Verified- Date
	As Received		Limit	Method		
Sample ID: Lot 202003051	Lab Number: 8814030 (con't)					
Butachlor (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bmi3-2020/10/08	akj2-2020/10/08
Oxadiazon (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bmi3-2020/10/08	akj2-2020/10/08
Hexazinone (QuEChERS)	n.d.	ppm	0.01	AOAC 2007.01 (mod)	bmi3-2020/10/08	akj2-2020/10/08

All results are reported on an AS RECEIVED basis., n.d. = not detected , ppm = parts per million, ppm = mg/kg

For questions please contact:


Kaley Parr
Account Manager
kparr@midwestlabs.com (402)829-9883

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Appendix G Literature Search

Search Strategy

Databases: BIOSIS® Toxicology, BIOSIS Previews®, CAB ABSTRACTS, Embase®, Lancet Titles, MEDLINE®, New England Journal of Medicine, NTIS: National Technical Information Service, Registry of Toxic Effects of Chemical Substances (RTECS®), ToxFile®, TOXLINE, TOXLINE

Set#	Searched for	Results
S1	((("steviol glycoside" or steviol or rebaudioside or stevioside or steviolbioside or rubusoside or dulcoside)) and (pd(>20190327) and ud(>20190327)))	266°

° Duplicates are removed from the search and from the result count.

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Metabolic fate in adult and pediatric population of steviol glycosides produced from stevia leaf extract by different production technologies

Author: Purkayastha, Sidd 1 ; Kwok, David 2 1 PureCircle Limited, 200 W Jackson Blvd, 8th Floor, Chicago, IL, 60606, United States sidd.purkayastha@purecircle.com 2 BRI Biopharmaceutical Research Inc., 101-8898 Heather St., Vancouver, BC V6P 3S8, Canada

Publication info: Regulatory Toxicology and Pharmacology 116 (Oct 2020).

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 2 of 266

Structural dependence of antidiabetic effect of steviol glycosides and their metabolites on streptozotocin-induced diabetic mice

Author: Myint, Khaing Zar 1 ; Chen, Jun-Ming 2 ; Zhou, Zhuo-Yu 1 ; Xia, Yong-Mei 1 ; Lin, Jianguo 3 ; Zhang, Jue 3 1 State Key Laboratory of Food Science and Technology, Jiangnan University, Wuxi, China, Key Laboratory of Synthetic and Biological Colloids (Ministry of Education), School of Chemical and Materials Engineering, Jiangnan University, Wuxi, China, China 2 State Key Laboratory of Food Science and Technology, Jiangnan University, Wuxi, China, China 3 Key Laboratory of Nuclear Medicine of Ministry of Health, Jiangsu Institute of Nuclear Medicine, Wuxi, China, China

Publication info: Journal of the science of food and agriculture 100.10 (Aug 2020): 3841-3849.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 3 of 266

Rapid and economic determination of 13 steviol glycosides in market-available food, dietary supplements, and ingredients: single-laboratory validation of an HPLC method.

Author: Liu, ZhiYan; Ren KangZi; Ye, Feng; Uong, T; Krepich, S; You, Hong

Publication info: Journal of Agricultural and Food Chemistry 68.37 (2020): 10142-10148.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 4 of 266

Physiological and morphological responses to abiotic stresses in two cultivars of *Stevia rebaudiana* (Bert.) Bertoni.

Author: Debnath, M; Ashwath, N; Midmore, D J

Publication info: South African Journal of Botany 123 (2019): 124-132.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

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A Validated RP-HPLC Method for Quantification of Steviol Glycoside: Rebaudioside A in Extracts of Stevia Rebaudiana Leaf

Author: Kolate, Nikhil S. 1 ; Mishra, Himanshu 1 ; Kini, Suvarna G. 1 ; Raghavan, Govindarajan 2 ; Vyas, Tejas B. 2 1

Department of Pharmaceutical Chemistry, Manipal College of Pharmaceutical Sciences, MAHE, Manipal, Karnataka, 576104, India suvarna.gk@manipal.edu 2 Analytical Development Laboratory, Zydus Wellness Ltd, Ahmedabad, Gujarat, 380015, India

Publication info: Chromatographia (2020).

[ProQuest document link](#)

Databases: Embase® (1947 - current)

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Factors affecting organogenesis of *Stevia rebaudiana* and *in vitro* accumulation of steviol glycosides.

Author: Blinstrubienė, A; Burbulis, N; Juškevičiūtė, N; Žūkienė, R

Publication info: Ūkio žurnalas (Agriculture) 107.2 (2020): 171-178.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

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Microwave-Assisted Subcritical Water Extraction of Steviol Glycosides from Stevia rebaudiana Leaves

Author: Yang, Zheng 1 ; Uhler, Brandon 1 ; Lipkie, Tristan 1 1 Cargill Inc., 14800 28th Ave N, Minneapolis, MN, 55447, United States zheng_yang@cargill.com

Publication info: Natural Product Communications 14.6 (2019).

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 8 of 266

Enzymatic synthesis and characterization of a novel α -1->6-glucosyl rebaudioside C derivative sweetener.

Author: Yang, Zheng; Uhler, B; Zheng, Ted; Adams, K M

Publication info: Biomolecules 9.1 (2019): 27-27.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

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***Stevia Rebaudiana* Bertoni, a source of high-potency natural sweetener-biochemical and genetic characterization.**

Author: Dyduch-Siemińska, M; Najda, A; Gawroński, J; Balant, S; świca, K; Żaba, A

Publication info: Molecules 25.4 (2020).

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

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Interaction of Organic Anion Transporter 3-Mediated Uptake of Steviol Acyl Glucuronide, a Major Metabolite of Rebaudioside A, with Selected Drugs

Author: Zhou, Dandan 1 ; Xu, Yunting 1 ; Wang, Yedong 1 ; Li, Jiajun 1 ; Gui, Chunshan 1 ; Zhang, Hongjian 1 1 College of Pharmaceutical Sciences, Soochow University, Suzhou, 215006, China

Publication info: Journal of agricultural and food chemistry 68.6 (Feb 12, 2020): 1579-1587.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 11 of 266

Comparison of uridine diphosphate-glycosyltransferase UGT76G1 genes from some varieties of *Stevia rebaudiana* Bertoni

Author: Abdelsalam, Nader R.; Botros, William A.; Khaled, Ahmed E.; Ghonema, Mohamed A.; Hussein, Shima G.; Ali, Hayssam M.; Elshikh, Mohamed S.

Publication info: Scientific Reports 9 (Jun 12, 2019): Article No.: 8559.

[ProQuest document link](#)

Databases: BIOSIS Previews® (1926 - current)

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Expression and characterization of a recombinant stevioside hydrolyzing beta-glycosidase from *Enterococcus casseliflavus*

Author: Boonkaew, Bootsakorn; Udompaisarn, Somsiri; Arthan, Dumrongkiet; Somana, Jamorn

Publication info: Protein Expression and Purification 163 (Nov 2019): Article No.: 105449.

[ProQuest document link](#)

Databases: BIOSIS Previews® (1926 - current)

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Steviol glycosides profile in *Stevia rebaudiana* Bertoni hairy roots cultured under oxidative stress-inducing conditions

Author: Libik-Konieczny, Marta 1 ; Michalec-Warzecha, Żaneta 1 ; Dziurka, Michał 1 ; Zastawny, Olga 2 ; Konieczny, Robert 2 ; Rozpądek, Piotr 3 ; Pistelli, Laura 4 1 The Franciszek Górski Institute of Plant Physiology, Polish Academy of Sciences, ul. Niezapominajek 21, 30-239, Krakow, Poland, Poland m.libik@ifr-pan.edu.pl 2 Department of Plant Cytology and Embryology, Jagiellonian University, Gronostajowa 9, 30-387, Krakow, Poland, Poland 3 Malopolska Centre of Biotechnology, Jagiellonian University, Gronostajowa 7A, 30-387, Krakow, Poland, Poland 4 Department of Agriculture, Food and Environment, University of Pisa, Via del Borghetto 80, I-56124, Pisa, Italy, Italy

Publication info: Applied microbiology and biotechnology 104.13 (Jul 2020): 5929-5941.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 14 of 266

Effect of gibberellic acid on production of biomass, polyphenolics and steviol glycosides in adventitious root cultures of *Stevia rebaudiana* (Bert.).

Author: Ashfaq, Ahmad; Haider, Ali; Khan, Habiba; Begam, Almas; Khan, Sheraz; Ali, S S; Ahmad, Naveed; Fazal, Hina; Ali, Mohammad; Hano, C; Nisar, Ahmad; Abbasi, B H

Publication info: Plants 9.4 (2020).

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

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EXTRACTION OF STEVIOL GLYCOSIDES FROM DRIED STEVIA REBAUDIANA BY PRESSURIZED HOT WATER EXTRACTION

Author: Nemeth, A.; Janosi, S. Z.

Publication info: Acta Alimentaria 48.2 (Jun 2019): 241-252.

[ProQuest document link](#)

Databases: BIOSIS Previews® (1926 - current)

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Bioconversion of Stevioside to Rebaudioside E Using Glycosyltransferase UGTSL2

Author: Chen, Liangliang 1 ; Pan, Huayi 1 ; Cai, Ruxin 1 ; Li, Yan 1 ; Jia, Honghua 1 ; Chen, Kequan 1 ; Yan, Ming 1 ; Ouyang, Pingkai 1 1 College of Biotechnology and Pharmaceutical Engineering, Nanjing Tech University, Nanjing, 211800, China
liyan@njtech.edu.cn

Publication info: Applied biochemistry and biotechnology (Oct 15, 2020).

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 17 of 266

Total Synthesis and Structural Revision of Rebaudioside S, a Steviol Glycoside

Author: Wen, Guo-En 1 ; Liu, Hui 1 ; Yin, Qi-Shuang 1 ; Liao, Jin-Xi 1 ; Tu, Yuan-Hong 1 ; Zhang, Qing-Ju 1 ; Sun, Jian-Song 1 1 National Research Centre for Carbohydrate Synthesis, Jiangxi Normal University, 99 Ziyang Avenue, Nanchang 330022, China, China

Publication info: The Journal of organic chemistry (Apr 23, 2020).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 18 of 266

Extraction of steviol glycosides from *Stevia rebaudiana* (Bertoni) leaves by high-speed shear homogenization extraction.

Author: Xu ShaoHe; Wang, GuanYu; Guo RuiLi; Zhong, Wei; Zhang, JinLi

Publication info: Journal of Food Processing and Preservation 43.12 (2019): e14250.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 19 of 266

Deciphering performances of fifteen genotypes of *Stevia rebaudiana* in southwestern France through dry biomass and steviol glycoside evaluation.

Author: Hastoy, C; Cosson, P; Cavaignac, S; Boutié, P; Waffo-Teguo, P; Rolin, D; Schurdi-Levraud, V

Publication info: Industrial Crops and Products 128 (2019): 607-619.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 20 of 266

Photosynthetic performance and stevioside concentration are improved by the arbuscular mycorrhizal symbiosis in *Stevia rebaudiana* under different phosphate concentrations

Author: Sarmiento-López, Luis G. 1 ; López-Meyer, Melina 2 ; Sepúlveda-Jiménez, Gabriela 1 ; Cárdenas, Luis 3 ; Rodríguez-Monroy, Mario 1 1 Departamento de Biotecnología, Centro de Desarrollo de Productos Bióticos, Instituto Politécnico Nacional, Yautepec, Morelos, Mexico mrmonroy@ipn.mx 2 Departamento de Biotecnología Agrícola, Centro Interdisciplinario de Investigación para el Desarrollo Integral Regional Unidad Sinaloa, Instituto Politécnico Nacional, Guasave, Sinaloa, Mexico 3 Departamento de Biología Molecular de Plantas, Instituto de Biotecnología, Universidad Nacional Autónoma de México, Cuernavaca, Morelos, Mexico

Publication info: PeerJ 8 (Oct 19, 2020).

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 21 of 266

An in-silico layer-by-layer adsorption study of the interaction between Rebaudioside A and the T1R2 human sweet taste receptor: modelling and biosensing perspectives

Author: Arodola, Olayide A 1 ; Kanchi, Suvadhan 1 ; Hloma, Phathisanani 1 ; Bisetty, Krishna 1 ; Asiri, Abdullah M 2 ; Inamuddin 3 1 Department of Chemistry, Durban University of Technology, P.O Box 1334, Durban, 4000, South Africa, South Africa Olayide.rodola@gmail.com ; bisettyk@dut.ac.za 2 Chemistry Department, Faculty of Science, King Abdulaziz University, Jeddah, 21589, Saudi Arabia, Saudi Arabia 3 Chemistry Department, Faculty of Science, King Abdulaziz University, Jeddah, 21589, Saudi Arabia, Advanced Functional Materials Laboratory, Department of Applied Chemistry, Faculty of Engineering and Technology, Aligarh Muslim University, Aligarh, 202 002, India, India inamuddin@zhcet.ac.in

Publication info: Scientific reports 10.1 (Oct 27, 2020): 18391.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 22 of 266

Increase in steviol glycosides production from *Stevia rebaudiana* Bertoni under organo-mineral fertilization.

Author: Díaz-Gutiérrez, C; Hurtado, A; Ortiz, A; Poschenrieder, C; Arroyave, C; Peláez, C

Publication info: Industrial Crops and Products 147 (2020).

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 23 of 266

Effects of *Septoglomus viscosum* inoculation on biomass yield and steviol glycoside concentration of some *Stevia rebaudiana* chemotypes.

Author: Tedone, L; Ruta, C; Cillis, F de; Mastro, G de

Publication info: Scientia Horticulturae 262 (2020): 109026.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 24 of 266

Iron nano modulated growth and biosynthesis of steviol glycosides in *Stevia rebaudiana*.

Author: Khan, M A; Ali, Amir; Sher Mohammad; Ali, Huma; Khan, Tariq; Zia-Ur-Rehman Mashwani; Asif, Jan; Pervaiz Ahmad

Publication info: Plant Cell, Tissue and Organ Culture 143.1 (2020): 121-130.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 25 of 266

Arbuscular mycorrhizal symbiosis and phosphorus supply induce changes in photosynthetic rate, sugar partitioning and steviol glycosides composition in stevia.

Author: Tavarini, S; Martini, A; Guglielminetti, L; Scartazza, A; Avio, L; Angelini, L G

Publication info: Agrochimica 63.1 (2019): 85-102.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 26 of 266

Determination of potential compounds of stevia leaves (Stevia rebaudiana Bertoni) against DPP4 as candidates for antidiabetic drugs

Author: Junaedi, Effan Cahyati 1 ; Megantara, Sandra 2 ; Mustarichie, Resmi 2 1 Department of Pharmaceutical Analysis and Medicinal Chemistry, Faculty of Pharmacy, Universitas Padjadjaran, Indonesia, Department of Pharmaceutical Chemical Analysis, Faculty of Mathematics and Natural Science, Universitas Garut, Indonesia 2 Department of Pharmaceutical Analysis and Medicinal Chemistry, Faculty of Pharmacy, Universitas Padjadjaran, Indonesia

Publication info: Journal of Global Pharma Technology 12.6 (Jun 1, 2020): 327-334.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

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Stevioside Has the Maximum Biological Activity among Natural Stevia Diterpenes

Author: Ogorodnova, U.A. 1 ; Sapunova, A.S. 1 ; Timofeeva, O.A. 1 ; Mironov, V.F. 2 1 Kazan Federal University, Kazan, Russian Federation 2 Arbuzov Institute of Organic and Physical Chemistry, Kazan Scientific Center, Russian Academy of Sciences, Kazan, Tatarstan, Russian Federation

Publication info: Doklady biological sciences : proceedings of the Academy of Sciences of the USSR, Biological sciences sections 492.1 (May 1, 2020): 79-82.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 28 of 266

Mutations in the uridine diphosphate glucosyltransferase 76G1 gene result in different contents of the major steviol glycosides in Stevia rebaudiana

Author: Zhang, Shao-Shan 1 ; Chen, Hong 1 ; Xiao, Jie-Yu 1 ; Liu, Qiong 1 ; Xiao, Ren-Feng 1 ; Wu, Wei 1 1 Agronomy College, Sichuan Agricultural University, Chengdu, 611130, China ewuwei@sicau.edu.cn

Publication info: Phytochemistry 162 (Jun 1, 2019): 141-147.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 29 of 266

Development of rebaudioside D polymorphs with improved solubility.

Author: Urai, S; Kudo, S; Takiyama, H

Publication info: Food Science and Technology Research 26.1 (2020): 17-23.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 30 of 266

Immunohistochemical evaluation of the influence of rebaudioside A on neurons containing acetylcholinesterase (AChE) in the rat's hippocampus and striatum.

Author: Rycerz, K; Jaworska-Adamu, J

Publication info: Medycyna Weterynaryjna 75.10 (2019): 627-631.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 31 of 266

Production of rebaudioside D from stevioside using a UGTSL2 Asn358Phe mutant in a multi-enzyme system

Author: Chen, Liangliang 1 ; Cai, Ruxin 1 ; Weng, Jingyuan 1 ; Li, Yan 1 ; Jia, Honghua 1 ; Chen, Kequan 1 ; Yan, Ming 1 ; Ouyang, Pingkai 1 1 College of Biotechnology and Pharmaceutical Engineering, Nanjing Tech University, Nanjing, 211800, China liyan@njtech.edu.cn

Publication info: Microbial Biotechnology 13.4 (Jul 1, 2020): 974-983.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 32 of 266

Effects of cytokinins, gibberellic acid 3, and gibberellic acid 4/7 on in vitro growth, morphological traits, and content of steviol glycosides in Stevia rebaudiana

Author: Pazuki, Arman 1 ; Aflaki, Fatemeh 1 ; Yücesan, Buhara 2 ; Gürel, Songül 1 1 Department of Biology, Faculty of Arts and Sciences, Bolu Abant İzzet Baysal University, Bolu, Turkey, Turkey songul_gurel@yahoo.com 2 Faculty of Natural and Agricultural Sciences, Department of Seed Science and Technology, Bolu Abant İzzet Baysal University, Bolu, Turkey, Turkey

Publication info: Plant physiology and biochemistry : PPB 137 (Apr 2019): 154-161.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 33 of 266

Co-expression of anti-miR319g and miRStv_11 lead to enhanced steviol glycosides content in Stevia rebaudiana

Author: Saifi, Monica; Yogindran, Sneha; Nasrullah, Nazima; Nissar, Umara; Gul, Irum; Abdin, M. Z.

Publication info: BMC Plant Biology 19 (Jun 24, 2019): Article No.: 274.

[ProQuest document link](#)

Databases: BIOSIS Previews® (1926 - current)

Document 34 of 266

Optimization of microwave assisted enzymatic extraction of steviol glycosides and phenolic compounds from stevia leaf.

Author: Görgüç, A; Gençdağ, E; Yılmaz, F M

Publication info: Acta Periodica Technologica 50 (2019): 69-76.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 35 of 266

Consumer-based sensory characterization of steviol glycosides (Rebaudioside A, D, and M).

Author: Tao Ran; Cho SunGeun

Publication info: Foods 9.8 (2020).

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 36 of 266

Effect of steviol glycosides on human health with emphasis on type 2 diabetic biomarkers: a systematic review and meta-analysis of randomized controlled trials.

Author: Anker, C C B; Rafiq, S; Jeppesen, P B

Publication info: Nutrients 11.9 (2019): 1965.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 37 of 266

***Agrobacterium rhizogenes*-mediated transformation enhances steviol glycosides production and growth in *Stevia rebaudiana* plantlets.**

Author: Sánchez-Cordova, Á. de J.; Capataz-Tafur, J; Barrera-Figueroa, B E; López-Torres, A; Sanchez-Ocampo, P M; García-López, E; Huerta-Heredia, A A

Publication info: Sugar Tech 21.3 (2019): 398-406.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 38 of 266

Molecular basis for branched steviol glucoside biosynthesis

Author: Lee, Soon Goo 1 ; Salomon, Eitan 2 ; Yu, Oliver 3 ; Jez, Joseph M. 4 1 Department of Biology, Washington University in St. Louis, St. Louis, MO, 63130, United States, Department of Chemistry and Biochemistry, University of North Carolina Wilmington, Wilmington, NC, 28403, United States 2 Department of Biology, Washington University in St. Louis, St. Louis, MO, 63130, United States, National Center for Mariculture, Israel Oceanographic and Limnological Research, Eilat, 8811201, Israel 3 Conagen, Inc., Bedford, MA, 01730, United States 4 Department of Biology, Washington University in St. Louis, St. Louis, MO, 63130, United States jjez@wustl.edu

Publication info: Proceedings of the National Academy of Sciences of the United States of America 116.26 (2019): 13131-13136.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 39 of 266

Crop and Steviol Glycoside Improvement in Stevia by Breeding

Author: Tavarini, S.; Passera, B.; Angelini, L. G.

Publication info: In Steviol Glycosides: Cultivation, Processing, Analysis and Applications in Food , 1-31. ROYAL SOC CHEMISTRY, (2019).

[ProQuest document link](#)

Databases: BIOSIS Previews® (1926 - current)

Document 40 of 266

Overexpression of SrUGT76 G1 in Stevia alters major steviol glycosides composition towards improved quality.

Author: Kim, MiJung; Zheng JunShi; Liao MingHui; Jang InCheol

Publication info: Plant Biotechnology Journal 17.6 (2019): 1037-1047.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 41 of 266

Safety of a proposed amendment of the specifications for steviol glycosides (E 960) as a food additive: to expand the list of steviol glycosides to all those identified in the leaves of *Stevia rebaudiana* bertoni.

Author: Younes, M; Aquilina, G; Engel, K H; Fowler, P; Fernandez, M J F; Fürst, P; Gürtler, R; Gundert-Remy, U; Husøy, T; Manco, M; Mennes, W; Moldeus, P; Passamonti, S; Shah, R; Waalkens-Berendsen, I; Wölfle, D; Wright, M; Degen, G; Giarola, A; Rincon, A M; Castle, L

Publication info: EFSA Journal 18.4 (2020).

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 42 of 266

Melatonin promotes seed germination under salinity and enhances the biosynthesis of steviol glycosides in *Stevia rebaudiana* Bertoni leaves

Author: Simlat, Magdalena 1 ; Szewczyk, Agnieszka 2 ; Ptak, Agata 1 1 Department of Plant Breeding, Physiology and Seed Science, University of Agriculture in Krakow, Krakow, Poland, Poland 2 Department of Pharmaceutical Botany, Faculty of Pharmacy, Jagiellonian University Medical College, Krakow, Poland, Poland

Publication info: PloS one 15.3 (Mar 27, 2020): e0230755.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 43 of 266

A comparative morphological and transcriptomic study on autotetraploid *Stevia rebaudiana* (bertoni) and its diploid

Author: Xiang, Zeng-xu; Tang, Xing-li; Liu, Wei-hu; Song, Chang-nian

Publication info: Plant Physiology and Biochemistry (Paris) 143 (Oct 2019): 154-164.

[ProQuest document link](#)

Databases: BIOSIS Previews® (1926 - current)

Document 44 of 266

Salt stress-induced changes in in vitro cultured *Stevia rebaudiana* Bertoni: effect on metabolite contents, antioxidant capacity and expression of steviol glycosides-related biosynthetic genes.

Author: Lucho, S R; Amaral, M. N. do; Auler, P A; Bianchi, V J; Ferrer, M Á; Calderón, A A; Braga, E J B

Publication info: Journal of Plant Growth Regulation 38.4 (2019): 1341-1353.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 45 of 266

Fractionation of *Stevia rebaudiana* aqueous extracts via two-step ultrafiltration process: towards rebaudioside a extraction.

Author: Díaz-Montes, E; Gutiérrez-Macías, P; Orozco-Álvarez, C; Castro-Muñoz, R

Publication info: Food and Bioproducts Processing 123 (2020): 111-122.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 46 of 266

Sustainable micropropagation of selected *Stevia rebaudiana* Bertoni genotypes.

Author: Kaplan, B; Duraklioglu, S; Turgut, K

Publication info: Acta Scientiarum Polonorum - Hortorum Cultus 18.6 (2019): 47-56.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 47 of 266

RQ3, A Natural Rebaudioside D Isomer, Was Obtained from Glucosylation of Rebaudioside A Catalyzed by the CGTase Toruzyme 3.0 L

Author: Guo, Qingbin 1 ; Zhang, Tongtong 2 ; Wang, Nifei 3 ; Xia, Yongmei 2 ; Zhou, Zhuoyu 2 ; Wang, Jian-Rong 4 ; Mei, Xuefeng 4 1 State Key Laboratory of Food Science and Technology, Jiangnan University, Wuxi, Jiangsu, 214122, China, State Key Laboratory of Food Nutrition and Safety, Tianjin University of Science and Technology, Ministry of Education, Tianjin, 300457, China 2 State Key Laboratory of Food Science and Technology, Jiangnan University, Wuxi, Jiangsu, 214122, China, School of Chemical and Materials Engineering, Jiangnan University, Wuxi, Jiangsu, 214122, China 3 State Key Laboratory of Food Nutrition and Safety, Tianjin University of Science and Technology, Ministry of Education, Tianjin, 300457, China 4 Pharmaceutical Analytical & Solid-State Chemistry Research Center, Shanghai Institute of Materia Medica, Chinese Academy of Sciences, Shanghai, 201203, China

Publication info: Journal of agricultural and food chemistry 67.28 (Jul 17, 2019): 8020-8028.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 48 of 266

Kinetics and thermodynamics of rebaudioside A adsorption on a strongly acidic cation exchange resin

Author: Chen, Bin 1 ; He, Jiaxin 1 ; Xiao, Xia 1 ; Li, Rong 1 1 School of Chemical Engineering, Northwest University, Xi'an, P. R. China, P. R. China

Publication info: Journal of separation science (Jul 6, 2020).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 49 of 266

Diversity-Oriented Library Synthesis from Steviol and Isosteviol-Derived Scaffolds

Author: Holth, Trinh A D 1 ; Walters, Michael A. 1 ; Hutt, Oliver E. 1 ; Georg, Gunda I. 1 1 Department of Medicinal Chemistry and Institute for Therapeutics Discovery and Development, College of Pharmacy, University of Minnesota, 717 Delaware Street Southeast, Minneapolis, MN, 55414, United States

Publication info: ACS combinatorial science 22.3 (Mar 9, 2020): 150-155.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 50 of 266

Effect of silver nanoparticle treatment on the expression of key genes involved in glycosides biosynthetic pathway in *Stevia rebaudiana* B. plant.

Author: Ramezani, M; Asghari, S; Gerami, M; Ramezani, F; Abdolmaleki, M K

Publication info: Sugar Tech 22.3 (2020): 518-527.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 51 of 266

Elicitation directed growth and production of steviol glycosides in the adventitious roots of *Stevia rebaudiana* Bertoni.

Author: Kazmi, A; Khan, M A; Mohammad, S; Ali, A; Kamil, A; Arif, M; Ali, H

Publication info: Industrial Crops and Products 139 (2019): 111530.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 52 of 266

Characterization of a new hemihydrate rebaudioside B crystal having lower aqueous solubility

Author: Dong, Jinping 1 ; Yang, Zheng 1 1 Cargill Inc., 14800 28th Ave N, Plymouth, MN, 55447, United States

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Publication info: Food Chemistry 304 (Jan 30, 2020).

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 53 of 266

Efficient Biocatalytic Preparation of Rebaudioside KA: Highly Selective Glycosylation Coupled with UDPG Regeneration

Author: Zhang, Yunyun 1 ; Xu, Shaohua 1 ; Jin, Yue 1 ; Dai, Yan 1 ; Chen, Yijun 1 ; Wu, Xuri 1 1 State Key Laboratory of Natural Medicines and Laboratory of Chemical Biology, China Pharmaceutical University, Nanjing, 211198, China yjchen@cpu.edu.cn ; xuriwu@cpu.edu.cn

Publication info: Scientific reports 10.1 (Apr 10, 2020): 6230.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 54 of 266

***Stevia rebaudiana* Bertoni responses to salt stress and chitosan elicitor**

Author: Gerami, Mahyar 1 ; Majidian, Parastoo 2 ; Ghorbanpour, Akram 3 ; Alipour, Zeinab 1 1 Department of Biology, Sana Institute of Higher Education, Sari, Iran, Iran 2 Crop and Horticultural Science Research Department, Mazandaran Agricultural and Natural Resources Research and Education Center, Agricultural Research, Education and Extension Organization (AREEO), Sari, Iran, Iran 3 3Department of Plant Biology, Kharazmi University, Tehran, Iran, Iran

Publication info: Physiology and molecular biology of plants : an international journal of functional plant biology 26.5 (May 2020): 965-974.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 55 of 266

***Stevia rebaudiana* Bertoni bioactive effects: From in vivo to clinical trials towards future therapeutic approaches**

Author: Salehi, Bahare 1 ; López, Maria Dolores 2 ; Martínez-López, Sara 3 ; Victoriano, Montserrat 4 ; Sharifi-Rad, Javad 5 ; Martorell, Miquel 4 ; F Rodrigues, Célia 6 ; Martins, Natália 7 1 Student Research Committee, School of Medicine, Bam University of Medical Sciences, Bam, Iran, Iran 2 Department of Plant Production, Faculty of Agronomy, Universidad de Concepción, Chillán, Chile, Chile 3 Department of Pharmacy, Biotechnology, and Nutrition, School of Biomedical Sciences, European University of Madrid (UEM), Madrid, Spain, Spain 4 Department of Nutrition and Dietetics, Faculty of Pharmacy, University of Concepcion, Concepcion, Chile, Chile 5 Zabol Medicinal Plants Research Center, Zabol University of Medical Sciences, Zabol, Iran, Iran 6 LEPABE-Department of Chemical Engineering, Faculty of Engineering, University of Porto, Porto, Portugal, Portugal 7 Faculty of Medicine, University of Porto, Porto, Portugal, Institute for Research and Innovation in Health (i3S), University of Porto, Porto, Portugal, Portugal

Publication info: Phytotherapy research : PTR 33.11 (Nov 2019): 2904-2917.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 56 of 266

Do Steviol Glycosides Provide Ecological Fitness to *Stevia rebaudiana* through Impact on Dietary Preference of Plant Pests and Herbivores?

Author: Deguzman, Ria R. 1 ; Midmore, David J. 1 ; Walsh, Kerry B. 1 1 Central Queensland University, Rockhampton, QLD, 4702, Australia d.midmore@cqu.edu.au

Publication info: Journal of Natural Products 82.5 (May 24, 2019): 1200-1206.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 57 of 266

Stevioside inhibits unilateral ureteral obstruction-induced kidney fibrosis and upregulates renal PPAR γ expression in mice

Author: Shen, Wei 1 ; Fan, Ke 1 ; Zhao, Ying 1 ; Zhang, Junyan 1 ; Xie, Meilin 1 1 Department of Pharmacology, College of Pharmaceutical Sciences, Soochow University, Suzhou, China, China

Publication info: Journal of food biochemistry (Oct 12, 2020): e13520.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 58 of 266

Method for increasing glycosylation of a composition comprising steviol glycosides

Author: Anonymous

Publication info: Van Den Berg, Marco Alexander. Madern, Marcella Katharina. Winter, Remko Tsjibbe. US 10472661. 12 November 2019 (20191112). US.

[ProQuest document link](#)

Databases: BIOSIS Previews® (1926 - current)

Document 59 of 266

Enzymatic Synthesis of Glucosyl Rebaudioside A and its Characterization as a Sweetener

Author: Lee, So-Hyeon 1 ; Ko, Jin-A 2 ; Kim, Hae-Soo 1 ; Jo, Min-Ho 1 ; Kim, Joong-Su 3 ; Kim, Doman 4 ; Cho, Jeong-Yong 1 ; Wee, Young-Jung 5 ; Kim, Young-Min 1 1 Dept. of Food Science & Technology, Chonnam National Univ., Gwangju, 61186, Republic of Korea, Republic of Korea 2 Radiation Breeding Research Center, Advanced Radiation Technology Inst., Korea Atomic Energy Research Inst., Jeongeup, Republic of Korea, Republic of Korea 3 Bio-industrial Process Research Center, Korea Research Inst. of Bioscience and Biotechnology, Jeongeup, 56212, Republic of Korea, Republic of Korea 4 Research Inst. of Food Industrialization, Inst. of Green Bio Science & Technology, Seoul National Univ., Pyeongchang, 25354, Korea, Korea 5 Dept. of Food Science and Technology, Yeungnam Univ., Gyeongsan, Gyeongbuk, 38541, Republic of Korea, Republic of Korea

Publication info: Journal of food science 84.11 (Nov 2019): 3186-3193.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 60 of 266

The SrWRKY71 transcription factor negatively regulates SrUGT76G1 expression in *Stevia rebaudiana*

Author: Zhang, Ting 1 ; Xu, Xiaoyang 2 ; Sun, Yuming 1 ; Gu, Chunsun 1 ; Hou, Menglan 3 ; Guan, Yunxiao 4 ; Yuan, Haiyan 1 ; Yang, Yongheng 5 1 Institute of Botany, Jiangsu Province and the Chinese Academy of Sciences, Nanjing, 210014, China; The Jiangsu Provincial Platform for Conservation and Utilization of Agricultural Germplasm, China zhangting901014@163.com ; sunyuminggagw@163.com ; chunsungu@126.com ; yuanhaiyan416@163.com 2 Institute of Botany, Jiangsu Province and the Chinese Academy of Sciences, Nanjing, 210014, China; The Jiangsu Provincial Platform for Conservation and Utilization of Agricultural Germplasm, China. Electronic address: intergoole@126.com 3 Institute of Botany, Jiangsu Province and the Chinese Academy of Sciences, Nanjing, 210014, China; The Jiangsu Provincial Platform for Conservation and Utilization of Agricultural Germplasm, China. Electronic address: hml2016t@163.com 4 College of Horticulture, Nanjing Agricultural University, Nanjing, 210095, China 5 Institute of Botany, Jiangsu Province and the Chinese Academy of Sciences, Nanjing,

210014, China; The Jiangsu Provincial Platform for Conservation and Utilization of Agricultural Germplasm, China. Electronic address: yyh8576@126.com

Publication info: Plant physiology and biochemistry : PPB 148 (Mar 1, 2020): 26-34.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 61 of 266

Establishment of proficient *in vitro* mass multiplication and regeneration system for enhanced production of stevioside and rebaudioside a in *Stevia rebaudiana*.

Author: Tufail, M B; Mustafa, G; Joyia, F A; Mushtaq, Z; Ghazala; Khan, M S

Publication info: JAPS, Journal of Animal and Plant Sciences 29.3 (2019): 796-802.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 62 of 266

Biotechnological production of natural calorie free steviol glycosides in stevia rebaudiana: An update on current scenario

Author: Kazmi, Abeer 1 ; Khan, Mubarak Ali 1 ; Mohammad, Sher 2 ; Ali, Amir 2 ; Ali, Huma 1 1 Department of Biotechnology, Faculty of Chemical and Life Sciences, Abdul Wali Khan University Mardan (AWKUM), Mardan, 23390, Pakistan makhan@awkum.edu.pk 2 Biotechnology Lab. Agricultural Research Institute (ARI), Tarnab, Peshawar, Pakistan

Publication info: Current Biotechnology 8.2 (2019): 70-84.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 63 of 266

Evaluation of the behavior of phenolic compounds and steviol glycosides of sonicated strawberry juice sweetened with stevia (*Stevia rebaudiana* Bertoni).

Author: Zlabur, J Š; Dobričević, N.; Brnčić, M.; Barba, F J; Lorenzo, J M; Franco, D; Atanasov, A G; Voc' a, S.; Brnčić, S. R.

Publication info: Molecules 24.7 (2019): 1202.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 64 of 266

Impact of cultivation factors in vitro on the growth and the biosynthesis of steviol glycosides in *Stevia rebaudiana* cell cultures.

Author: Bondarev, N; Reshetnyak, O; Bondareva, T; Il'in, M; Nosov, A

Publication info: Physiology and Molecular Biology of Plants 25.4 (2019): 1091-1096.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

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Stevioside protects against rhabdomyolysis-induced acute kidney injury through PPAR-γ agonism in rats

Author: Kaur, Tajpreet 1 ; Singh, Damanpreet 2 ; Singh, Amrit P 1 ; Pathak, Devendra 3 ; Arora, Saroj 4 ; Singh, Brahmjot 1 ; Kaur, Sarabjit 1 ; Singh, Balbir 1 1 Department of Pharmaceutical Sciences, Guru Nanak Dev University, Amritsar, India, India 2 Pharmacology and Toxicology Laboratory, CSIR-Institute of Himalayan Bioresource Technology, Palampur, India, India 3 Department of Veterinary Anatomy, Guru Angad Dev Veterinary and Animal Sciences University, Ludhiana, India, India 4 Department of Botanical and Environmental Sciences, Guru Nanak Dev University, Amritsar, India, India

Publication info: Drug development research (Jul 31, 2020).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 66 of 266

Genetic diversity and population structure of the sweet leaf herb, *Stevia rebaudiana* B., cultivated and landraces germplasm assessed by EST-SSRs genotyping and steviol glycosides phenotyping

Author: Cosson, Patrick; Hastoy, Cecile; Errazzu, Luis Ernesto; Budeguer, Carlos Jorge; Boutie, Philippe; Rolin, Dominique; Schurdi-Levraud, Valerie

Publication info: BMC Plant Biology 19.1 (Oct 21, 2019): Article No.: 436.

[ProQuest document link](#)

Databases: BIOSIS Previews® (1926 - current)

Document 67 of 266

Steviol Glycosides in Dentistry

Author: Frentzen, M.; Besrukow, P.; Ackermann, A.; Pierog, S.; Schiermeyer, B.; Winter, J.; Woelwer-Rieck, U.; Kraus, D.

Publication info: In Steviol Glycosides: Cultivation, Processing, Analysis and Applications in Food , 162-184. ROYAL SOC CHEMISTRY, (2019).

[ProQuest document link](#)

Databases: BIOSIS Previews® (1926 - current)

Document 68 of 266

Low-dose stevia (Rebaudioside A) consumption perturbs gut microbiota and the mesolimbic dopamine reward system

Author: Nettleton, Jodi E. 1 ; Klancic, Teja 1 ; Schick, Alana 2 ; Choo, Ashley C. 1 ; Shearer, Jane 3 ; Borgland, Stephanie L. 4 ; Chleilat, Faye 1 ; Mayengbam, Shyamchand 1 ; Reimer, Raylene A. 3 1 Faculty of Kinesiology, University of Calgary, 2500 University Drive NW, Calgary, AB, T2N 1N4, Canada jenettle@ucalgary.ca ; teja.klancic@ucalgary.ca ; acchoo@ucalgary.ca ; fatima.chleilat1@ucalgary.ca ; shyamchandsingh.maye@ucalgary.ca 2 International Microbiome Centre, Cumming School of Medicine, University of Calgary, 3300 Hospital Drive NW, Calgary, AB, T2N 4N1, Canada a.schick@ucalgary.ca 3 Faculty of Kinesiology, University of Calgary, 2500 University Drive NW, Calgary, AB, T2N 1N4, Canada, Department of Biochemistry and Molecular Biology, Cumming School of Medicine, University of Calgary, 3300 Hospital Drive NW, Calgary, AB, T2N 4N1, Canada jshearer@ucalgary.ca 4 Hotchkiss Brain Institute, University of Calgary, 3300 Hospital Drive NW, Calgary, AB, T2N 4N1, Canada slborgla@ucalgary.ca

Publication info: Nutrients 11.6 (Jun 2019).

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 69 of 266

Effect of Growth Regulators on *Stevia rebaudiana* Bertoni Callus Genesis and Influence of Auxin and Proline to Steviol Glycosides, Phenols, Flavonoids Accumulation, and Antioxidant Activity In Vitro

Author: Blinstrubienė, Aušra 1 ; Burbulis, Natalija 1 ; Juškevičiūtė, Neringa 1 ; Vaitkevičienė, Nijolė 1 ; Žūkienė, Rasa 1 1 Institute of Biology and Plant Biotechnology, Agriculture Academy, Vytautas Magnus University, Donelaicio str. 58, Kaunas, 44248, Lithuania

Publication info: Molecules (Basel, Switzerland) 25.12 (Jun 15, 2020).

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 70 of 266

Study on sweetener selection in RTD tea beverages.

Author: Fadlillah, H N; Ramadhan, H S; Hermanianto, J; Felanesa, L

Publication info: Jurnal Teknologi dan Industri Pangan 31.1 (2020): 1-8.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 71 of 266

Effect of pruning in biomass production and steviol glycosides contents of *Stevia rebaudiana* Bertoni var. *Morita* II

Author: Gonzalez, Marielys; Daquinta, Marcos; Pina, Danilo; Portal, Nayansi; Mosqueda, Osbel; Andujar, Ivan; Gonzalez, Luzgrey; Gonzalez, Susett; Perez, Lianny; Lezcano, Yarianne; Concepcion, Oscar; Escalona, Maritza

Publication info: *Biotechnologia Vegetal* 19.3 (Jul 2019 - Sep 2019): 155-164.

[ProQuest document link](#)

Databases: BIOSIS Previews® (1926 - current)

Document 72 of 266

Comparative transcriptome analysis revealed gamma-irradiation mediated disruption of floral integrator gene(s) leading to prolonged vegetative phase in *Stevia rebaudiana* Bertoni

Author: Singh, Gopal; Pal, Poonam; Masand, Mamta; Seth, Romit; Kumar, Ashok; Singh, Sanatsujat; Sharma, Ram Kumar

Publication info: *Plant Physiology and Biochemistry (Paris)* 148 (Mar 2020): 90-102.

[ProQuest document link](#)

Databases: BIOSIS Previews® (1926 - current)

Document 73 of 266

Effect of nutrient solutions and different planting bed on some growth characteristics of *Stevia rebaudiana* in inoculum conditions with mycorrhizal fungus.

Author: Seyedmohammadi, N S; Barmaki, M; Davari, M; Majd, K H

Publication info: *Applied Ecology and Environmental Research* 17.3 (2019): 5641-5648.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 74 of 266

Molecular docking studies and pharmacophore modeling of some insulin mimetic agents from herbal sources: A rational approach towards designing of orally active insulin mimetic agents

Author: Pradhan, Joohee 1 ; Panchawat, Sunita 1 1 Department of Pharmaceutical Sciences, Mohanlal Sukhadia University, Udaipur, Rajasthan, 313001, India Juhipradhan123@gmail.com

Publication info: *Current Traditional Medicine* 6.2 (2020): 121-133.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 75 of 266

Identification of GH1 gene family fgt members in *Stevia rebaudiana* and their expression when grown in darkness

Author: Yang, Yongheng 1 ; Zhang, Ting 1 ; Xu, Xiaoyang 1 ; Sun, Yuming 1 ; Zhang, Yongxia 1 ; Hou, Menglan 1 ; Huang, Suzhen 1 ; Yuan, Haiyan 1 ; Tong, Haiying 1 1 Institute of Botany, Jiangsu Province and Chinese Academy of Sciences, Nanjing, 210014, China, The Jiangsu Provincial Platform for Conservation and Utilization of Agricultural Germplasm, Nanjing, 210014, China, China yuanhaiyan416@163.com ; njtonghy@163.com

Publication info: *Molecular biology reports* (Oct 24, 2020).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 76 of 266

Reaction coupling separation for isosteviol production from stevioside catalyzed by acidic ion-exchange resin

Author: Hu, Xueyi 1 ; Zhou, Zhuoyu 1 ; Zhang, Zongying 1 ; Wang, Xiaoxia 1 ; Sui, Xiaochen 1 ; Chen, Junming 1 ; Xia, Yongmei 1 ; Zhang, Jue 2 ; Lin, Jianguo 2 1 State Key Laboratory of Food Science and Technology, School of Chemical and Materials Engineering, Jiangnan University, 1800 Lihu Avenue, Wuxi, 214122, Jiangsu, China, Key Laboratory of Synthetic and Biological Colloids (Ministry of Education), School of Chemical and Materials Engineering, Jiangnan University, 1800 Lihu Avenue, Wuxi, 214122, Jiangsu, China, China ymxia@jiangnan.edu.cn 2 Key Laboratory of Nuclear Medicine of Ministry of Health, Jiangsu Institute of Nuclear Medicine, Wuxi, 214063, Jiangsu, China, China

Publication info: Bioprocess and biosystems engineering (Aug 28, 2020).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 77 of 266

A bioeconomy perspective for natural sweetener Stevia.

Author: Ciriminna, R; Meneguzzo, F; Pecoraino, M; Pagliaro, M

Publication info: Biofuels, Bioproducts & Biorefining 13.3 (2019): 445-452.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

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Diterpenes and their derivatives as promising agents against dengue virus and dengue vectors: A literature-based review

Author: Islam, Muhammad Torequl 1 ; Mubarak, Mohammad S. 2 1 Department for Management of Science and Technology Development, Ton Duc Thang University, Ho Chi Minh City, Viet Nam, Faculty of Pharmacy, Ton Duc Thang University, Ho Chi Minh City, Viet Nam muhammad.torequl.islam@tdtu.edu.vn 2 Department of Chemistry, The University of Jordan, Amman, Jordan

Publication info: Phytotherapy Research 34.4 (Apr 1, 2020): 674-684.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 79 of 266

Response to the Letter to the Editor by S. Schiffman and H. Nagle: Revisiting the data and information that has collectively established the safety of low/no-calorie sweeteners, including sucralose

Author: Roberts, Ashley; Lobach, Alexandra R.

Publication info: Food and Chemical Toxicology 132 (Oct 2019): Article No.: 110691.

[ProQuest document link](#)

Databases: BIOSIS Previews® (1926 - current)

Document 80 of 266

Highly sweet compounds of plant origin: from ethnobotanical observations to wide utilization.: Special Issue: JEP 40 years.

Author: Soejarto, D D; Addo, E M; Kinghorn, A D

Publication info: Journal of Ethnopharmacology 243 (2019): 112056.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 81 of 266

Plant growth retardants (PGRs) affect growth and secondary metabolite biosynthesis in Stevia rebaudiana Bertoni under drought stress

Author: Karimi, M.; Ahmadi, A.; Hashemi, J.; Abbasi, A.; Tavarini, S.; Pompeiano, A.; Guglielminetti, L.; Angelini, L. G.

Publication info: South African Journal of Botany 121 (Mar 2019): 394-401.

[ProQuest document link](#)

Databases: BIOSIS Previews® (1926 - current)

Document 82 of 266

Effect of air drying on quality characteristics and mass transfer kinetics of osmotically dehydrated sea buckthorn by stevia

Author: Lentzou, D. 1 ; Templalexis, C. 1 ; Xanthopoulos, G. 1 1 Department of Natural Resources Management and Agricultural Engineering, Faculty of School of Environment and Agricultural Engineering, Agricultural University of Athens, 75 Iera Odos Str., Athens, 11855, Greece xanthopoulos@aua.gr

Publication info: Food Research 4.4 (Aug 2020): 1140-1150.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 83 of 266

Bactericidal activity and structural studies of the steviol derivative 17-hydroxy-16-hydroxyiminobayeran-19-oic acid

Author: Boreiko, Sheila 1 ; Machado, Agnes T.P. 1 ; Stiirmer, Júlio C. 2 ; Iulek, Jorge 1 ; Silva, Marcio 3 1 Department of Chemistry, State University of Ponta Grossa, Ponta Grossa, PR, Brazil 2 Department of Chemical Engineering, Federal Technological University of Paraná, Ponta Grossa, PR, Brazil 3 Department of Education, Federal Technological University of Paraná, Ponta Grossa, PR, Brazil marcios@utfpr.edu.br

Publication info: Current Bioactive Compounds 16.2 (2020): 96-101.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 84 of 266

Angiotensin-converting enzyme inhibiting ability of ethanol extracts, steviol glycosides and protein hydrolysates from stevia leaves

Author: Wang, Limin 1 ; Wu, Wenbiao 1 College of Food Science, Southwest University, No. 2 Tian Shengqiao, Beibei, Chongqing, China

Publication info: Food & function 10.12 (Dec 11, 2019): 7967-7972.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 85 of 266

Development of a new particulate 4- μ m adsorbent layer for ultrathin-layer chromatography (miniaturized chromatogram).

Author: Kirchert, S; Schulz, M; Oberle, M; Morlock, G E

Publication info: Journal of Chromatography, A 1587 (2019): 247-255.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 86 of 266

Controlled elicitation increases steviol glycosides (SGs) content and gene expression-associated to biosynthesis of SGs in *Stevia rebaudiana* B. cv. Morita II.

Author: Vazquez-Hernandez, C; Feregrino-Perez, A A; Perez-Ramirez, I; Ocampo-Velazquez, R V; Rico-García, E; Torres-Pacheco, I; Guevara-Gonzalez, R G

Publication info: Industrial Crops and Products 139 (2019): 111479.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 87 of 266

Anti-inflammatory effect of stevioside abates Freund's complete adjuvant (FCA)-induced adjuvant arthritis in rats

Author: Alavala, Sateesh 1 ; Nalban, Nasiruddin 1 ; Sangaraju, Rajendra 1 ; Kuncha, Madhusudana 1 ; Jerald, Mahesh Kumar 2 ; Kilari, Eswar Kumar 3 ; Sistla, Ramakrishna 1 1 Department of Applied Biology, CSIR-Indian Institute of Chemical Technology (IICT), Hyderabad, 500 007, India sistla@iict.res.in 2 Animal House Facility, CSIR-Centre for Cellular and Molecular Biology (CCMB), Hyderabad, 500 007, India 3 Department of Pharmacology, A.U. College of Pharmaceutical Sciences, Andhra University, Visakhapatnam, 530 033, India

Publication info: Inflammopharmacology 28.6 (Dec 1, 2020): 1579-1597.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 88 of 266

***Stevia rebaudiana* extract attenuate metabolic disorders in diabetic rats via modulation of glucose transport and antioxidant signaling pathways and aquaporin-2 expression in two extrahepatic tissues**

Author: Bayat, Elahe 1 ; Rahpeima, Zahra 2 ; Dastghaib, Sanaz 3 ; Gholizadeh, Fatemeh 1 ; Erfani, Mehran 1 ; Asadikaram, Gholamreza 2 ; Mokarram, Pooneh 4 1 Department of Biochemistry, School of Medicine, Shiraz University of Medical Sciences, Shiraz, Iran, Iran 2 Department of Biochemistry, School of Medicine, Kerman University of Medical Sciences, Kerman, Iran, Iran 3 Department of Biochemistry, School of Medicine, Shiraz University of Medical Sciences, Shiraz, Iran, Endocrinology and Metabolism Research Center, Shiraz University of Medical Sciences, Shiraz, Iran, Iran 4 Department of Biochemistry, School of Medicine, Shiraz University of Medical Sciences, Shiraz, Iran, Autophagy Research Center, Shiraz University of Medical Sciences, Shiraz, Iran, Iran

Publication info: Journal of food biochemistry 44.8 (Aug 2020): e13252.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 89 of 266

Synthesis and characterization of stevioside having low degree polymerized glucosides using dextranucrase and dextranase

Author: Son, Gyumin 1 ; Nguyen, Thi Thanh Hanh 2 ; Park, Byeongsu 1 ; Kwak, Sohyung 1 ; Jin, Juhui 1 ; Kim, Young-Min 3 ; Moon, Young-Hwan 4 ; Park, Sunghee 5 ; Kim, Seong-Bo 5 ; Kim, Doman 6 1 Graduate School of International Agricultural Technology, Seoul National University, Pyeongchang-gun, Gangwon-do, 25354, South Korea, South Korea 2 Institute of Food Industrialization, Institutes of Green Bio Science & Technology, Seoul National University, Pyeongchang-gun, Gangwon-do, 25354, South Korea, South Korea 3 Department of Food Science & Technology, Chonnam National University, Gwangju, 61186, South Korea, South Korea 4 Audubon Sugar Institute, Louisiana State University Agricultural Center, Gabriel, LA, 70776, USA, USA 5 CJ CheilJedang, Life Ingredient & Material Research Institute, Suwon, 16495, South Korea, South Korea 6 Graduate School of International Agricultural Technology, Seoul National University, Pyeongchang-gun, Gangwon-do, 25354, South Korea; Institute of Food Industrialization, Institutes of Green Bio Science & Technology, Seoul National University, Pyeongchang-gun, Gangwon-do, 25354, South Korea, South Korea kimdm@snu.ac.kr

Publication info: Enzyme and microbial technology 132 (Jan 2020): 109412.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 90 of 266

Impact of Steviol Glycosides and Erythritol on the Human and Cebus apella Gut Microbiome

Author: Mahalak, Karley K 1 ; Firrman, Jenni 1 ; Tomasula, Peggy M 1 ; Nuñez, Alberto 2 ; Lee, Jung-Jin 3 ; Bittinger, Kyle 3 ; Rinaldi, William 4 ; Liu, Lin Shu 1 1 United States Department of Agriculture, Dairy and Functional Foods Research Unit, Agricultural Research Service, Eastern Regional Research Center, 600E Mermaid Lane, Wyndmoor, Pennsylvania 19038, United States, United States 2 United States Department of Agriculture, Agricultural Research Service, Eastern Regional Research Center, 600E Mermaid Lane, Wyndmoor, Pennsylvania 19038, United States, United States 3 Division of Gastroenterology, Hepatology, and Nutrition, Children's Hospital of Philadelphia, 3401 Civic Center Boulevard, Philadelphia, Pennsylvania 19104, United States, United States 4 Alpha Genesis Inc., 95 Castle Hall Road, Yemassee, South Carolina 29945, United States, United States

Publication info: Journal of agricultural and food chemistry (Jan 7, 2020).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 91 of 266

Stereoselective synthesis and antiproliferative activity of steviol-based diterpen aminodiols

Author: Ozsvár, Dániel 1 ; Nagy, Viktória 2 ; Zupkó, István 3 ; Szakonyi, Zsolt 4 1 Institute of Pharmaceutical Chemistry, University of Szeged, Interdisciplinary Excellent Center, Szeged, H-6720, Hungary daniel.ozsvar@pharm.u-szeged.hu 2 Department of Pharmacodynamics and Biopharmacy, University of Szeged, Szeged, H-6720, Hungary nagy.viktoria@pharm.u-szeged.hu 3 Department of Pharmacodynamics and Biopharmacy, University of Szeged, Szeged, H-6720, Hungary, Interdisciplinary Centre of Natural Products, University of Szeged, Szeged, H-6720, Hungary zupko@pharm.u-szeged.hu 4 Institute of Pharmaceutical Chemistry, University of Szeged, Interdisciplinary Excellent Center, Szeged, H-6720, Hungary, Interdisciplinary Centre of Natural Products, University of Szeged, Szeged, H-6720, Hungary szakonyi@pharm.u-szeged.hu

Publication info: International Journal of Molecular Sciences 21.1 (Jan 1, 2020).

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 92 of 266

Advances in enzymatic glycosylation of rubusoside

Author: Wang, Yao 1 ; Chen, Yi-Jun 1 ; Wu, Xu-Ri 1 1 Laboratory of Chemical Biology, School of Life Science and Technology, China Pharmaceutical University, Nanjing, 211198, China wangy94@yeah.net ; yjchen@cpu.edu.cn ; xuriwu@cpu.edu.cn

Publication info: Chinese Journal of Pharmaceutical Biotechnology 26.4 (Aug 1, 2019): 337-342.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 93 of 266

Stevia (*Stevia rebaudiana* Bertoni) responses to NaCl stress: growth, photosynthetic pigments, diterpene glycosides and ion content in root and shoot.

Author: Shahverdi, M A; Omid, H; Tabatabaei, S J

Publication info: Journal of the Saudi Society of Agricultural Sciences 18.4 (2019): 355-360.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 94 of 266

Stevia as a natural sweetener: A review

Author: Arumugam, Balakrishnan 1 ; Subramaniam, Arunambiga 1 ; Alagaraj, Praveena 1 1 Centre for Biological Sciences, Department of Biochemistry, K.S. Ramgasamy College of Arts and Science (Autonomous), Namakkal, Tiruchengode, Tamilnadu, 637215, India biosciencebala@gmail.com

Publication info: Cardiovascular and Hematological Agents in Medicinal Chemistry 18.2 (2020): 94-103.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 95 of 266

Spray drying encapsulation of stevia extract with maltodextrin and evaluation of the physicochemical and functional properties of produced powders

Author: Zorzenon, Maria Rosa T 1 ; Formigoni, Maysa 1 ; da Silva, Sandra B 2 ; Hodas, Fabiane 3 ; Piovan, Silvano 4 ; Ciotta, Simone R 1 ; Jansen, Cler A 5 ; Dacome, Antonio S 3 ; Pilau, Eduardo J 6 ; Mareze-Costa, Cecília E 4 ; Milani, Paula G 3 ; Costa, Silvio C 3 1 Postgraduate Program in Food Science, State University of Maringá (UEM), 5790, Colombo Avenue, Zip-code 87020-900, Maringá, Paraná, Brazil, Biochemistry Department, State University of Maringá (UEM), 5790, Colombo Avenue, Zip-code 87020-900, Maringá, Paraná, Brazil, Brazil 2 Postgraduate Program in Food Science, State University of Maringá (UEM), 5790, Colombo Avenue, Zip-code 87020-900, Maringá, Paraná, Brazil, Brazil 3 Biochemistry Department, State University of Maringá (UEM), 5790, Colombo Avenue, Zip-code 87020-900, Maringá, Paraná, Brazil, Brazil 4 Physiological Sciences Department, State University of Maringá (UEM), 5790, Colombo Avenue, Zip-code 87020-900, Maringá, Paraná, Brazil, Brazil 5 Laboratory of Biomolecules and Mass Spectrometry, Chemistry Department, State University of Maringá (UEM), 5790, Colombo Avenue, Zip-code 87020-900, Maringá, Paraná, Brazil, Postgraduate Program in Cell Biology, State University of Maringá (UEM), 5790, Colombo Avenue, Zip-code 87020-900, Maringá, Paraná, Brazil, Brazil 6 Laboratory of Biomolecules and Mass Spectrometry, Chemistry Department, State University of Maringá (UEM), 5790, Colombo Avenue, Zip-code 87020-900, Maringá, Paraná, Brazil, Brazil

Publication info: Journal of food science 85.10 (Oct 2020): 3590-3600.

[ProQuest document link](#) **Databases:** MEDLINE® (1946 - current)

Document 96 of 266

The first total synthesis of rebaudioside R

Author: Wen, Guo-En 1 ; Qiao, Zhi; Liu, Hui; Zeng, Zhi-Yong; Tu, Yuan-Hong; Xia, Jian-Hui; Zhang, Qing-Ju; Sun, Jian-Song 1 National Research Centre for Carbohydrate Synthesis, Jiangxi Normal University, Nanchang, 330022, China

Publication info: Organic & biomolecular chemistry 18.1 (Dec 18, 2019): 108-126.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 97 of 266

Engineered ZnO and CuO Nanoparticles Ameliorate Morphological and Biochemical Response in Tissue Culture Regenerants of Candyleaf (Stevia rebaudiana)

Author: Ahmad, Muhammad Arslan 1 ; Javed, Rabia 2 ; Adeel, Muhammad 3 ; Rizwan, Muhammad 4 ; Ao, Qiang 2 ; Yang, Yuesuo 5 1 Key Lab of Eco-restoration of Regional Contaminated Environment (Shenyang University), Ministry of Education, Shenyang, 110044, China, Department of Tissue Engineering, China Medical University, Shenyang, 110122, China 2 Department of Tissue Engineering, China Medical University, Shenyang, 110122, China 3 Beijing Key Laboratory of Farmland Soil Pollution Prevention and Remediation, College of Resources and Environmental Sciences, China Agricultural University, Beijing, 100193, China 4 Microelement research center, College of Resources and Environment, Huazhong Agricultural University, Wuhan, 430070, China 5 Key Lab of Eco-restoration of Regional Contaminated Environment (Shenyang University), Ministry of Education, Shenyang, 110044, China, Key Lab of Groundwater and Environment (Jilin University), Ministry of Education, Changchun, 130021, China

Publication info: Molecules (Basel, Switzerland) 25.6 (Mar 17, 2020).

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 98 of 266

Dietary stevioside supplementation increases feed intake by altering the hypothalamic transcriptome profile and gut microbiota in broiler chickens

Author: Jiang, Jingle 1 ; Qi, Lina 1 ; Lv, Zengpeng 1 ; Wei, Quanwei 1 ; Shi, Fangxiong 1 1 College of Animal Science and Technology, Nanjing Agricultural University, Nanjing, China, China

Publication info: Journal of the science of food and agriculture (Sep 27, 2020).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 99 of 266

The effect of the elicitors on the steviol glycosides biosynthesis pathway in Stevia rebaudiana

Author: Tavakoli, Hourieh 1 ; Tavakoli, Nasibeh 1 ; Moradi, Foad 2 1 Department of Agronomy and Plant Breeding, Faculty of Agriculture and Natural Resource, University of Mohaghegh Ardabili, Agricultural Biotechnology Research Institute of Iran (ABRII), Karaj, Iran 2 Agricultural Biotechnology Research Institute of Iran (ABRII), Agricultural Research Education and Extension Organisation (AREEO), Karaj, Iran; and Corresponding author. Email: fmoradi@abrii.ac.ir

Publication info: Functional plant biology : FPB 46.9 (Aug 1, 2019): 787-795.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 100 of 266

Antifungal Activity against Fusarium culmorum of Stevioside, Silybum marianum Seed Extracts, and Their Conjugate Complexes

Author: Buzon-Duran, Laura; Martin-Gil, Jesus; del Carmen Ramos-Sanchez, Maria; Perez-Lebena, Eduardo; Luis Marcos-Robles, Jose; Fombellida-Villafuela, Angel; Martin-Ramos, Pablo

Publication info: Antibiotics-Basel 9.8 (Aug 2020): Article No.: 440.

[ProQuest document link](#)

Databases: BIOSIS® Toxicology (1969 - current)

Document 101 of 266

Redesign and reconstruction of a steviol-biosynthetic pathway for enhanced production of steviol in Escherichia coli

Author: Moon, Jun Ho 1 ; Lee, Kunjoong 1 ; Lee, Jun Ho 1 ; Lee, Pyung Cheon 1 1 Department of Molecular Science and Technology and Department of Applied Chemistry and Biological Engineering, Ajou University, Woncheon-dong, Yeongtong-gu, Suwon, 16499, South Korea, South Korea pcleee@ajou.ac.kr

Publication info: Microbial cell factories 19.1 (Feb 3, 2020): 20.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 102 of 266

Effect of dietary stevioside supplementation on growth performance, nutrient digestibility, serum parameters, and intestinal microflora in broilers

Author: Wu, Xuezhuang 1 ; Yang, Peilong; Sifa, Dai; Wen, Zhiguo 1 College of Animal Science, Anhui Science and Technology University, Bengbu, 233100, China wuxuezhuang@126.com

Publication info: Food & function 10.5 (May 22, 2019): 2340-2346.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 103 of 266

Volatile organic compounds of tobacco leaves versus waste (scrap, dust, and midrib): extraction and optimization

Author: Banožić, Marija 1 ; Aladić, Krunoslav 1 ; Jerković, Igor 2 ; Jokić, Stela 1 1 Faculty of Food Technology Osijek, Josip Juraj Strossmayer University of Osijek, Osijek, Croatia, Croatia 2 Faculty of Chemistry and Technology, University of Split, Split, Croatia, Croatia

Publication info: Journal of the science of food and agriculture (Sep 7, 2020).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 104 of 266

Preparing molecularly imprinted nanoparticles of saponins via cooperative imprinting strategy

Author: Zhang, Hao 1 ; Zhang, Yanan 2 ; Wang, Hailing 2 ; Wen, Han 2 ; Yan, Zhifeng 3 ; Huang, Ailan 3 ; Bie, Zijun 4 ; Chen, Yang 4 1 School of Pharmacy, Bengbu Medical University, Bengbu, P. R. China, Department of Pharmacy, First Affiliated Hospital of Bengbu Medical University, Bengbu, P. R. China, P. R. China 2 School of Pharmacy, Bengbu Medical University, Bengbu, P. R. China, P. R. China 3 Department of Chemistry, Bengbu Medical University, Bengbu, P. R. China, P. R. China 4 School of Pharmacy, Bengbu Medical University, Bengbu, P. R. China, Department of Chemistry, Bengbu Medical University, Bengbu, P. R. China, P. R. China

Publication info: Journal of separation science 43.11 (Jun 2020): 2162-2171.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 105 of 266

Adjusting the chromatographic properties of poly(ionic liquid)-modified stationary phases by substitution on the imidazolium cation

Author: Wang, Jie 1 ; Tang, Yuqi 1 ; Chu, Huiyuan 1 ; Shen, Jiwei 1 ; Wang, Chaozhan 1 ; Wei, Yinmao 1 1 Key Laboratory of Synthetic and Natural Functional Molecule Chemistry of Ministry of Education, College of Chemistry and Materials Science, Northwest University, Xi'an, P. R. China, P. R. China

Publication info: Journal of separation science (Apr 19, 2020).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

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Detecting kinsenoside from *Anoectochilus roxburghii* by HPLC-ELSD with dual columns of NH₂ and AQ-C18

Author: Wei, Mi 1 ; Chen, Xuemin 2 ; Yi, Liwen 3 ; Yuan, Yuanyuan 3 ; Zhang, Hua 3 ; Fu, Chunhua 3 ; Yu, Longjiang 3 1 Institute of Resource Biology and Biotechnology, Department of Biotechnology, College of Life Science and Technology, Huazhong University of Science and Technology, Wuhan, 430074, China, Key Laboratory for Quality Control of Characteristic Fruits and Vegetables of Hubei Province, College of Life Science and Technology, Hubei Engineering University, Xiaogan, 432000, China, China 2 Institute of Resource Biology and Biotechnology, Department of Biotechnology, College of Life Science and Technology, Huazhong University of Science and Technology, Wuhan, 430074, China, China 3 Institute of Resource Biology and Biotechnology, Department of Biotechnology, College of Life Science and Technology, Huazhong University of Science and Technology, Wuhan, 430074, China, Hubei Engineering Research Center for both Edible and Medicinal Resources, Wuhan, 430074, China, China

Publication info: Phytochemical analysis : PCA (Mar 26, 2020).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

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Stevioside, a diterpenoid glycoside, shows anti-inflammatory property against Dextran Sulphate Sodium-induced ulcerative colitis in mice

Author: Alavala, Sateesh 1 ; Sangaraju, Rajendra 1 ; Nalban, Nasiruddin 1 ; Sahu, Bidya Dhar 1 ; Jerald, Mahesh Kumar 2 ; Kilari, Eswar Kumar 3 ; Sistla, Ramakrishna 1 1 Department of Applied Biology, CSIR-Indian Institute of Chemical Technology (IICT), Hyderabad, 500 007, India sistla@iict.res.in 2 Animal House Facility, CSIR-Centre for Cellular and Molecular Biology(CCMB), Hyderabad, 500 007, India 3 Department of Pharmacology, A.U College of Pharmaceutical Sciences, Andhra University, Visakhapatnam, 530 033, India

Publication info: European Journal of Pharmacology 855 (Jul 15, 2019): 192-201.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 108 of 266

Evaluation of cardioprotection and bio-efficacy enhancement of stevioside and diltiazem in rats

Author: Bhatt, Laxit 1 ; Amrutia, Jay 2 ; Chakraborty, Manodeep 2 ; Kamath, Jagadish 2 1 Department of Pharmacology and Toxicology, Zydus Research Centre, Ahmedabad, India, Department of Pharmacology, Shree Devi College of Pharmacy, Mangalore, India bkltox@gmail.com 2 Department of Pharmacology, Shree Devi College of Pharmacy, Mangalore, India jayamrutia@gmail.com ; Manodeep.chakraborty@gmail.com ; sdcpharm@rediffmail.com

Publication info: Future Journal of Pharmaceutical Sciences 6.1 (Dec 1, 2020).

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 109 of 266

Modeling and optimization of ultrasound-assisted green extraction and rapid HPTLC analysis of stevioside from *Stevia Rebaudiana*.

Author: Rouhani, M

Publication info: Industrial Crops and Products 132 (2019): 226-235.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 110 of 266

Antioxidant and immunomodulatory activity induced by stevioside in liver damage: *in vivo*, *in vitro* and *in silico* assays.

Author: Casas-Grajales, S; Ramos-Tovar, E; Chávez-Estrada, E; Alvarez-Suarez, D; Hernández-Aquino, E; Reyes-Gordillo, K; Cerda-García-Rojas, C M; Camacho, J; Tsutsumi, V; Lakshman, M R; Muriel, P

Publication info: Life Sciences 224 (2019): 187-196.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 111 of 266

Modeling of aerosol coating of sugar crystals based on study of physical and chemical properties of stevioside solutions.

Author: Petrov, S M; Podgornova, N M; Petrov, K S; Ryazhskih, V I

Publication info: Journal of Food Engineering 255 (2019): 61-68.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 112 of 266

In vitro evaluation of the cariogenic potential of rebaudioside A compared to sucrose and xylitol

Author: Ganter, Johanna 1 ; Hellwig, Elmar 2 ; Doerken, Sam 3 ; Al-Ahmad, Ali 2 1 Department of Operative Dentistry and Periodontology, Center for Dental Medicine, Faculty of Medicine, Medical Center-University of Freiburg, University of Freiburg, Freiburg, Germany, Department of Orthodontics, Center for Dental Medicine, Faculty of Medicine, Medical Center-University of Freiburg, University of Freiburg, Hugstetter Str. 55, Freiburg, 79106, Germany johanna.ganter@uniklinik-freiburg.de 2 Department of Operative Dentistry and Periodontology, Center for Dental Medicine, Faculty of Medicine, Medical Center-University of Freiburg, University of Freiburg, Freiburg, Germany 3 Institute of Medical Biometry and Statistics, Faculty of Medicine, Medical Center-University of Freiburg, University of Freiburg, Freiburg, Germany

Publication info: Clinical oral investigations 24.1 (Jan 1, 2020): 113-122.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 113 of 266

Antifungal activity against fusarium culmorum of stevioside, silybum marianum seed extracts, and their conjugate complexes

Author: Buzón-Durán, Laura 1 ; Martín-Gil, Jesús 1 ; Ramos-Sánchez, María Del Carmen 2 ; Pérez-Lebeña, Eduardo 1 ; Marcos-Robles, José Luis 1 ; Fombellida-Villafruela, Ángel 1 ; Martín-Ramos, Pablo 3 1 ETSIIAA, Universidad de Valladolid, Avenida de Madrid 44, Palencia, 34004, Spain laura.buzon@uva.es ; mgil@iaf.uva.es ; eplebena@gmail.com ; jlmarcos@iaf.uva.es ; afv@pvs.uva.es 2 Servicio de Microbiología y Parasitología, Hospital Universitario Río Hortega, SACYL, Calle Dulzaina, 2, Valladolid, 47012, Spain mramoss@saludcastillayleon.es 3 Instituto Universitario de Investigación en Ciencias Ambientales de Aragón (IUCA), EPS, Universidad de Zaragoza, Carretera de Cuarte, s/n, Huesca, 22071, Spain pmr@unizar.es

Publication info: Antibiotics 9.8 (Aug 2020): 1-14.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

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Development of Bacillus safensis-based liquid bioformulation to augment growth, stevioside content, and nutrient uptake in Stevia rebaudiana

Author: Prakash, Jai 1 ; Arora, Naveen Kumar 2 1 Department of Environmental Microbiology (DEM), School for Environmental Sciences (SES), Babasaheb Bhimrao Ambedkar University (A Central University), Vidya Vihar ,Raebareli Road, Lucknow, Uttar Pradesh, 226025, India 2 Department of Environmental Science (DES), School for Environmental Sciences (SES), Babasaheb Bhimrao Ambedkar University (A Central University), Vidya Vihar ,Raebareli Road, Lucknow, Uttar Pradesh, 226025, India nkarora_net@rediffmail.com

Publication info: World journal of microbiology & biotechnology 36.1 (Dec 19, 2019): 8.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

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HI-HPTLC-UV/Vis/FLD-HESI-HRMS and bioprofiling of steviol glycosides, steviol, and isosteviol in Stevia leaves and foods

Author: Morlock, Gertrud E. 1 ; Heil, Julia 1 1 Institute of Nutritional Science, Chair of Food Science, TransMIT Center for Effect-Directed Analysis, Justus Liebig University Giessen, Heinrich-Buff-Ring 26-32, Giessen, 35392, Germany Gertrud.Morlock@uni-giessen.de

Publication info: Analytical and bioanalytical chemistry 412.24 (Sep 1, 2020): 6431-6448.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 116 of 266

Ceramide–rubusoside nanomicelles, a potential therapeutic approach to target cancers carrying p53 missense mutations

Author: Khiste, Sachin K. 1 ; Liu, Zhijun 2 ; Roy, Kartik R. 1 ; Uddin, Mohammad B. 1 ; Hosain, Salman B. 1 ; Gu, Xin 3 ; Nazzal, Sami 4 ; Hill, Ronald A. 1 ; Liu, Yong-Yu 1 1 School of Basic Pharmaceutical and Toxicological Sciences, College of Pharmacy, University of Louisiana at Monroe, Monroe, LA, United States yliu@ulm.edu 2 School of Renewable Natural Resources, Louisiana State University Agricultural Center, Baton Rouge, LA, United States 3 Department of Pathology, Louisiana State University Health Sciences Center, Shreveport, LA, United States 4 Department of Pharmaceutical Sciences, Texas Tech University Health Science Center, Dallas, TX, United States

Publication info: Molecular Cancer Therapeutics 19.2 (Feb 2020): 564-574.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

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The effects of the addition of starter cultures and stevioside on technological low-fat fermented sherbet ice-cream without sugar

Author: Arseneva, Tamara Pavlovna 1 ; Evstigneeva, Tatiana Nikolaevna 1 ; Iakovchenko, Natalia Vladimirovna 1 ; Vitalevna, Lugova Margarita 1 ; Kurganova, Ekaterina Vladimirovna 1 1 Department of Applied Biotechnology, ITMO University. Saint Petersburg, Russian Federation

Publication info: Acta scientiarum polonorum. Technologia alimentaria 18.4 (Oct 1, 2019): 361-371.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 118 of 266

Rubusoside-assisted solubilization of poorly soluble C₆-Ceramide for a pilot pharmacokinetic study

Author: Chen, Jianzhong 1 ; Khiste, Sachin K 2 ; Fu, Xiaomei 3 ; Roy, Kartik R. 2 ; Dong, Yixuan 1 ; Zhang, Jian 4 ; Liu, Mei 5 ; Liu, Yong-Yu 2 ; Liu, Zhijun 1 1 School of Renewable Natural Resources, LSU Agricultural Center, Louisiana State University, Baton Rouge, 70803, United States zliu@agcenter.lsu.edu 2 Basic Pharmaceutical Sciences, School of Pharmacy, University of Louisiana at Monroe, Monroe, LA, 71203, United States 3 School of Renewable Natural Resources, LSU Agricultural Center, Louisiana State University, Baton Rouge, 70803, United States, School of Pharmacy, Jiangxi University of Traditional Chinese Medicine, Nanchang, 330004, China 4 School of Renewable Natural Resources, LSU Agricultural Center, Louisiana State University, Baton Rouge, 70803, United States, School of Perfume and Aroma Technology, Shanghai Institute of Technology, Shanghai, 201418, China 5 School of Renewable Natural Resources, LSU Agricultural Center, Louisiana State University, Baton Rouge, 70803, United States, School of Pharmacy, Guangzhou University of Traditional Chinese Medicine, Guangzhou, 330004, China

Publication info: Prostaglandins and Other Lipid Mediators 146 (Feb 2020).

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 119 of 266

Development of *Bacillus safensis*-based liquid bioformulation to augment growth, stevioside content, and nutrient uptake in *Stevia rebaudiana*.

Author: Prakash, Jai; Arora, N K

Publication info: World Journal of Microbiology & Biotechnology 36.1 (2020).

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 120 of 266

Metabolomics analysis of the protective effect of rubusoside on palmitic acid-induced lipotoxicity in INS-1 cells using UPLC-Q/TOF MS

Author: Zheng, Hua 1 ; Wu, Jinxia 2 ; Huang, Hong 3 ; Meng, Chunmei 1 ; Li, Weidong 1 ; Wei, Tianli 2 ; Su, Zhiheng 2 1 Life Sciences Institute, Guangxi Medical University, Nanning, 530021, China 2 Pharmaceutical College, Guangxi Medical University, Nanning, 530021, China suzhiheng@gxmu.edu.cn 3 First Affiliated Hospital, Guangxi Medical University, Nanning, 530021, China

Publication info: Molecular Omics 15.3 (2019): 222-232.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 121 of 266

Effect of methyl jasmonate and salicylic acid on in vitro growth, stevioside production, and oxidative defense system in *Stevia rebaudiana*.

Author: Moharramnejad, S; Azam, A T; Panahandeh, J; Dehghanian, Z; Ashraf, Muhammad

Publication info: Sugar Tech 21.6 (2019): 1031-1038.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 122 of 266

Computational and pharmacological evaluation of stevioside derivatives for antinociceptive and anti-inflammatory potential

Author: Ahmad, Sadaf 1 ; Khan, Arifullah 1 ; Hossain, Mohammad Akbar 2 ; Ullah, Asad 3 ; Faheem, Muhammad 1 ; Shahid, Muhammad 4 ; Ahmad, Shabir 3 1 Riphah Institute of Pharmaceutical Sciences, Riphah International University, Islamabad, Pakistan mohammad.faheem@riphah.edu.pk 2 Department of Pharmacology and Toxicology, College of Pharmacy, Umm Al-Qura University, Makkah al Mukarramah, Saudi Arabia 3 Department of Pharmacology and Toxicology, College of Medicine, Umm Al-Qura University, Makkah, Saudi Arabia 4 Department of Clinical Pharmacy, College of Pharmacy, Prince Sattam Bin Abdulaziz University, Alkharj, Saudi Arabia

Publication info: Tropical Journal of Pharmaceutical Research 19.8 (Aug 2020): 1677-1684.

[ProQuest document link](#) **Databases:** Embase® (1947 - current)

Document 123 of 266

Efficacy of stevioside sweetener on pH of plaque among young adults

Author: Saira Siraj, E. 1 ; Pushpanjali, K. 2 ; Manoranjitha, B.S. 3 1 Department of Public Health Dentistry, Azeezia College of Dental Sciences and Research, Meeyannoor P.O, Kollam, Kerala, 691 537, India sairasirajsk@gmail.com 2 Department of Public Health Dentistry, Faculty of Dental Sciences, MSRUAS, Bengaluru, Karnataka, India 3 Public Health Dentistry, Moulaali, Hyderabad, Telangana, India

Publication info: Dental Research Journal 16.2 (Mar 2019 - Apr 2019): 104-109.

[ProQuest document link](#) **Databases:** Embase® (1947 - current)

Document 124 of 266

Nitrogen drives plant growth to the detriment of leaf sugar and steviol glycosides metabolisms in *Stevia (Stevia rebaudiana Bertonii)*

Author: Sun, Yuming 1 ; Hou, Menglan 1 ; Mur, Luis A J 2 ; Yang, Yongheng 1 ; Zhang, Ting 1 ; Xu, Xiaoyang 1 ; Huang, Suzhen 1 ; Tong, Haiying 1 1 Institute of Botany, Jiangsu Province and Chinese Academy of Sciences, Nanjing, 210014, China sunyumingagw@163.com ; njtonghy@163.com 2 Institute of Biological, Environmental and Rural Sciences, Aberystwyth University, Aberystwyth, SY23 3DA, United Kingdom

Publication info: Plant physiology and biochemistry : PPB 141 (Aug 1, 2019): 240-249.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 125 of 266

Novel nanomicelles based on rebaudioside A: A potential nanoplatform for oral delivery of honokiol with enhanced oral bioavailability and antitumor activity

Author: Wang, Jun 1 ; Yang, Hui 1 ; Li, Qiqi 1 ; Wu, Xianggen 1 ; Di, Guohu 2 ; Fan, Junting 3 ; Wei, Dongxu 4 ; Guo, Chuanlong 1
1 Department of Pharmacy, College of Chemical Engineering, Qingdao University of Science and Technology, Qingdao, 266042, China guochuanlong@qust.edu.cn 2 School of Basic Medicine, Qingdao University, Qingdao, 266071, China 3 Department of Pharmaceutical Analysis, School of Pharmacy, Nanjing Medical University, Nanjing, 211166, China 4 Department of Neonatal Intensive Care Unit, Qingdao Women and Children's Hospital, Qingdao, 266034, China
Publication info: International Journal of Pharmaceutics 590 (Nov 30, 2020).

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 126 of 266

Novel self-nanomicellizing solid dispersion based on rebaudioside A: A potential nanoplatform for oral delivery of curcumin

Author: Hou, Yuzhen 1 ; Wang, Hui 1 ; Zhang, Fan 1 ; Sun, Fengyuan 1 ; Xin, Meng 2 ; Li, Mengshuang 3 ; Li, Jun 4 ; Wu, Xianggen 1
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Publication info: International Journal of Nanomedicine 14 (2019): 557-571.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 127 of 266

Computational and pharmacological evaluation of stevioside derivatives for antinociceptive and antiinflammatory potential.

Author: Ahmad, Sadaf; Arif-Ullah Khan; Faheem, Muhammad; Iqbal, M S; Hossain, M A; Ullah, Asad; Ahmad, Shabir
Publication info: Tropical Journal of Pharmaceutical Research 19.8 (2020): 1677-1684.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 128 of 266

Inactivation of soybean Bowman-Birk inhibitor by stevioside: interaction studies and application to soymilk.

Author: Liu, Chun; Luo LiJuan; Wu, Ying; Yang, XiaoQuan; Dong Jie; Luo FeiJun; Zou Yuan; Shen YingBin; Lin QinLu
Publication info: Journal of Agricultural and Food Chemistry 67.8 (2019): 2255-2264.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 129 of 266

Hydrophobic recognition allows the glycosyltransferase UGT76G1 to catalyze its substrate in two orientations

Author: Yang, Ting 1 ; Zhang, Jinzhu 1 ; Ke, Dan 1 ; Yang, Wenxian 1 ; Tang, Minghai 2 ; Jiang, Jian 1 ; Cheng, Guo 3 ; Li, Jianshu 4 ; Cheng, Wei 2 ; Wei, Yuquan 2 ; Li, Qintong 5 ; Naismith, James H 6 ; Zhu, Xiaofeng 1 1 Key Laboratory of Bio-Resource and Eco-Environment of Ministry of Education, College of Life Sciences, Sichuan University; State Key Laboratory of Biotherapy and Cancer Center, West China Hospital, Sichuan University, 610064, Chengdu, China, China zhuxiaofeng@scu.edu.cn 2 State Key Laboratory of Biotherapy and Cancer Center, West China Hospital, Sichuan University, 610041, Chengdu, China, China 3 West China School of Public Health, Healthy Food Evaluation Research Center and State Key Laboratory of Biotherapy and Cancer Center, Sichuan University, 610041, Chengdu, China, China 4 Department of Biomedical Polymers and Artificial Organs, College of Polymer Science and Engineering and State Key Laboratory of Polymer Materials Engineering, Sichuan University, 610065, Chengdu, China, China 5 Department of Pediatrics, Obstetrics and Gynecology, West China Second University Hospital, Key Laboratory of Birth Defects and Related Diseases of Women and Children, Ministry of Education, Sichuan University, 610041, Chengdu, China, China liqintong@scu.edu.cn 6 State Key Laboratory of Biotherapy and Cancer Center, West China Hospital, Sichuan University, 610041, Chengdu, China, Division of Structural Biology, Wellcome Trust Centre of Human Genomics, Oxford, OX3 7BN, UK, Rosalind Franklin Institute, Harwell Campus, Didcot, OX11 0FA, UK, UK naismith@strubi.ox.ac.uk

Publication info: Nature communications 10.1 (Jul 19, 2019): 3214.

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Databases: MEDLINE® (1946 - current)

Document 130 of 266

Separation and purification of hydroxytyrosol and oleuropein from *Olea europaea* L. (olive) leaves using macroporous resins and a novel solvent system

Author: Liu, Baoqian 1 ; Liu, Jianfei 1 ; Huang, Dongdong 2 ; Pei, Dong 3 ; Di, Duolong 4 1 CAS Key Laboratory of Chemistry of Northwestern Plant Resources and Key Laboratory for Natural Medicine of Gansu Province, Lanzhou Institute of Chemical Physics, Chinese Academy of Sciences, Lanzhou, P. R. China, University of Chinese Academy of Sciences, 19A Yuquan Road, Beijing, 100049, P. R. China, P. R. China 2 CAS Key Laboratory of Chemistry of Northwestern Plant Resources and Key Laboratory for Natural Medicine of Gansu Province, Lanzhou Institute of Chemical Physics, Chinese Academy of Sciences, Lanzhou, P. R. China, College of Chemistry and Chemical Engineering, Northwest Normal University, Lanzhou, P. R. China, P. R. China 3 Centre of Resource Chemical and New Material, Lanzhou Institute of Chemical Physics, Chinese Academy of Sciences, Qingdao, P. R. China, P. R. China 4 CAS Key Laboratory of Chemistry of Northwestern Plant Resources and Key Laboratory for Natural Medicine of Gansu Province, Lanzhou Institute of Chemical Physics, Chinese Academy of Sciences, Lanzhou, P. R. China, University of Chinese Academy of Sciences, 19A Yuquan Road, Beijing, 100049, P. R. China, College of Chemistry and Chemical Engineering, Northwest Normal University, Lanzhou, P. R. China, P. R. China

Publication info: Journal of separation science 43.13 (Jul 2020): 2619-2625.

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Databases: MEDLINE® (1946 - current)

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Effect of Rubusoside, a Natural Sucrose Substitute, on Streptococcus mutans Biofilm Cariogenic Potential and Virulence Gene Expression In Vitro

Author: Guan, Chunru 1 ; Che, Faai 2 ; Zhou, Huoxiang 3 ; Li, Yiwei 2 ; Li, Yaru 2 ; Chu, Jinpu 2 1 Department of Stomatology, First Affiliated Hospital of Zhengzhou University, Zhengzhou, China, Academy of Medical Sciences, Zhengzhou University, Zhengzhou, China 2 Department of Stomatology, First Affiliated Hospital of Zhengzhou University, Zhengzhou, China 3 Laboratory of Microbiology and Immunology, Henan Institute of Medical and Pharmaceutical Sciences, Zhengzhou University, Zhengzhou, China, Marshall B. J. Medical Research Center, Fifth Affiliated Hospital of Zhengzhou University, Zhengzhou, China

Publication info: Applied and environmental microbiology 86.16 (Aug 3, 2020).

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 132 of 266

Beneficial effects of stevioside on AGEs, blood glucose, lipid profile and renal status in streptozotocin-induced diabetic rats

Author: Aswar, Urmila 1 ; Gogawale, Vinayak 2 ; Miniyar, Pankaj 2 ; Patil, Yugendra 3 1 Bharati Vidyapeeth (Deemed to be University), Poona College of Pharmacy, Erandwane, Pune, Maharashtra, India aswarurmila@gmail.com 2 STES's Sinhgad Institute of Pharmacy, Savitribai Phule Pune University, Narhe, Pune, Maharashtra, India 3 National Chemical Laboratory, Pune, Maharashtra, India

Publication info: Journal of Applied Biomedicine 17.3 (2019): 190-197.

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Databases: Embase® (1947 - current)

Document 133 of 266

Enhanced Heterologous Production of Glycosyltransferase UGT76G1 by Co-Expression of Endogenous prpD and malK in Escherichia coli and Its Transglycosylation Application in Production of Rebaudioside

Author: Shu, Wenju 1 ; Zheng, Hongchen 2 ; Fu, Xiaoping 3 ; Zhen, Jie 3 ; Tan, Ming 3 ; Xu, Jianyong 3 ; Zhao, Xingya 1 ; Yang, Shibin 1 ; Song, Hui 2 ; Ma, Yanhe 4 1 University of Chinese Academy of Sciences, Beijing 100049, China, Industrial Enzymes National Engineering Laboratory, Tianjin Institute of Industrial Biotechnology, Chinese Academy of Sciences, Tianjin 300308, China, China 2 University of Chinese Academy of Sciences, Beijing 100049, China, Industrial Enzymes National Engineering Laboratory, Tianjin Institute of Industrial Biotechnology, Chinese Academy of Sciences, Tianjin 300308, China, Tianjin Key Laboratory for Industrial Biological Systems and Bioprocessing Engineering, Tianjin Institute of Industrial Biotechnology, Chinese Academy of Sciences, Tianjin 300308, China, China 3 Industrial Enzymes National Engineering Laboratory, Tianjin Institute of Industrial Biotechnology, Chinese Academy of Sciences, Tianjin 300308, China, Tianjin Key Laboratory for Industrial Biological Systems and Bioprocessing Engineering, Tianjin Institute of Industrial Biotechnology, Chinese Academy of Sciences, Tianjin 300308, China, China 4 Industrial Enzymes National Engineering Laboratory, Tianjin Institute of Industrial Biotechnology, Chinese Academy of Sciences, Tianjin 300308, China, China

Publication info: International journal of molecular sciences 21.16 (Aug 11, 2020).

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Databases: MEDLINE® (1946 - current)

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¹H-NMR Metabolomics Analysis of the Effect of Rubusoside on Serum Metabolites of Golden Hamsters on a High-Fat Diet

Author: Li, Li 1 ; Jiang, Manjing 1 ; Li, Yaohua 1 ; Su, Jian 2 ; Qu, Xiaosheng 3 ; Fan, Lanlan 1 1 School of Pharmacy, Guangxi University of Chinese Medicine, Nanning 530001, China, China 2 Guangxi Scientific Research Center of Traditional Chinese Medicine, Guangxi University of Chinese Medicine, Nanning 530001, China, China 3 National Engineering Laboratory of Southwest Endangered Medicinal Resources Development, Guangxi Botanical Garden of Medicinal Plants, Nanning 530023, China, China

Publication info: Molecules (Basel, Switzerland) 25.6 (Mar 11, 2020).

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Databases: MEDLINE® (1946 - current)

Document 135 of 266

Enzymatic Monoglycosylation of Rubusoside and the Structure-Sweetness/Taste Relationship of Monoglycosyl Derivatives

Author: Zhao, Ling 1 ; Wang, Yao 1 ; Li, Zhenlin 2 ; Wang, Xiaonan 3 ; Chen, Yijun 1 ; Wu, Xuri 3 1 Laboratory of Chemical Biology, College of Life Sciences and Technology, China Pharmaceutical University, Jiangsu Province 211198, 639 Longmian Road, Nanjing, China 2 Department of Pharmaceutical Analysis and Metabolomics, Jiangsu Province Academy of Traditional Chinese Medicine, 100 Shizi St. Hongshan Rd. Nanjing, Jiangsu Province 210028, China 3 Department of Biochemistry, College of Life Sciences and Technology, China Pharmaceutical University, Jiangsu Province 211198, 639 Longmian Road, Nanjing, China

Publication info: Journal of agricultural and food chemistry 68.32 (Aug 12, 2020): 8702-8709.

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Databases: Embase® (1947 - current)

Document 136 of 266

Do steviol glycosides affect the oxidative and genotoxicity parameters in BALB/c mice?

Author: Yılmaz, Şemsi Gül 1 ; Uçar, Aslı 1 ; Yılmaz, Serkan 2 1 Faculty of Health Sciences, Department of Nutrition and Dietetics, Ankara University, Ankara, Turkey, Turkey 2 Faculty of Health Sciences, Ankara University, Ankara, Turkey, Turkey
Publication info: Drug and chemical toxicology (Jan 21, 2020): 1-6.

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Databases: MEDLINE® (1946 - current)

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Steviol glucuronide, a metabolite of steviol glycosides, potently stimulates insulin secretion from isolated mouse islets: Studies in vitro

Author: Gu, Wenqian 1 ; Rebsdorf, Andreas 1 ; Anker, Camilla 1 ; Gregersen, Søren 2 ; Hermansen, Kjeld 3 ; Geuns, Jan M C 4 ; Jeppesen, Per Bendix 1 1 Department of Clinical Medicine Aarhus University Aarhus N Denmark 2 Steno Diabetes Center Aarhus Aarhus University Hospital Aarhus N Denmark, Department of Endocrinology and Internal Medicine Aarhus University Hospital Aarhus N Denmark 3 Department of Clinical Medicine Aarhus University Aarhus N Denmark, Department of Endocrinology and Internal Medicine Aarhus University Hospital Aarhus N Denmark 4 Laboratory of Functional Biology KU Leuven Leuven Belgium

Publication info: Endocrinology, diabetes & metabolism 2.4 (Sep 6, 2019): e00093.

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Databases: MEDLINE® (1946 - current)

Document 138 of 266

Stevioside mediated chemosensitization studies and cytotoxicity assay on breast cancer cell lines MDA-MB-231 and SKBR3

Author: Khare, Noopur 1 ; Chandra, Sheela 1 1 Department of Bioengineering, BIT Mesra, Ranchi, India, India

Publication info: Saudi journal of biological sciences 26.7 (Nov 2019): 1596-1601.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

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ROLE OF MICROBIOME IN HEPATOPROTECTIVE EFFECT OF REBAUDIOSIDE FOR NONALCOHOLIC STEATOHEPATITIS

Author: Xi, Dong; Bhattacharjee, Jashdeep; Salazar-Gonzalez, Rosa-Maria; Warren, Mikako; Merritt, Russell; Kohli, Rohit

Publication info: Gastroenterology, suppl. S1 156.6 (2019): S-1294.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 140 of 266

Rebaudioside affords hepatoprotection ameliorating sugar sweetened beverage- induced nonalcoholic steatohepatitis

Author: Xi, Dong 1 ; Bhattacharjee, Jashdeep 1 ; Salazar-Gonzalez, Rosa-Maria 1 ; Park, Soyoun 2 ; Jang, Alice 2 ; Warren, Mikako 3 ; Merritt, Russell 1 ; Michail, Sonia 1 ; Bouret, Sebastien 2 ; Kohli, Rohit 1 1 Gastroenterology, Hepatology and Nutrition, 90027, Los Angeles, CA, USA rokohli@chla.usc.edu 2 Developmental Neuroscience Program & Diabetes and Obesity Program, Center for Endocrinology, Diabetes and Metabolism, VA, Richmond, 23298, United States 3 Pathology and Laboratory Medicine, Children's Hospital Los Angeles & University of Southern California Keck School of Medicine, Los Angeles, CA, 90027, USA

Publication info: Scientific reports 10.1 (Apr 21, 2020): 6689.

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Databases: Embase® (1947 - current)

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Elimination of bitter-off taste of stevioside through structure modification and computational interventions

Author: Bhardwaj, Vijay 1 ; Singh, Rahul 1 ; Singh, Pooja 2 ; Purohit, Rituraj 3 ; Kumar, Sanjay 4 1 Structural Bioinformatics Lab, CSIR-Institute of Himalayan Bioresource Technology (CSIR-IHBT), Palampur, HP 176061, India; Biotechnology division, CSIR-IHBT, Palampur, HP 176061, India, India 2 Pharmacoinformatics unit, National Institute of Pharmaceutical Education and Research, Hajipur, Bihar 844102, India, India 3 Structural Bioinformatics Lab, CSIR-Institute of Himalayan Bioresource Technology (CSIR-IHBT), Palampur, HP 176061, India; Biotechnology division, CSIR-IHBT, Palampur, HP 176061, India; Academy of Scientific & Innovative Research (AcSIR), CSIR-IHBT Campus, Palampur, HP 176061, India, India rituraj@ihbt.res.in 4 Biotechnology division, CSIR-IHBT, Palampur, HP 176061, India, India

Publication info: Journal of theoretical biology 486 (Feb 7, 2020): 110094.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

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Dietary Stevioside Supplementation Alleviates Lipopolysaccharide-Induced Intestinal Mucosal Damage through Anti-Inflammatory and Antioxidant Effects in Broiler Chickens

Author: Jiang, Jingle 1 ; Qi, Lina 1 ; Lv, Zengpeng 1 ; Jin, Song 2 ; Wei, Xihui 1 ; Shi, Fangxiong 1 1 National Experimental Teaching Demonstration Center of Animal Science, College of Animal Science and Technology, Nanjing Agricultural University, Nanjing 210095, China, China 2 Changzhou Animal Disease Control Center, Bureau of Agriculture and Rural Affairs of Changzhou, Changzhou 213003, China, China

Publication info: Antioxidants (Basel, Switzerland) 8.12 (Nov 21, 2019).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 143 of 266

Rubusoside alleviates the ovalbumin-induced mice allergic asthma by modulating the NF-κB activation

Author: Wang, Chengqiang 1 ; Tang, Jia 2 ; Qian, Bo 1 ; Zeng, Zhen 1 ; Gao, Yang 3 ; Song, Jia-Le 4 1 Department of Nutrition and Food Hygiene, School of Public Health, Guilin Medical University, Guilin, China 2 Institution of Documentation of Chinese Traditional Medicine Research, Nanjing University of Chinese Medicine, Nanjing, China 3 Department of Pharmacy, Northern Jiangsu People's Hospital, Yangzhou, China 4 Department of Nutrition and Food Hygiene, School of Public Health, Guilin Medical University, Guilin, China, Department of Surgery, School of Medicine, University of Maryland, MD, Baltimore, United States

Publication info: Journal of food biochemistry 44.5 (May 1, 2020): e13187.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 144 of 266

A beta-glucosidase gene from Stevia rebaudiana may be involved in the steviol glycosides catabolic pathway

Author: Yang, Yongheng 1 ; Hou, Menglan 1 ; Zhang, Ting 1 ; Sun, Yuming 1 ; Zhang, Yongxia 1 ; Huang, Suzhen 1 ; Xu, Xiaoyang 1 ; Yuan, Haiyan 1 1 Institute of Botany, Jiangsu Province and Chinese Academy of Sciences, Nanjing, 210014, China, The Jiangsu Provincial Platform for Conservation and Utilization of Agricultural Germplasm, Nanjing, 210014, China, China intergoogle@126.com ; yuanhaiyan416@163.com

Publication info: Molecular biology reports 47.5 (May 2020): 3577-3584.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 145 of 266

Biofunctionalization of magnetite nanoparticles with stevioside: effect on the size and thermal behaviour for use in hyperthermia applications

Author: Gupta, Ruby 1 ; Sharma, Deepika 1 1 a Institute of Nano Science and Technology , Mohali , Punjab , India, India

Publication info: International journal of hyperthermia : the official journal of European Society for Hyperthermic Oncology, North American Hyperthermia Group 36.1 (2019): 302-312.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 146 of 266

Heterologous expression of EUGT11 from *Oryza sativa* in *Pichia pastoris* for highly efficient one-pot production of rebaudioside D from rebaudioside A

Author: Wang, Zhenyang 1 ; Hong, Jiefang 2 ; Ma, Siyuan 3 ; Huang, Tong 3 ; Ma, Yuanyuan 4 ; Liu, Wei 5 ; Liu, Wenbin 5 ; Liu, Zhiming 6 ; Song, Hao 7 1 College of Material Science and Engineering, Northeast Forestry University, Harbin 150040, China; R&D Division, Sinochem Health Company Ltd., Qingdao 266071, China, China 2 Biomass Conversion Laboratory, Tianjin R&D Center for Petrochemical Technology, Tianjin University, Tianjin 300072, China, China 3 Biomass Conversion Laboratory, Tianjin R&D Center for Petrochemical Technology, Tianjin University, Tianjin 300072, China; School of Chemical Engineering and Technology, Tianjin University, Tianjin 300072, China, China 4 Biomass Conversion Laboratory, Tianjin R&D Center for Petrochemical Technology, Tianjin University, Tianjin 300072, China; Frontier Science Center for Synthetic Biology, Key Laboratory of Systems Bioengineering (Ministry of Education), Collaborative Innovation Center of Chemical Science and Engineering (Tianjin), Tianjin 300072, China; Frontier Technology Institute (Wuqing), Tianjin University, Tianjin 30072, China, China myy@tju.edu.cn 5 School of Chemical Engineering and Technology, Tianjin University, Tianjin 300072, China, China 6 College of Material Science and Engineering, Northeast Forestry University, Harbin 150040, China, China 7 School of Chemical Engineering and Technology, Tianjin University, Tianjin 300072, China; Frontier Science Center for Synthetic Biology, Key Laboratory of Systems Bioengineering (Ministry of Education), Collaborative Innovation Center of Chemical Science and Engineering (Tianjin), Tianjin 300072, China; Frontier Technology Institute (Wuqing), Tianjin University, Tianjin 30072, China, China hsong@tju.edu.cn

Publication info: International journal of biological macromolecules 163 (Nov 15, 2020): 1669-1676.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 147 of 266

Anti-cariogenic Characteristics of Rubusoside

Author: Kim, Jeessoo; Thi Thanh Hanh Nguyen; Jin, Juhui; Septiana, Iis; Son, Gyu-Min; Lee, Gang-Hee; Jung, You-Jin; Qureshi, Dilshad; Mok, Il Kyoon; Pal, Kunal; Yang, Soo-Yeon; Kim, Seong-Bo; Kim, Doman

Publication info: Biotechnology and Bioprocess Engineering 24.2 (Mar 2019): 282-287.

[ProQuest document link](#)

Databases: BIOSIS Previews® (1926 - current)

Document 148 of 266

Stevioside inhibits experimental fibrosis by down-regulating profibrotic Smad pathways and blocking hepatic stellate cell activation

Author: Casas-Grajales, Sael 1 ; Alvarez-Suarez, Diana 1 ; Ramos-Tovar, Erika 1 ; Dayana Buendía-Montaño, Laura 1 ; Reyes-Gordillo, Karina 2 ; Camacho, Javier 1 ; Tsutsumi, Víctor 3 ; Lakshman, M Raj 2 ; Muriel, Pablo 1 1 Department of Pharmacology, Cinvestav-IPN, Mexico City, Mexico, Mexico 2 Department of Biochemistry and Molecular Biology, School of Medicine and Health Science, The George Washington University Medical Center, Washington, District of Columbia, Lipid Research Laboratory, VA Medical Center, Washington, District of Columbia, District of Columbia 3 Department of Infectomics and Molecular Pathogenesis, Cinvestav-IPN, Mexico City, Mexico, Mexico

Publication info: Basic & clinical pharmacology & toxicology 124.6 (Jun 2019): 670-680.

[ProQuest document link](#)

Databases: ToxFile® (1900 - current)

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High affinity of 4-(4-(dimethylamino)styryl)-N-methylpyridinium transport for assessing organic cation drugs in hepatocellular carcinoma cells

Author: Jinakote, Metee 1 ; Ontawong, Atcharaporn 2 ; Soodvilai, Sunhapas 3 ; Pimta, Jeerawat 3 ; Pasachan, Tiphida 4 ; Chatsudthipong, Varanuj 3 ; Srimaroeng, Chutima 4 1 Faculty of Oriental Medicine, Chiang Rai College, Chiang Rai, 57000, Thailand, Department of Physiology, Faculty of Medicine, Chiang Mai University, Chiang Mai, 50200, Thailand, Thailand 2 Unit of Excellence of Coffee, Division of Physiology, School of Medical Sciences, University of Phayao, Phayao, 56000, Thailand, Thailand 3 Research Center of Transport Protein for Medical Innovation, Department of Physiology, Faculty of Science, Mahidol University, Bangkok, 10400, Thailand, Thailand 4 Department of Physiology, Faculty of Medicine, Chiang Mai University, Chiang Mai, 50200, Thailand, Thailand

Publication info: Fundamental & clinical pharmacology (Dec 27, 2019).

[ProQuest document link](#)

Databases: ToxFile® (1900 - current)

Document 150 of 266

Stevia residue extract alone and combination with allopurinol attenuate hyperuricemia in fructose-PO-induced hyperuricemic mice

Author: Mehmood, Arshad 1 ; Zhao, Lei 1 ; Wang, Chengtao 1 ; Hossen, Imam 1 ; Nadeem, Muhammad 2 1 Beijing Advance Innovation Center for Food Nutrition and Human Health, Beijing Technology and Business University, Beijing, China, Beijing Engineering and Technology Research Center of Food Additives, School of Food and Chemical Technology, Beijing Technology and Business University, Beijing, China, China 2 Institute of Food Science and Nutrition, University of Sargodha, Sargodha, Pakistan, Pakistan

Publication info: Journal of food biochemistry 44.1 (Jan 2020): e13087.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 151 of 266

Selective profiling of steviol-catalyzing UDP-glycosyltransferases with a metabolically synthesized probe

Author: Wong, Nai-Kei 1 ; Zhong, Suyun; Li, Weichao; Zhou, Fugui; Deng, Zhangshuang; Zhou, Yiqing 1 School of Biotechnology and Food Engineering, Changshu Institute of Technology, Changshu 215500, China, China zhyq2012@gmail.com

Publication info: Chemical communications (Cambridge, England) 56.82 (Oct 21, 2020): 12387-12390.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 152 of 266

Rebaudioside A administration prevents experimental liver fibrosis: an in vivo and in vitro study of the mechanisms of action involved

Author: Casas-Grajales, Sael 1 ; Reyes-Gordillo, Karina 2 ; Cerda-García-Rojas, Carlos M 3 ; Tsutsumi, Víctor 4 ; Lakshman, M Raj 2 ; Muriel, Pablo 1 1 Department of Pharmacology, Cinvestav-IPN, Av. Instituto Politécnico Nacional 2508, Col. San Pedro Zacatenco, 07360, Apartado Postal 14-740, Mexico City, Mexico, Mexico 2 Department of Biochemistry and Molecular Biology, School of Medicine and Health Science, The George Washington University Medical Center, 2300 I St. NW, Washington, DC, 20052, USA, Lipid Research Laboratory, VA Medical Center, 50 Irving St., Washington, DC, 20422, USA, USA 3 Department of Chemistry, Cinvestav-IPN, Av. Instituto Politécnico Nacional 2508, Col. San Pedro Zacatenco, 07360, Apartado Postal 14-740, Mexico City, Mexico, Mexico 4 Department of Infectomics and Molecular Pathogenesis, Cinvestav-IPN, Av. Instituto Politécnico Nacional 2508, Col. San Pedro Zacatenco, 07360, Apartado Postal 14-740, Mexico City, Mexico, Mexico

Publication info: Journal of applied toxicology : JAT 39.8 (Aug 2019): 1118-1131.

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Databases: MEDLINE® (1946 - current)

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Membrane separation processes for the extraction and purification of steviol glycosides: an overview

Author: Castro-Muñoz, Roberto 1 ; Díaz-Montes, Elsa 2 ; Cassano, Alfredo 3 ; Gontarek, Emilia 4 1 Tecnológico de Monterrey, Toluca de Lerdo, Campus Toluca, Mexico 2 Unidad Profesional Interdisciplinaria de Biotecnología, Instituto Politécnico Nacional, México City, Mexico 3 Institute on Membrane Technology, ITM-CNR, c/o University of Calabria, Rende, Italy 4 Faculty of Chemistry, Department of Process Engineering and Chemical Technology, Gdansk University of Technology, Gdansk, Poland

Publication info: Critical reviews in food science and nutrition (Jun 4, 2020): 1-23.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 154 of 266

The impact of steviol glycosides and erythritol on the human and Cebus apella gut microbiome

Author: Mahalak, Karley; Firrman, Jenni; Tomasula, Peggy M.; Nunez, Alberto; Lee, Jung-Jin; Bittinger, Kyle; Rinaldi, William; Liu, Linshu

Publication info: Journal of agricultural and food chemistry (Dec 23, 2019).

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 155 of 266

Glycaemic effects and plasma exposure of steviol administration in type 2 diabetic mice

Author: Simoens, C.; Wuyts, C.; Philippaert, K.; Beunen, K.; Khodaparast, L.; Gosciny, S.; Van Loco, J.; Van der Schueren, B.; Vennekens, R.

Publication info: Diabetologia 63.SUPPL 1, Suppl. 1, Sp. Iss. SI (Sep 2020): S234.

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Databases: BIOSIS Previews® (1926 - current)

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Erratum: Correction to RQ3, A Natural Rebaudioside D Isomer, Was Obtained from Glucosylation of Rebaudioside A Catalyzed by the CGTase Toruzyme 3.0 L (Journal of agricultural and food chemistry (2019) 67 28 (8020-8028))

Author: Guo, Qingbin; Zhang, Tongtong; Wang, Nifei; Xia, Yongmei; Zhou, Zhuoyu; Wang, Jian-Rong; Mei, Xuefeng

Publication info: Journal of agricultural and food chemistry 68.42 (Oct 21, 2020): 11875-11876.

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Databases: Embase® (1947 - current)

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Impact of Natural and Artificial Sweeteners Compounds in the Sensory Profile and Preference Drivers Applied to Traditional, Lactose-Free, and Vegan Frozen Desserts of Chocolate Flavor

Author: de Medeiros, Alessandra Cazelatto 1 ; Filho, Elson Rogério Tavares 1 ; Bolini, Helena Maria André 1 1 State Univ. of Campinas-FEA, Food and Nutrition Dept., Campinas, 13083-862, Brazil, Brazil

Publication info: Journal of food science 84.10 (Oct 2019): 2973-2982.

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Databases: MEDLINE® (1946 - current)

Document 158 of 266

Sensory Influence of Sweetener Addition on Traditional and Decaffeinated Espresso

Author: Cusiello, Kalinca Vitoria Cardoso 1 ; da Silva, Alessandra Cazelatto de Medeiros Lins 1 ; Tavares-Filho, Elson Rogerio 1 ; Bolini, Helena Maria André 1 1 Dept. of Food and Nutrition, School of Food Engineering, Univ. of Campinas, Campinas, Brazil
Publication info: Journal of food science 84.9 (Sep 1, 2019): 2628-2637.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

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A comparison of the effects of Stevia extract and metformin on metabolic syndrome indices in rats fed with a high-fat, high-sucrose diet

Author: Ranjbar, Tahereh 1 ; Nekooeian, Ali Akbar 2 ; Tanideh, Nader 3 ; Koohi-Hosseinabadi, Omid 4 ; Masoumi, Seyed Jalil 5 ; Amanat, Sasan 6 ; Azarpira, Negar 7 ; Monabati, Ahmad 8 1 Department of Clinical Nutrition, School of Nutrition and Food Science, Shiraz University of Medical Sciences, Shiraz, Iran 2 Department of Pharmacology, School of Medicine, Shiraz University of Medical Sciences, Shiraz, Iran 3 Department of Pharmacology, School of Medicine, Shiraz University of Medical Sciences, Shiraz, Iran, Stem Cells Technology Research Center, Shiraz University of Medical Sciences, Shiraz, Iran, Iran 4 Department of Central Research Laboratory, Shiraz University of Medical Sciences, Shiraz, Iran, Iran 5 Department of Clinical Nutrition, School of Nutrition and Food Science, Shiraz University of Medical Sciences, Shiraz, Iran, Nutrition Research Center, School of Nutrition and Food Sciences, Shiraz University of Medical Sciences, Shiraz, Iran, Iran 6 Student Research Committee, Larestan University of Medical Sciences, Larestan, Iran, Iran 7 Transplant Research Center, Shiraz University of Medical Sciences, Shiraz, Iran, Iran 8 Department of Pathology and Hematology Research Center, Shiraz University of Medical Sciences, Shiraz, Iran, Iran

Publication info: Journal of food biochemistry 44.8 (Aug 2020): e13242.

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Databases: MEDLINE® (1946 - current)

Document 160 of 266

Plausible biological interactions of low- and non-calorie sweeteners with the intestinal microbiota: An update of recent studies

Author: Plaza-Díaz, Julio 1 ; Pastor-Villaescusa, Belén 2 ; Rueda-Robles, Ascensión 3 ; Abadia-Molina, Francisco 4 ; Ruiz-Ojeda, Francisco Javier 5 1 Department of Biochemistry and Molecular Biology II, School of Pharmacy, University of Granada, Granada, 18071, Spain, Institute of Nutrition and Food Technology “José Mataix”, Center of Biomedical Research, University of Granada, Avda. del Conocimiento s/n, Armilla Granada, 18016, Spain, Instituto de Investigación Biosanitaria IBS.GRANADA, Complejo Hospitalario Universitario de Granada, Granada, 18014, Spain 2 Department of Biochemistry and Molecular Biology II, School of Pharmacy, University of Granada, Granada, 18071, Spain, LMU–Ludwig-Maximilians-University of Munich, Division of Metabolic and Nutritional Medicine, von Hauner Children’s Hospital, University of Munich Medical Center, Munich, 80337, Germany, Institute of Epidemiology, Helmholtz ZentrumMünchen–German Research Centre for Environmental Health, Neuherberg, 85764, Germany 3 Department of Biochemistry and Molecular Biology II, School of Pharmacy, University of Granada, Granada, 18071, Spain, Institute of Nutrition and Food Technology “José Mataix”, Center of Biomedical Research, University of Granada, Avda. del Conocimiento s/n, Armilla Granada, 18016, Spain 4 Institute of Nutrition and Food Technology “José Mataix”, Center of Biomedical Research, University of Granada, Avda. del Conocimiento s/n, Armilla Granada, 18016, Spain, Department of Cell Biology, School of Sciences, University of Granada, Granada, 18071, Spain 5 Department of Biochemistry and Molecular Biology II, School of Pharmacy, University of Granada, Granada, 18071, Spain, Instituto de Investigación Biosanitaria IBS.GRANADA, Complejo Hospitalario Universitario de Granada, Granada, 18014, Spain, RG Adipocytes and metabolism, Institute for Diabetes and Obesity, Helmholtz Diabetes Center at Helmholtz Center Munich, Neuherberg, Munich, 85764, Germany francisco.ruiz@helmholtz-muenchen.de

Publication info: Nutrients 12.4 (Apr 2020).

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Databases: Embase® (1947 - current)

Document 161 of 266

Consumption of non-nutritive sweeteners by pre-schoolers of the food and environment Chilean cohort (FECHIC) before the implementation of the Chilean food labelling and advertising law.

Author: Venegas Hargous, C.; Reyes, M; Taillie, L S; González, C G; Corvalán, C

Publication info: Nutrition Journal 19.69 (2020): (10 July 2020).

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

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Polyphenols from Stevia rebaudiana (Bertoni) leaves and their functional properties

Author: Myint, Khaing Zar 1 ; Wu, Ke 1 ; Xia, Yongmei 1 ; Fan, Ye 2 ; Shen, Jie 2 ; Zhang, Peter 3 ; Gu, Jianxin 4 1 State Key Laboratory of Food Science and Technology, Jiangnan Univ., 1800 Lihu Avenue, Wuxi, Jiangsu, 214122, China, Key Laboratory of Synthetic and Biological Colloids of Ministry of Education, School of Chemical and Materials Engineering, Jiangnan Univ., 1800 Lihu Avenue, Wuxi, Jiangsu, 214122, China, China 2 Key Laboratory of Synthetic and Biological Colloids of Ministry of Education, School of Chemical and Materials Engineering, Jiangnan Univ., 1800 Lihu Avenue, Wuxi, Jiangsu, 214122, China, China 3 Nascent Health Science LLC, 325 East 80th Street, 4E, New York, NY, 10075, U.S.A, U.S.A 4 Dept. of Biochemistry and Molecular Biology, School of Basic Medical Sciences, Fudan Univ., 130 Dong'an Road, Shanghai, 200032, China, China

Publication info: Journal of food science 85.2 (Feb 2020): 240-248.

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Databases: MEDLINE® (1946 - current)

Document 163 of 266

Divergent synthesis of complex diterpenes through a hybrid oxidative approach

Author: Zhang, Xiao 1 ; King-Smith, Emma 1 ; Dong, Liao-Bin 1 ; Yang, Li-Cheng 1 ; Rudol, Jeffrey D. 1 ; Shen, Ben 2 ; Renata, Hans 1 1 Department of Chemistry, Scripps Research Institute, Jupiter, FL, 33458, United States hrenata@scripps.edu 2 Department of Chemistry, Scripps Research Institute, Jupiter, FL, 33458, United States, Department of Molecular Medicine, Natural Products Discovery Center at Scripps Research, Jupiter, FL, 33458, United States

Publication info: Science 369.6505 (Aug 14, 2020): 799-806.

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Databases: Embase® (1947 - current)

Document 164 of 266

Protective and therapeutic effects of natural products against diabetes mellitus via regenerating pancreatic β -cells and restoring their dysfunction

Author: Semwal, Deepak Kumar 1 ; Kumar, Ankit 2 ; Aswal, Sonali 2 ; Chauhan, Ashutosh 3 ; Semwal, Ruchi Badoni 4 1 Department of Phytochemistry, Faculty of Biomedical Sciences, Uttarakhand Ayurved University, Dehradun, India, India 2 Research and Development Centre, Faculty of Biomedical Sciences, Uttarakhand Ayurved University, Dehradun, India, India 3 Department of Biotechnology, Faculty of Biomedical Sciences, Uttarakhand Ayurved University, Dehradun, India, India 4 Department of Chemistry, Pt. Lalit Mohan Sharma Government Postgraduate College, Rishikesh, India, India

Publication info: Phytotherapy research : PTR (Sep 28, 2020).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 165 of 266

Challenges in confectionery industry: Development and storage stability of innovative white tea-based candies

Author: Šeremet, Danijela 1 ; Mandura, Ana 1 ; Cebin, Aleksandra Vojvodić 1 ; Martinić, Arijana 1 ; Galić, Kata 1 ; Komes, Draženka 1 1 Faculty of Food Technology and Biotechnology, Department of Food Engineering, University of Zagreb, Pietrotti St 6, Zagreb, Croatia, Croatia

Publication info: Journal of food science 85.7 (Jul 2020): 2060-2068.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

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Use of mycorrhizal fungi releases the application of organic fertilizers to increase the production of leaf vitexin in yellow passion fruit

Author: de Oliveira, Paula Tarcila Félix 1 ; Dos Santos, Emanuela Lima 2 ; da Silva, Wliana Alves Viturino 3 ; Ferreira, Magda Rhayanny Assunção 3 ; Soares, Luiz Alberto Lira 3 ; da Silva, Francineyde Alves 4 ; da Silva, Fábio Sérgio Barbosa 2 1 Programa de Pós-Graduação em Biologia Celular e Molecular Aplicada, Instituto de Ciências Biológicas, Universidade de Pernambuco, Recife, Pernambuco, Brasil. Laboratório de Análises, Pesquisas e Estudos em Micorrizas, Universidade de Pernambuco, Recife, Brazil, Laboratório de Tecnologia Micorrízica, Universidade de Pernambuco, Petrolina, Brazil, Brazil 2 Programa de Pós-Graduação em Biologia Celular e Molecular Aplicada, Instituto de Ciências Biológicas, Universidade de Pernambuco, Recife, Pernambuco, Brasil. Laboratório de Análises, Pesquisas e Estudos em Micorrizas, Universidade de Pernambuco, Recife, Brazil, Brazil 3 Núcleo de Desenvolvimento Analítico e Tecnológico de Fitoterápicos, Universidade Federal de Pernambuco, Recife, Brazil, Brazil 4 Laboratório de Tecnologia Micorrízica, Universidade de Pernambuco, Petrolina, Brazil, Brazil

Publication info: Journal of the science of food and agriculture 100.4 (Mar 15, 2020): 1816-1821.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 167 of 266

In vitro cytotoxicity evaluation of steviosid on cancerous liver (Hep G2), colon (HT29), breast (MCF7) cells and normal kidney cell (Hek293) in comparison with cisplatin.

Author: Abolhasani, A; Heidari, F; Raminfar, R; Mousavi, S; Abolhasani, H

Publication info: Qom University of Medical Sciences Journal 14.3 (2020): fa27-fa34, en26.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 168 of 266

Sweet taste receptor agonists alter ovarian functions and ovarian cycles in aged mice

Author: Ngekure M X, Kavita 1 ; Jiang, Jingle 1 ; Enayatullah, Hamdard 1 ; Ennab, Wael 1 ; Mustafa, Sheeraz 1 ; Rodeni, Saif 1 ; Wei, Quanwei 1 ; Shi, Fangxiong 1 1 College of Animal Science and Technology, Nanjing Agricultural University, Nanjing, 210095, China fxshi@njau.edu.cn

Publication info: Reproductive biology 19.3 (Sep 1, 2019): 230-236.

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Databases: Embase® (1947 - current)

Document 169 of 266

Characterization of low calorie ready-to-serve peach beverage using natural sweetener, Stevia (Stevia rebaudiana Bertoni)

Author: Ahmad, Uswa 1 ; Ahmad, Rabia Shabir 2 ; Imran, Ali 2 ; Mushtaq, Zarina 2 ; Hussian, Syed Makhdoom 3 1

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Publication info: Progress in Nutrition 21 (2019): 435-444.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 170 of 266

Phytochemical and pharmacological importance of stevia: a calorie-free natural sweetener.: Special issue on green management for sustainable sugar industry.

Author: Singh, D P; Kumari, Meenakshi; Prakash, H G; Rao, G P; Solomon, S

Publication info: Sugar Tech 21.2 (2019): 227-234.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 171 of 266

Erythrocyte Partitioning Profile of Isosteviol in Human and Rat Blood

Author: Adehin, Ayorinde 1 ; Tan, Keai Sinn 2 ; Tan, Wen 2 1 Institute of Biomedical and Pharmaceutical Sciences, Guangdong University of Technology, Guangzhou, China, Department of Pharmaceutical Chemistry, Faculty of Pharmacy, Obafemi Awolowo University, Ile-Ife, Nigeria ayoadehin@outlook.com 2 Institute of Biomedical and Pharmaceutical Sciences, Guangdong University of Technology, Guangzhou, China went@gdut.edu.cn

Publication info: Current Therapeutic Research - Clinical and Experimental 91 (2019): 1-4.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 172 of 266

Rate-All-That-Apply (RATA) comparison of taste profiles for different sweeteners in black tea, chocolate milk, and natural yogurt

Author: Tan, Vicki Wei Kee 1 ; Wee, May Sui Mei 1 ; Tomic, Oliver 2 ; Forde, Ciarán G 3 1 Clinical Nutrition Research Center, A*STAR Singapore Inst. for Clinical Sciences, Medical Drive, Singapore, Singapore 2 Faculty of Science and Technology, Norwegian Univ. of Life Sciences, Ås, Norway, Norway 3 Clinical Nutrition Research Center, A*STAR Singapore Inst. for Clinical Sciences, Medical Drive, Singapore, Dept. of Physiology, Yong Loo Lin School of Medicine, Natl. Univ. of Singapore, Kent Ridge Road, Singapore, Singapore

Publication info: Journal of food science 85.2 (Feb 2020): 486-492.

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Databases: MEDLINE® (1946 - current)

Document 173 of 266

Tangential microfiltration of blackberry (*Rubus adenotrichus* Schltdl.) juice sweetened with stevia (*Stevia rebaudiana* Bertoni).

Author: González T, C; Vaillant B, F; Tapia G, M S

Publication info: Revista de Ciencias Agrícolas 36.2 (2019): 17-30.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 174 of 266

Effects of dietary sweeteners supplementation on growth performance, serum biochemicals, and jejunal physiological functions of broiler chickens

Author: Jiang, Jingle 1 ; Liu, Siyi 1 ; Jamal, Tuniyaz 1 ; Ding, Tengxin 1 ; Qi, Lina 1 ; Lv, Zengpeng 1 ; Yu, Debing 1 ; Shi, Fangxiong 1 1 National Experimental Teaching Demonstration Center of Animal Science, College of Animal Science and Technology, Nanjing Agricultural University, Nanjing, 210095, China fxshi@njau.edu.cn

Publication info: Poultry science 99.8 (Aug 1, 2020): 3948-3958.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 175 of 266

Plant-based antidiabetic nanoformulations: The emerging paradigm for effective therapy

Author: Dewanjee, Saikat 1 ; Chakraborty, Pratik 1 ; Mukherjee, Biswajit 2 ; De Feo, Vincenzo 3 1 Advanced Pharmacognosy Research Laboratory, Department of Pharmaceutical Technology, Jadavpur University, Kolkata, 700032, India saikat.dewanjee@jadavpuruniversity.in ; pratik.chakraborty88@yahoo.com 2 Pharmaceutics Research Laboratory, Department of Pharmaceutical Technology, Jadavpur University, Kolkata, 700032, India biswajit.mukherjee@jadavpuruniversity.in 3 Department of Pharmacy, University of Salerno, Fisciano, 84084, Italy defeo@unisa.it

Publication info: International Journal of Molecular Sciences 21.6 (Mar 2, 2020).

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 176 of 266

Biodegradability assessment of food additives using OECD 301F respirometric test

Author: Gatidou, Georgia 1 ; Vazaiou, Niki 1 ; Thomaidis, Nikolaos S. 2 ; Stasinakis, Athanasios S. 1 1 Department of Environment, Water and Air Quality Laboratory, University of the Aegean, University Hill, Mytilene, 81100, Greece ggatid@env.aegean.gr 2 Laboratory of Analytical Chemistry, Department of Chemistry, National and Kapodistrian University of Athens, Athens, 157 71, Greece

Publication info: Chemosphere 241 (Feb 2020).

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Databases: Embase® (1947 - current)

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Novel insights into the potential role of ion transport in sensory perception in Acanthamoeba

Author: Siddiqui, Ruqaiyyah 1 ; Roberts, Stephen K. 2 ; Ong, Timothy Yu Yee 3 ; Mungroo, Mohammad Ridwane 3 ; Anwar, Areeba 3 ; Khan, Naveed Ahmed 1 1 Department of Biology, Chemistry and Environmental Sciences, College of Arts and Sciences, American University of Sharjah, Sharjah, United Arab Emirates, Department of Biological Sciences, School of Science and Technology, Sunway University, Bandar Sunway, Malaysia ruqaiyyah_s@hotmail.com ; naveed5438@gmail.com 2 Biomedical and Life Sciences, Lancaster University, Lancaster, United Kingdom s.k.roberts@lancaster.ac.uk 3 Department of Biological Sciences, School of Science and Technology, Sunway University, Bandar Sunway, Malaysia timothy@sunway.edu.my ; ridwanemungroo@gmail.com ; areeba_anwar@ymail.com

Publication info: Parasites and Vectors 12.1 (Nov 14, 2019).

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 178 of 266

Sweeteners modulate bioactivity of endothelial progenitor cells but not induce detrimental effects both on inflammation and behavioural changes

Author: Schiano, Concetta 1 ; Grimaldi, Vincenzo 2 ; Boccella, Serena 3 ; Iannotta, Monica 3 ; Zullo, Alberto 4 ; Luongo, Livio 3 ; Mancini, Francesco Paolo 4 ; Maione, Sabatino 3 ; Napoli, Claudio 5 1 IRCCS SDN, Naples, Italy 2 Department of Medical, Surgical, Neurological, Metabolic and Geriatric Sciences, U.O.C. Immunohematology, Transfusion Medicine and Transplant Immunology, Azienda Ospedaliera Universitaria, Naples, Italy, Department of Sciences and Technologies, University of Sannio, Benevento, Italy 3 Department of Experimental Medicine, Section of Pharmacology 'L. Donatelli', Naples, Italy 4 Department of Sciences and Technologies, University of Sannio, Benevento, Italy 5 IRCCS SDN, Naples, Italy, Department of Medical, Surgical, Neurological, Metabolic and Geriatric Sciences, U.O.C. Immunohematology, Transfusion Medicine and Transplant Immunology, Azienda Ospedaliera Universitaria, Naples, Italy

Publication info: International journal of food sciences and nutrition 70.6 (Sep 1, 2019): 725-737.

[ProQuest document link](#) **Databases:** Embase® (1947 - current)

Document 179 of 266

Infection by the parasitic helminth *Trichinella spiralis* activates a Tas2r-mediated signaling pathway in intestinal tuft cells

Author: Luo, Xiao-Cui 1 ; Chen, Zhen-Huang 1 ; Xue, Jian-Bo 1 ; Zhao, Dong-Xiao 1 ; Lu, Chen 1 ; Li, Yi-Hong 1 ; Li, Song-Min 1 ; Du, Ya-Wen 1 ; Liu, Qun 1 ; Wang, Ping 2 ; Liu, Mingyuan 3 ; Huang, Liquan 4 1 College of Life Sciences, Zhejiang University, Hangzhou, 310058, China 2 Biosensor National Special Lab, Key Lab for Biomedical Engineering of Ministry of Education, Department of Biomedical Engineering, Zhejiang University, Hangzhou, 310027, China 3 Key Lab of Zoonosis Research, Institute of Zoonosis, Jilin University, Changchun, 130062, China, College of Veterinary Medicine, Jilin University, Changchun, 130062, China 4 College of Life Sciences, Zhejiang University, Hangzhou, 310058, China, Monell Chemical Senses Center, Philadelphia, PA, 19104, United States huangliquan@zju.edu.cn

Publication info: Proceedings of the National Academy of Sciences of the United States of America 116.12 (2019): 5564-5569.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

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Optimization of the silica-gel adsorption technique for the extraction of phytosterol glycosides from soybean lecithin powder using response surface methodology and artificial neural network models

Author: Kang, Jingjing 1 ; Cao, Dong 1 1 Natl. Engineering Laboratory for Food Science and Technology, Oil and Plant Protein Center, School of Food Science and Technology, Jiangnan Univ., 1800 Lihu Rd, Wuxi, Jiangsu Province, 214122, P. R. China, P. R. China

Publication info: Journal of food science 85.7 (Jul 2020): 1971-1982.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 181 of 266

Cross-modal effect of vanilla aroma on sweetness of different sweeteners among Chinese and Danish consumers.

Author: Bertelsen, A S; Zeng, Yan; Mielby, L A; Sun YuanXia; Byrne, D V; Kidmose, U

Publication info: Food Quality and Preference 87 (2021).

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 182 of 266

Anti-obesity effect of plant diterpenes and their derivatives: A review

Author: Islam, Muhammad Torequl 1 ; Ali, Eunos S. 2 ; Mubarak, Mohammad S. 3 1 Department for Management of Science and Technology Development, Ton Duc Thang University, Ho Chi Minh City, Viet Nam, Faculty of Pharmacy, Ton Duc Thang University, Ho Chi Minh City, Viet Nam muhammad.torequl.islam@tdtu.edu.vn 2 Gaco Pharmaceuticals and Research Laboratory, Dhaka, Bangladesh, College of Medicine and Public Health, Flinders University, Adelaide, SA, Australia 3 Department of Chemistry, The University of Jordan, Amman, Jordan

Publication info: Phytotherapy Research 34.6 (Jun 1, 2020): 1216-1225.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 183 of 266

Preparation technology and quality evaluation of *Schisandra chinensis* effervescent tablets.

Author: Qu ZhongYuan; Zhu ShiRu; Jiang, Xue; Zou Xiang; Wang, XinChen; Wu, Shuang; Li, WenLan

Publication info: Food Research and Development 41.6 (2020): 99-105.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 184 of 266

Development of grape polyphenol compound solid beverage.

Author: Li, JianHui; Lu, WenJing; Wang XiuJin; Liao ZhenYu

Publication info: Food Research and Development 41.17 (2020): 70-73.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 185 of 266

A randomized controlled trial contrasting the effects of 4 low-calorie sweeteners and sucrose on body weight in adults with overweight or obesity.

Author: Higgins, K A; Mattes, R D

Publication info: American Journal of Clinical Nutrition 109.5 (2019): 1288-1301.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 186 of 266

Rapid identification of some sweeteners and sugars by attenuated total reflectance-Fourier transform infrared (ATR-FTIR), near-infrared (NIR) and Raman spectroscopy.

Author: Guven, B; Velioglu, S D; Boyaci, I H

Publication info: GIDA - Journal of Food 44.2 (2019): 274-290.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 187 of 266

Glycoside-specific glycosyltransferases catalyze regio-selective sequential glucosylations for a sesame lignan, sesaminol trilucoside

Author: Ono, Eiichiro 1 ; Waki, Toshiyuki 2 ; Oikawa, Daiki 2 ; Murata, Jun 3 ; Shiraiishi, Akira 3 ; Toyonaga, Hiromi 1 ; Kato, Masako 4 ; Ogata, Naoki 4 ; Takahashi, Seiji 2 ; Yamaguchi, Masa-Atsu 5 ; Horikawa, Manabu 3 ; Nakayama, Toru 2 1 Suntory Global Innovation Center (SIC) Ltd., Research Institute, Soraku-gun, Kyoto, 619-0284, Japan, Japan 2 Graduate School of Engineering, Tohoku University, Sendai, Miyagi, 980-8579, Japan, Japan 3 Suntory Bioorganic Research Institute (SUNBOR), Suntory Foundation for Life Sciences, Soraku-gun, Kyoto, 619-0284, Japan, Japan 4 National Agriculture and Food Research Organization (NARO), Tsukuba, Ibaraki, 305-8517, Japan, Japan 5 Minami-Kyushu University, Miyakonojo, Miyazaki, 885-0035, Japan, Japan

Publication info: The Plant journal : for cell and molecular biology 101.5 (Mar 2020): 1221-1233.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 188 of 266

Diabetes and plant-derived natural products: From ethnopharmacological approaches to their potential for modern drug discovery and development

Author: Jugran, Arun K. 1 ; Rawat, Sandeep 2 ; Devkota, Hari P. 3 ; Bhatt, Indra D. 4 ; Rawal, Ranbeer S. 4 1 Garhwal Regional Centre, G. B. Pant National Institute of Himalayan Environment (NIHE), Srinagar, Uttarakhand, India arunjugran@gbpihed.nic.in 2 Sikkim Regional Centre, G. B. Pant National Institute of Himalayan Environment (NIHE), Gangtok, Sikkim, India sandeep_rawat15@rediffmail.com 3 Department of Instrumental Analysis, Graduate School of Pharmaceutical Sciences, Kumamoto University, Kumamoto, Japan devkotah@kumamoto-u.ac.jp 4 Center for Biodiversity Conservation and Management (CBCM), G. B. Pant National Institute of Himalayan Environment (NIHE), Kosi-Katarmal, Almora, Uttarakhand, India bhattid4@gmail.com ; ranbeerrawal4@gmail.com

Publication info: Phytotherapy Research (2020).

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 189 of 266

STAT signaling in polycystic kidney disease

Author: Strubl, Sebastian 1 ; Torres, Jacob A. 2 ; Spindt, Alison K. 2 ; Pellegrini, Hannah 2 ; Liebau, Max C. 3 ; Weimbs, Thomas 2 1 Department of Molecular, Cellular, and Developmental Biology, Neuroscience Research Institute, University of California Santa Barbara, Santa Barbara, CA, 93106-9625, United States, Department II of Internal Medicine, Faculty of Medicine and University Hospital Cologne, University of Cologne, Cologne, Germany 2 Department of Molecular, Cellular, and Developmental Biology, Neuroscience Research Institute, University of California Santa Barbara, Santa Barbara, CA, 93106-9625, United States weimbs@ucsb.edu 3 Department of Pediatrics and Center for Molecular Medicine Cologne, Faculty of Medicine and University Hospital Cologne, University of Cologne, Cologne, Germany, Department II of Internal Medicine, Faculty of Medicine and University Hospital Cologne, University of Cologne, Cologne, Germany

Publication info: Cellular Signalling 72 (Aug 2020).

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 190 of 266

Natural sweeteners: The relevance of food naturalness for consumers, food security aspects, sustainability and health impacts

Author: Saraiva, Ariana 1 ; Carrascosa, Conrado 1 ; Raheem, Dele 2 ; Ramos, Fernando 3 ; Raposo, António 4 1 Department of Animal Pathology and Production, Bromatology and Food Technology, Faculty of Veterinary, Universidad de Las Palmas de Gran Canaria, Trasmontaña s/n, Arucas, 35413, Spain 2 Northern Institute for Environmental and Minority Law (NIEM), Arctic Centre, University of Lapland, Rovaniemi, Lapland, 96101, Finland 3 Pharmacy Faculty, University of Coimbra, Azinhaga de Santa Comba, Coimbra, 3000-548, Portugal, REQUIMTE/LAQV, Apartado, R. D. Manuel II, Oporto, 55142, Portugal 4 Department for Management of Science and Technology Development, Ton Duc Thang University, Ho Chi Minh City, Viet Nam, Faculty of Environment and Labour Safety, Ton Duc Thang University, Ho Chi Minh City, Viet Nam antonio.raposo@tdtu.edu.vn

Publication info: International Journal of Environmental Research and Public Health 17.17 (Sep 1, 2020): 1-22.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

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Exploring novel endothelin receptor blocker as anti-hypertensive agents identified from a natural drugs library using induced fit docking and biological assay

Author: Patel, Jimish R. 1 ; Joshi, Hirak V. 2 ; Shah, Ujashkumar A. 2 ; Patel, Jayvadan K. 2 1 Department of Pharmaceutical Chemistry, Shri B.M. Shah College of Pharmaceutical Education and Research, College Campus, Dhansura Road, Modasa, Gujarat, 383315, India, Sankalchand Patel University, S.K. Campus, Kamana Cross Road, Visnagar, Gujarat, 384315, India 2 Department of Quality Assurance & Pharmaceutical Chemistry, Nootan Pharmacy College, Sankalchand Patel University, S.K. Campus, Kamana Cross Road, Visnagar, Gujarat, 384315, India jimish_patel_1986@yahoo.co.in

Publication info: Medicinal Plants 12.3 (Sep 2020): 405-413.

[ProQuest document link](#) **Databases:** Embase® (1947 - current)

Document 192 of 266

Nematicidal activity of Stevia rebaudiana (Bertoni) assisted by phytochemical analysis

Author: Ntalli, Nikoletta 1 ; Kasiotis, Konstantinos M. 2 ; Baira, Eirini 2 ; Stamatis, Christos L. 3 ; Machera, Kyriaki 2 1 Laboratory of Biological Control of Pesticides, Department of Pesticides Control and Phytopharmacy, Benaki Phytopathological Institute, 8 St. Delta Street, Athens, 14561, Greece n.ntalli@bpi.gr 2 Laboratory of Pesticides' Toxicology, Department of Pesticides Control and Phytopharmacy, Benaki Phytopathological Institute, 8 St. Delta Street, Athens, 14561, Greece k.kasiotis@bpi.gr ; e.baira@bpi.gr ; k.machera@bpi.gr 3 Stevia Hellas Coop, 6th klm Lamia-Karpenisi, Lamia, PS 35131, Greece christos.stamatis@stevianet.gr

Publication info: Toxins 12.5 (May 2020).

[ProQuest document link](#) **Databases:** Embase® (1947 - current)

Document 193 of 266

Anticancer property of hexane extract of Suaeda fruticose plant leaves against different cancer cell lines

Author: Saleh, Kamel A. 1 ; Albinhassan, Tahani H. 1 ; Al-Ghazzawi, Adel M. 2 ; Mohaya, Abdulrahman 3 ; Shati, Ali A. 1 ; Ayoub, Huda J. 4 ; Abdallah, Qasem M. 5 1 Department of Biology, Science College, King Khalid University, PO Box 9004, Abha, Saudi Arabia kasaleh@kku.edu.sa 2 Department of Chemistry, Science College, King Khalid University, PO Box 9004, Abha, Saudi Arabia 3 Asser Toxicology Center, King Abdullallah Street, PO Box 1988, Abha, 61441, Saudi Arabia 4 Department of Pharmacognosy, Pharmacy College, Alzaytoonah University, Amman, Jordan 5 Department of Pharmacology, Pharmacy College, University of Petra, Amman, Jordan

Publication info: Tropical Journal of Pharmaceutical Research 19.1 (2020): 129-136.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 194 of 266

Consumo de edulcorantes no calóricos en embarazadas chilenas: estudio transversal

Author: Fuentealba Arévalo, Fabiola 1 ; Espinoza Espinoza, Jonathan 1 ; Salazar Ibañeta, Carolina 1 ; Durán Agüero, Samuel 1 1 Facultad de Ciencias para el Cuidado de la Salud. Universidad San Sebastián

Publication info: Nutricion hospitalaria 36.4 (Aug 26, 2019): 890-897.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 195 of 266

Recent advancements in fungal-derived fuel and chemical production and commercialization

Author: Ekas, Holly 1 ; Deaner, Matthew 1 ; Alper, Hal S 2 1 McKetta Department of Chemical Engineering, The University of Texas at Austin, 200 E Dean Keeton St. Stop C0400, Austin, TX, 78712, United States 2 McKetta Department of Chemical Engineering, The University of Texas at Austin, 200 E Dean Keeton St. Stop C0400, Austin, TX, 78712, United States, Institute for Cellular and Molecular Biology, The University of Texas at Austin, 2500 Speedway Avenue, Austin, TX, 78712, United States halper@che.utexas.edu

Publication info: Current Opinion in Biotechnology 57 (Jun 2019): 1-9.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 196 of 266

The optimization of prebiotic sucrose-free mango nectar by response surface methodology: The effect of stevia and inulin on physicochemical and rheological properties

Author: Alizadeh, Ainaz 1 ; Oskuyi, Amin Seyedan 1 ; Amjadi, Sajed 2 1 Department of Food Science and Technology, Islamic Azad University, Tabriz Branch, Tabriz, Iran 2 Department of Food Science and Technology, Faculty of Agriculture, Urmia University, Urmia, Iran, Biotechnology Research Center, Tabriz University of Medical Sciences, Tabriz, Iran

Publication info: Food science and technology international = Ciencia y tecnologia de los alimentos internacional 25.3 (Apr 1, 2019): 243-251.

[ProQuest document link](#) **Databases:** Embase® (1947 - current)

Document 197 of 266

Bioproduction of the recombinant SWEET protein thaumatin: Current state of the art and perspectives

Author: Joseph, Jewel Ann 1 ; Akkermans, Simen 1 ; Nimmegeers, Philippe 1 ; Van Impe, Jan F.M. 1 1 BioTeC+, Chemical and Biochemical Process Technology and Control, Department of Chemical Engineering, KU Leuven, Leuven, Belgium, Optimization in Engineering Center-of-Excellence, KU Leuven, Leuven, Belgium, CPMF2, Flemish Cluster Predictive Microbiology in Foods, Leuven, Belgium jan.vanimpe@kuleuven.be

Publication info: Frontiers in Microbiology 10.APR (2019).

[ProQuest document link](#) **Databases:** Embase® (1947 - current)

Document 198 of 266

Sweetening agents and sweeteners in dietary supplements for children-analysis of the Polish market

Author: Piekara, Agnieszka 1 ; Krzywonos, Małgorzata 1 ; Szymańska, Anna 1 1 Department of Bioprocess Engineering, Wrocław University of Economics and Business, room 301H, Komandorska 118/120, Wrocław, 53-345, Poland
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Publication info: Nutrients 12.8 (Aug 2020): 1-9.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 199 of 266

Comparison of stevia leaf extract (*Stevia rebaudiana*) and sucrose sugar addition on the textural properties of Kanomtian crust.

Author: Suwannarong, S; Chotthanom, P; Banlue, K; Niyom, B; Kanarun, K

Publication info: Kaen Kaset = Khon Kaen Agriculture Journal 47.Suppl. 1 (2019): 657-662.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 200 of 266

Cirrhosis induced by thioacetamide is prevented by stevia. Molecular mechanisms.

Author: Ramos-Tovar, E; Casas-Grajales, S; Hernández-Aquino, E; Flores-Beltrán, R E; Galindo-Gómez, S; Vera-Aguilar, E; Diaz-Ruiz, A; Montes, S; Camacho, J; Tsutsumi, V; Muriel, P

Publication info: Journal of Functional Foods 52 (2019): 552-564.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 201 of 266

Bioactivity profile of the diterpene isosteviol and its derivatives.

Author: Ullah, Asad; Munir, Sidra; Mabkhot, Yahia; Badshah, S L

Publication info: Molecules 24.4 (2019): 678.

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 202 of 266

Consumption Of The Sugar Substitute Stevia Leads To Cross-Reactivity Of CEDIA® Buprenorphine II Immunoassay

Author: Plattner, Sabine 1 ; Pavlic, Marion 1 ; Pitterl, Florian 1 ; Schubert, Birthe 1 1 Institute of Legal Medicine, Medical University of Innsbruck, Innsbruck, Austria, Austria

Publication info: Journal of analytical toxicology (Oct 26, 2020).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 203 of 266

Non-nutritive Sweeteners Induce Hypothalamic ER Stress Causing Abnormal Axon Outgrowth

Author: Park, Soyung 1 ; Sethi, Sunjay 1 ; Bouret, Sebastien G 2 1 Development Neuroscience Program, Children's Hospital Los Angeles, Los Angeles, CA, United States, United States 2 INSERM, Laboratory of Development and Plasticity of the Neuroendocrine Brain, Jean-Pierre Aubert Research Centre, UMR-S 1172, Lille, France, University of Lille, FHU 1, 000 Days for Health, Lille, France, France

Publication info: Frontiers in endocrinology 10 (Dec 17, 2019): 876.

[ProQuest document link](#) **Databases:** MEDLINE® (1946 - current)

Document 204 of 266

Changes in nutrient and calorie intake, adipose mass, triglycerides and TNF- α concentrations after non-caloric sweetener intake: Pilot study

Author: Sánchez-Delgado, Marcela 1 ; Estrada, José Antonio 1 ; Paredes-Cervantes, Vladimir 2 ; Kaufer-Horwitz, Martha 3 ; Contreras, Irazú, Phd 1 1 Laboratorio de Neuroquímica, Facultad de Medicina, Universidad Autónoma Del Estado de México, Paseo Tollocan s/n esq. Jesús Carranza, Colonia Moderna de la Cruz, Toluca, C.P. 50180, Mexico icontrerasg@uaemex.mx 2 UMAE, Hospital General, Centro Médico Nacional la Raza, Instituto Mexicano Del Seguro Social, Mexico 3 Clínica de Obesidad y Trastornos de la Conducta Alimentaria, Instituto Nacional de Ciencias Médicas y Nutrición, Salvador Zubirán, Mexico
Publication info: International Journal for Vitamin and Nutrition Research (2019).

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 205 of 266

Changes in Appetite Regulation-Related Signaling Pathways in the Brain of Mice Supplemented with Non-nutritive Sweeteners

Author: Contreras-Chavez, Gerson G 1 ; Estrada, José A 1 ; Contreras, Irazú 1 1 Laboratorio de Neuroquímica, Facultad de Medicina, Universidad Autónoma del Estado de México, Paseo Tollocan S/N Esq. Jesús Carranza, Colonia Moderna de La Cruz, Estado de México, 50180, Toluca, Mexico, Mexico icontrerasg@uaemex.mx
Publication info: Journal of molecular neuroscience : MN (Oct 31, 2020).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 206 of 266

De novo transcriptome dataset of Stevia rebaudiana accession MS007.

Author: Samsulrizal, Nurul H 1 ; Khadzran, Khairul S 1 ; Shaarani, Siti Hn 2 ; Noh, Abdul L 1 ; Sundram, Tamil Cm 1 ; Naim, Mohd Azrul 3 ; Zainuddin, Zarina 1 1 Department of Plant Science, Kulliyah of Science, International Islamic University Malaysia, Jln. Sultan Ahmad Shah, Kuantan, Pahang, Malaysia. 2 Faculty of Chemical & Natural Resources Engineering, Universiti Malaysia Pahang, Lebuhraya Tun Razak, Gambang, Pahang, Malaysia. 3 Research Unit for Bioinformatics and Computational Biology (RUBIC), Kulliyah of Science, International Islamic University Malaysia, Jln. Sultan Ahmad Shah, Kuantan, Pahang, Malaysia.

Publication info: Data in brief 28 (February 2020): 104811.

[ProQuest document link](#)

Databases: TOXLINE (1999 - current)

Document 207 of 266

De novo transcriptome dataset of Stevia rebaudiana accession MS007

Author: Samsulrizal, Nurul H 1 ; Khadzran, Khairul S 2 ; Shaarani, Siti Hn 3 ; Noh, Abdul L 2 ; Sundram, Tamil Cm 2 ; Naim, Mohd Azrul 4 ; Zainuddin, Zarina 2 1 Department of Plant Science, Kulliyah of Science, International Islamic University Malaysia, Jln. Sultan Ahmad Shah, Kuantan, Pahang, Malaysia, Research Unit for Bioinformatics and Computational Biology (RUBIC), Kulliyah of Science, International Islamic University Malaysia, Jln. Sultan Ahmad Shah, Kuantan, Pahang, Malaysia, Malaysia 2 Department of Plant Science, Kulliyah of Science, International Islamic University Malaysia, Jln. Sultan Ahmad Shah, Kuantan, Pahang, Malaysia, Malaysia 3 Faculty of Chemical & Natural Resources Engineering, Universiti Malaysia Pahang, Lebuhraya Tun Razak, Gambang, Pahang, Malaysia, Malaysia 4 Research Unit for Bioinformatics and Computational Biology (RUBIC), Kulliyah of Science, International Islamic University Malaysia, Jln. Sultan Ahmad Shah, Kuantan, Pahang, Malaysia, Malaysia

Publication info: Data in brief 28 (Nov 15, 2019): 104811.

[ProQuest document link](#)

Databases: ToxFile® (1900 - current)

Document 208 of 266

Stevia-containing beverage

Author: Anonymous

Publication info: Nakajima, Makoto. Kobayashi, Yasuyuki. US 10750764. 25 August 2020 (20200825). US.

[ProQuest document link](#)

Databases: BIOSIS Previews® (1926 - current)

Document 209 of 266

Overuse of Non-caloric Sweeteners in Foods and Beverages in Chile: A Threat to Consumers' Free Choice?

Author: Samba, Verónica 1 ; López-Arana, Sandra 1 ; Cáceres, Paola 1 ; Abrigo, Karen 1 ; Collinao, Javiera 2 ; Espinoza, Alexandra 2 ; Valenzuela, Sabrina 2 ; Carvajal, Bielka 3 ; Prado, Gabriel 1 ; Peralta, Rebeca 1 ; Gotteland, Martin 4 1 Department of Nutrition, Faculty of Medicine, University of Chile, Santiago, Chile, Chile 2 Faculty of Medicine, School of Nutrition and Dietetics, University of Chile, Santiago, Chile, Chile 3 Department of Women and Newborn's Health Promotion, Faculty of Medicine, University of Chile, Santiago, Chile, Chile 4 Department of Nutrition, Faculty of Medicine, University of Chile, Santiago, Chile, Human Nutrition Unit, Institute of Nutrition and Food Technology (INTA), University of Chile, Santiago, Chile, Chile

Publication info: Frontiers in nutrition 7 (Jun 17, 2020): 68.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 210 of 266

Microorganisms for diterpene production

Author: Anonymous

Publication info: Boer, Viktor Marius. Suir, Erwin. US 10689681. 23 June 2020 (20200623). US.

[ProQuest document link](#)

Databases: BIOSIS Previews® (1926 - current)

Document 211 of 266

Heat treatment to produce glycosides

Author: Anonymous

Publication info: Anderson, James C.. Brower, Robert J.. Carlson, Ting Liu. Flores, Belit. Gaspard, Dan S.. Mortenson, Kristopher T.. Nygaard, Richard. Paulson, Nicole. Rasmussen, Maribeth. US 10612065. 07 April 2020 (20200407). US.

[ProQuest document link](#)

Databases: BIOSIS Previews® (1926 - current)

Document 212 of 266

Optimization of regeneration and Agrobacterium-mediated transformation of Stevia (Stevia rebaudiana Bertoni): a commercially important natural sweetener plant

Author: Taak, Pooja 1 ; Tiwari, Siddharth 2 ; Koul, Bhupendra 1 1 School of Bioengineering and Biosciences, Lovely Professional University, Jalandhar-Delhi G.T. Road (NH-1), Phagwara, Punjab, 144411, India bhupendra.18673@lpu.co.in 2 Department of Biotechnology, Govt. of India, National Agri-Food Biotechnology Institute (NABI), Knowledge City, Sector 81S.A.S. Nagar, Mohali, Punjab, 140306, India siddharth@nabi.res.in

Publication info: Scientific reports 10.1 (Oct 1, 2020): 16224.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 213 of 266

Development of screening methods for functional characterization of UGTs from *Stevia rebaudiana*

Author: Petit, Eva 1 ; Berger, Monique 1 ; Camborde, Laurent 2 ; Vallejo, Veronica 3 ; Daydé, Jean 1 ; Jacques, Alban 1 1 Equipe Physiologie, Pathologie et Génétique Végétales (PPGV), INP-PURPAN, Université de Toulouse, 75 voie du TOEC ,BP 57611, 31076, France monique.berger@purpan.fr 2 Laboratoire de Recherche en Sciences Végétales (LRSV), CNRS, Université Paul Sabatier (UPS), Toulouse, France 3 PepsiCo, 700 Anderson Hill Rd., Purchase, NY, 10577, USA

Publication info: Scientific reports 10.1 (Sep 15, 2020): 15137.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 214 of 266

Greenhouse evaluation of branching, leaf yield and biochemical compositions of *Stevia rebaudiana* Bertoni to decapitation and foliar application of abscisic acid and fluridone

Author: Tavakoli Hasanaklou, Nasibeh; Sedghi, Mohammad; Moradi, Foad; Ebadi Khazineh Ghadim, Ali; Jahanbakhsh Ghodehkahriz, Sodabeh

Publication info: Functional plant biology : FPB (Jul 31, 2020).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 215 of 266

An Efficient *Stevia rebaudiana* Transformation System and In vitro Enzyme Assays Reveal Novel Insights into UGT76G1 Function

Author: Wu, Qian 1 ; La Hovary, Christophe 2 ; Chen, Han-Yi 3 ; Li, Xu 3 ; Eng, Hayde 1 ; Vallejo, Veronica 4 ; Qu, Rongda 5 ; Dewey, Ralph E 5 1 Department of Crop and Soil Sciences, North Carolina State University, Raleigh, NC, 27695, USA, Elo Life Systems, 3054 East Cornwallis Road, Durham, NC, 27709, USA, USA 2 Department of Crop and Soil Sciences, North Carolina State University, Raleigh, NC, 27695, USA, Johnson County Community College, 12345 College Blvd, Overland Park, KS, 66210, USA, USA 3 Department of Plant and Microbial Biology, North Carolina State University, Raleigh, NC, 27695, USA, USA 4 PepsiCo, 700 Anderson Hill Rd, Purchase, NY, 10577, USA, USA 5 Department of Crop and Soil Sciences, North Carolina State University, Raleigh, NC, 27695, USA, USA ralph_dewey@ncsu.edu

Publication info: Scientific reports 10.1 (Feb 28, 2020): 3773.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

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A Novel Urinary Biomarker Approach Reveals Widespread Exposure to Multiple Low-Calorie Sweeteners in Adults

Author: Logue, Caomhan 1 ; Dowey, Le Roy C 1 ; Verhagen, Hans 2 ; Strain, J.J. 1 ; O'Mahony, Maeve 1 ; Kapsokefalou, Maria 3 ; Athanasatou, Adelais 3 ; Gallagher, Alison M. 1 1 Nutrition Innovation Centre for Food and Health (NICHE), School of Biomedical Sciences, Ulster University, Coleraine, United Kingdom 2 Nutrition Innovation Centre for Food and Health (NICHE), School of Biomedical Sciences, Ulster University, Coleraine, United Kingdom, European Food Safety Authority, Parma, Italy 3 Department of Food Science and Human Nutrition, Agricultural University of Athens, Athens, Greece

Publication info: The Journal of nutrition 150.9 (Sep 1, 2020): 2435-2441.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 217 of 266

Diterpene production in Yarrowia

Author: Anonymous

Publication info: Broers, Nicolette Jasmijn. Boer, Viktor Marius. Lawrence, Adam G.. US 10273519. 30 April 2019 (20190430). US.

[ProQuest document link](#)

Databases: BIOSIS Previews® (1926 - current)

Document 218 of 266

Chinese Sweet Leaf Tea (*Rubus suavissimus*) Mitigates LPS-Induced Low-Grade Chronic Inflammation and Reduces the Risk of Metabolic Disorders in a C57BL/6J Mouse Model

Author: Zhang, Hua; Qi, Ruili; Zeng, Yuhan; Tsao, Rong; Mine, Yoshinori

Publication info: Journal of Agricultural and Food Chemistry 68.1 (Jan 8, 2020): 138-146.

[ProQuest document link](#)

Databases: BIOSIS Previews® (1926 - current)

Document 219 of 266

Influences of non-nutritive sweeteners on ovarian and uterine expression of T1R2 and T1R3 in peripubertal female guinea pigs

Author: Li, Junrong 1 ; Shen, Ting 2 ; Shi, Fangxiong 3 ; Fu, Yan 4 1 College of Agriculture and Bio-engineering, Jinhua Polytechnic, Jinhua, China, College of Animal Science, Zhejiang University, Hangzhou, China, China 2 College of Agriculture and Bio-engineering, Jinhua Polytechnic, Jinhua, China, China 3 College of Animal Science and Technology, Nanjing Agricultural University, Nanjing, China, China 4 College of Animal Science, Zhejiang University, Hangzhou, China, China

Publication info: Animal science journal = Nihon chikusan Gakkaiho 91.1 (Jan 2020 - Dec 2020): e13348.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 220 of 266

Temporal sweetness and side tastes profiles of 16 sweeteners using temporal check-all-that-apply (TCATA)

Author: Tan, Vicki Wei Kee; Wee, May Sui Mei; Tomic, Oliver; Forde, Ciaran G.

Publication info: Food Research International 121 (Jul 2019): 39-47.

[ProQuest document link](#) **Databases:** BIOSIS Previews® (1926 - current)

Document 221 of 266

Types and Amounts of Nonnutritive Sweeteners Purchased by US Households: A Comparison of 2002 and 2018 Nielsen Homescan Purchases

Author: Dunford, Elizabeth K.; Miles, Donna R.; Ng, Shu Wen; Popkin, Barry

Publication info: Journal of the Academy of Nutrition and Dietetics 120.10 (Oct 1, 2020): 1662-1671.

[ProQuest document link](#) **Databases:** Embase® (1947 - current)

Document 222 of 266

Production of low-calorie apricot nectar sweetened with stevia: Impact on qualitative, sensory, and nutritional profiles

Author: Reale, Anna 1 ; Di Renzo, Tiziana 1 ; Russo, Antonio 1 ; Niro, Serena 2 ; Ottobrino, Antonio 1 ; Pellicano, Mario Paolo 1 1 Institute of Food Science National Research Council ISA-CNR Avellino Italy 2 Department of Agricultural, Environmental and Food Sciences DiAAA University of Molise Campobasso Italy, Environmental and Food Sciences DiAAA University of Molise Campobasso Italy

Publication info: Food science & nutrition 8.4 (Mar 3, 2020): 1837-1847.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 223 of 266

Phytol, (E)-nerolidol and spathulenol from *Stevia rebaudiana* leaf essential oil as effective and eco-friendly botanical insecticides against *Metopolophium dirhodum*.

Author: Benelli, G; Pavela, R; Drenaggi, E; Desneux, N; Maggi, F

Publication info: Industrial Crops and Products 155 (2020).

[ProQuest document link](#)

Databases: CAB ABSTRACTS (1910 - current)

Document 224 of 266

When the Beverage Is Sweet, How Does the Liver Feel?

Author: Xi, Dong 1 ; Kohli, Rohit 1 1 Division of Gastroenterology, Hepatology and Nutrition, Children's Hospital Los Angeles, University of Southern California, 4650 Sunset Blvd, MS 78, Los Angeles, CA, United States rokohli@chla.usc.edu

Publication info: Current Treatment Options in Pediatrics 5.4 (Dec 1, 2019): 458-465.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 225 of 266

Discovery of NDM-1 inhibitors from natural products

Author: Shi, Cheng 1 ; Chen, Jiaying 1 ; Xiao, Bin 1 ; Kang, Xinyue 1 ; Lao, Xingzhen 1 ; Zheng, Heng 1 1 School of Life Science and Technology, China Pharmaceutical University, Nanjing 210009, PR China, PR China lao@cpu.edu.cn ; zhengh18@hotmail.com

Publication info: Journal of global antimicrobial resistance 18 (Sep 2019): 80-87.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 226 of 266

A case report on the probable herb-drug interaction between stevia products and etoricoxib relating to declined kidney function

Author: Tangkiatkumjai, M. 1 ; Janma, J. 2 ; Changsirikulchai, S. 2 1 Department of Clinical pharmacy, Faculty of Pharmacy, Srinakharinwirot University, Nakhon Nayok, Thailand 2 Division of Nephrology, Department of Medicine, Faculty of Medicine, Srinakharinwirot University, Nakhon Nayok, Thailand siribha@g.swu.ac.th

Publication info: Journal of the Medical Association of Thailand, suppl. Supplement 6 102.7 (Jul 2019): 100-104.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 227 of 266

First Report of *Septoria* sp. Infecting *Stevia rebaudiana* in France and Screening of *Stevia rebaudiana* Genotypes for Host Resistance

Author: Hastoy, C 1 ; Le Bihan, Z 1 ; Gaudin, J 2 ; Cosson, P 3 ; Rolin, D 3 ; Schurdi-Levraud, V 3 1 1 INRA, Université de Bordeaux, UMR 1332 Biologie du Fruit et Pathologie, Villenave d'Ornon, France and Oviatis SA, Lacaussade, France, France 2 2 INRA, UMR 1065 Santé et Agroécologie du Vignoble, ISVV, Univ. Bordeaux, Bordeaux Sciences-Agro, Villenave d'Ornon, France, France 3 3 INRA, Université de Bordeaux, UMR 1332 Biologie du Fruit et Pathologie, Villenave d'Ornon, France, France

Publication info: Plant disease 103.7 (Jul 2019): 1544-1550.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 228 of 266

Drying Kinetics and Quality of Dehydrated Cranberries Pretreated by Traditional and Innovative Techniques

Author: Wiktor, A. 1 ; Nowacka, M. 1 ; Anuszevska, A. 1 ; Rybak, K. 1 ; Dadan, M. 1 ; Witrowa-Rajchert, D. 1 1 Dept. of Food Engineering and Process Management, Faculty of Food Sciences, Warsaw Univ. of Life Sciences (WULS-SGGW), Warsaw, Poland

Publication info: Journal of food science 84.7 (Jul 1, 2019): 1820-1828.

[ProQuest document link](#)

Databases: Embase® (1947 - current)

Document 229 of 266

Isosteviol Protects Free Fatty Acid- and High Fat Diet-Induced Hepatic Injury via Modulating PKC-β/p66Shc/ROS and Endoplasmic Reticulum Stress Pathways

Author: Yi, Hongwei 1 ; Xu, Deyi 1 ; Wu, Xudong 2 ; Xu, Fang 2 ; Lin, Lin 1 ; Zhou, Huiping 3 1 1 Department of Pharmacology, School of Medicine, Southeast University, Nanjing, China, China 2 2 State Key Laboratory of Pharmaceutical Biotechnology, School of Life Sciences, Nanjing University, Nanjing, China, China 3 3 Department of Microbiology and Immunology, Virginia Commonwealth University and McGuire Veterans Affairs Medical Center, Richmond, Virginia, Virginia

Publication info: Antioxidants & redox signaling 30.17 (Jun 10, 2019): 1949-1968.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 230 of 266

Contemporary Trends and Habits in the Consumption of Sugar and Sweeteners-A Questionnaire Survey among Poles

Author: Pielak, Marlena 1 ; Czarniecka-Skubina, Ewa 1 ; Trafiałek, Joanna 1 ; Głuchowski, Artur 1 1 Department of Food Gastronomy and Food Hygiene, Faculty of Human Nutrition and Consumer Sciences, Warsaw University of Life Sciences, 02-787 Warsaw, Poland, Poland marlenapielak@gmail.com ; ewa_czarniecka_skubina@sggw.pl ; joanna_trafialek@sggw.pl ; artur_gluchowski@sggw.pl

Publication info: International journal of environmental research and public health 16.7 (Apr 1, 2019).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 231 of 266

Phylogenomic analysis of UDP-dependent glycosyltransferases provides insights into the evolutionary landscape of glycosylation in plant metabolism

Author: Wilson, Alexander E 1 ; Tian, Li 1 1 Department of Plant Sciences, University of California, Davis, CA, 95616, USA, USA

Publication info: The Plant journal : for cell and molecular biology 100.6 (Dec 2019): 1273-1288.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 232 of 266

High affinity of 4-(4-(dimethylamino)styryl)-N-methylpyridinium transport for assessing organic cation drugs in hepatocellular carcinoma cells

Author: Jinakote, Metee 1 ; Ontawong, Atcharaporn 2 ; Soodvilai, Sunhapas 3 ; Pimta, Jeerawat 3 ; Pasachan, Tiphida 4 ; Chatsudthipong, Varanuj 3 ; Srimaroeng, Chutima 4 1 Faculty of Oriental Medicine, Chiang Rai College, Chiang Rai, 57000, Thailand, Department of Physiology, Faculty of Medicine, Chiang Mai University, Chiang Mai, 50200, Thailand, Thailand 2 Unit of Excellence of Coffee, Division of Physiology, School of Medical Sciences, University of Phayao, Phayao, 56000, Thailand, Thailand 3 Research Center of Transport Protein for Medical Innovation, Department of Physiology, Faculty of Science, Mahidol University, Bangkok, 10400, Thailand, Thailand 4 Department of Physiology, Faculty of Medicine, Chiang Mai University, Chiang Mai, 50200, Thailand, Thailand

Publication info: Fundamental & clinical pharmacology 34.3 (Jun 2020): 365-379.

[ProQuest document link](#) **Databases:** MEDLINE® (1946 - current)

Document 233 of 266

Isoniazid-phytochemical conjugation: A new approach for potent and less toxic anti-TB drug development

Author: Swain, Shasank S 1 ; Paidasetty, Sudhir K 2 ; Padhy, Rabindra N 3 ; Hussain, Tahziba 4 1 Division of Microbiology and NCDs, ICMR-Regional Medical Research Centre, Bhubaneswar, India, Central Research Laboratory, Institute of Medical Sciences and SUM Hospital, Siksha 'O' Anusandhan Deemed to be University, Bhubaneswar, India, India 2 Department of Pharmaceutical Chemistry, School of Pharmaceutical Sciences, Siksha 'O' Anusandhan Deemed to be University, Bhubaneswar, India, India 3 Central Research Laboratory, Institute of Medical Sciences and SUM Hospital, Siksha 'O' Anusandhan Deemed to be University, Bhubaneswar, India, India 4 Division of Microbiology and NCDs, ICMR-Regional Medical Research Centre, Bhubaneswar, India, India

Publication info: Chemical biology & drug design 96.2 (Aug 2020): 714-730.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 234 of 266

Effect of the homogenization process on the sensory and rheological properties in model system

Author: da Silva Pereira, Gerlândia 1 ; Leite, Thiago S 2 ; Schmidt, Flávio L 2 ; Cristianini, Marcelo 2 ; Bolini, Helena M A 1 1 Department of Food and Nutrition (DEPAN), School of Food Engineering (FEA), University of Campinas (UNICAMP), Campinas, Brazil, Brazil 2 Department of Food Technology (DTA), School of Food Engineering (FEA), University of Campinas (UNICAMP), Campinas, Brazil, Brazil

Publication info: Journal of texture studies 51.2 (Apr 2020): 352-360.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

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Protective effects of Stevia rebaudiana aqueous extract on experimental unilateral testicular ischaemia/reperfusion injury in rats

Author: Ganjiani, Vahid 1 ; Ahmadi, Nasrollah 2 ; Raayat Jahromi, Alireza 3 1 School of Veterinary Medicine, Shiraz University, Shiraz, Iran, Iran 2 Veterinary Pathology, School of Veterinary Medicine, Shiraz University, Shiraz, Iran, Iran 3 Veterinary Surgery, School of Veterinary Medicine, Shiraz University, Shiraz, Iran, Iran

Publication info: Andrologia 52.2 (Mar 2020): e13469.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 236 of 266

Etoposide Loaded SPION-PNIPAM Nanoparticles Improve the in vitro Therapeutic Outcome on Metastatic Prostate Cancer Cells via Enhanced Apoptosis

Author: Erkisa, Merve 1 ; Ari, Ferda 2 ; Ulku, Irem 3 ; Khodadust, Rouhollah 3 ; Yar, Yasemin 4 ; Yagci Acar, Havva 5 ; Ulukaya, Engin 6 1 Bursa Uludag University, Science and Art Faculty, Department of Biology, 16059, Bursa, Turkey, Turkey 2 Istinye University, Faculty of Medicine, Molecular Cancer Research Center, 34010, Istanbul, Turkey, Turkey 3 Bursa Uludag University, Science and Art Faculty, Department of Biology, 16059, Bursa, Turkey, Turkey 4 Koc University, Department of Chemistry, 34450, Istanbul, Turkey, Turkey 5 Koc University, Materials Science and Engineering, 34450, Istanbul, Turkey, Turkey 6 Istinye University, School of Medicine, Department of Clinical Biochemistry, 34010, Istanbul, Turkey, Turkey

Publication info: Chemistry & biodiversity (Sep 11, 2020).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 237 of 266

Sesquiterpene glucosylation mediated by glucosyltransferase UGT91Q2 is involved in the modulation of cold stress tolerance in tea plants

Author: Zhao, Mingyue 1 ; Zhang, Na 1 ; Gao, Ting 1 ; Jin, Jieyang 1 ; Jing, Tingting 1 ; Wang, Jingming 1 ; Wu, Yi 1 ; Wan, Xiaochun 1 ; Schwab, Wilfried 2 ; Song, Chuankui 1 1 State Key Laboratory of Tea Plant Biology and Utilization, International Joint Laboratory on Tea Chemistry and Health Effects, Anhui Agricultural University, Hefei, Anhui, 230036, China, China 2 State Key Laboratory of Tea Plant Biology and Utilization, International Joint Laboratory on Tea Chemistry and Health Effects, Anhui Agricultural University, Hefei, Anhui, 230036, China, Biotechnology of Natural Products, Technische Universität München, Liesel-Beckmann-Str. 1, Freising, 85354, Germany, Germany

Publication info: The New phytologist 226.2 (Apr 2020): 362-372.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 238 of 266

Mining of UDP-glucosyltransferases in licorice for controllable glycosylation of pentacyclic triterpenoids

Author: Zhang, Liang 1 ; Ren, Shichao 1 ; Liu, Xiaofei 1 ; Liu, Xiaochen 1 ; Guo, Fang 1 ; Sun, Wentao 1 ; Feng, Xudong 1 ; Li, Chun 2 1 Institute of Biochemical Engineering, Department of Chemical Engineering, School of Chemistry and Chemical Engineering, Beijing Institute of Technology, Beijing, China, China 2 Institute of Biochemical Engineering, Department of Chemical Engineering, School of Chemistry and Chemical Engineering, Beijing Institute of Technology, Beijing, China, Key Laboratory for Industrial Biocatalysis, Ministry of Education, Department of Chemical Engineering, Tsinghua University, Beijing, China, China

Publication info: Biotechnology and bioengineering (Jul 27, 2020).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 239 of 266

Functional genomics analysis reveals two novel genes required for littorine biosynthesis

Author: Qiu, Fei 1 ; Zeng, Junlan 1 ; Wang, Jing 1 ; Huang, Jian-Ping 2 ; Zhou, Wei 3 ; Yang, Chunxian 1 ; Lan, Xiaozhong 4 ; Chen, Min 5 ; Huang, Sheng-Xiong 2 ; Kai, Guoyin 3 ; Liao, Zhihua 1 1 Chongqing Key Laboratory of Plant Resource Conservation and Germplasm Innovation, SWU-TAAHC Medicinal Plant Joint R&D Centre, School of Life Sciences, Southwest University, Chongqing, 400715, China, China 2 State Key Laboratory of Phytochemistry and Plant Resources in West China, CAS Centre for Excellence in Molecular Plant Sciences, Kunming Institute of Botany, Chinese Academy of Sciences, Kunming, 650201, China, China 3 Laboratory of Medicinal Plant Biotechnology, College of Pharmacy, Zhejiang Chinese Medical University, Hangzhou, Zhejiang, 311402, China, China 4 TAAHC-SWU Medicinal Plant Joint R&D Centre, Tibetan Collaborative Innovation Centre of Agricultural and Animal Husbandry Resources, Xizang Agricultural and Animal Husbandry College, Nyingchi of Tibet, 860000, China, China 5 College of Pharmaceutical Sciences, Key Laboratory of Luminescent and Real-Time Analytical Chemistry (Ministry of Education), Southwest University, Chongqing, 400715, China, China

Publication info: The New phytologist 225.5 (Mar 2020): 1906-1914.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 240 of 266

Can the Creation of Tetraploids Improve the Productivity and Quality of Stevia rebaudiana?

Author: Shock, Clinton C.; Yilma, Solomon

Publication info: Hortscience 54.9, Suppl. S (Sep 2019): S76.

[ProQuest document link](#)

Databases: BIOSIS Previews® (1926 - current)

Document 241 of 266

Effects of sunset yellow on proliferation and differentiation of intestinal epithelial cells in murine intestinal organoids

Author: Kong, Xiunan 1 ; Wang, Xiu 1 ; Qin, Yumei 1 ; Han, Jianzhong 1 1 School of Food Science and Biotechnology, Zhejiang Gongshang University, Hangzhou, China, China

Publication info: Journal of applied toxicology : JAT (Oct 15, 2020).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 242 of 266

Optimization of extraction conditions of Angelica archangelica extract and activity evaluation in experimental fibromyalgia

Author: Kaur, Anudeep 1 ; Singh, Nirmal 2 ; Bhatti, Manpreet Singh 3 ; Bhatti, Rajbir 1 1 Department of Pharmaceutical Sciences, Guru Nanak Dev University, Amritsar, India, India 2 Department of Pharmaceutical Sciences and Drug Research, Punjabi University, Patiala, India, India 3 Department of Botanical and Environmental Sciences, Guru Nanak Dev University, Amritsar, India, India

Publication info: Journal of food science (Oct 13, 2020).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 243 of 266

Flax and hempseed oil functional ingredient stabilized by inulin and chia mucilage as a butter replacer in muffin formulations

Author: Gutiérrez-Luna, Katherine 1 ; Ansorena, Diana 2 ; Astiasarán, Iciar 2 1 Department of Nutrition, Food Science and Physiology, Faculty of Pharmacy and Nutrition, Universidad de Navarra, C/Irunlarrea s/n, Pamplona, 31008, Spain, Spain 2 Department of Nutrition, Food Science and Physiology, Faculty of Pharmacy and Nutrition, Universidad de Navarra, C/Irunlarrea s/n, Pamplona, 31008, Spain, Universidad de Navarra, IDISNA - Instituto de Investigación Sanitaria de Navarra, Pamplona, Spain, Spain

Publication info: Journal of food science 85.10 (Oct 2020): 3072-3080.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 244 of 266

Effects of starch type and concentration on the physicochemical properties of bilayer-stabilized oil-in-water emulsion gels enriched with β -1,3/1,6-D-glucans

Author: Bortnowska, Grażyna 1 ; Bartkowiak, Artur 2 ; Tokarczyk, Grzegorz 1 ; Przybylska, Sylwia 1 ; Iwański, Robert 1 1 Department of Fish, Plant and Gastronomy Technology, West Pomeranian University of Technology in Szczecin, Szczecin, Poland, Poland 2 Center of Bioimmobilisation and Innovative Packaging Materials, West Pomeranian University of Technology in Szczecin, Szczecin, Poland, Poland

Publication info: Journal of the science of food and agriculture 100.13 (Oct 2020): 4879-4886.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 245 of 266

Recent progress in separation prediction of counter-current chromatography

Author: Guo, Yuru 1 ; Tong, Shengqiang 1 ; Zhang, Keqing 1 ; Yan, Jizhong 1 1 College of Pharmaceutical Science, Zhejiang University of Technology, Hangzhou, P. R. China, P. R. China

Publication info: Journal of separation science (Sep 14, 2020).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 246 of 266

Purple non-sulphur bacteria and plant production: benefits for fertilization, stress resistance and the environment

Author: Sakarika, Myrsini 1 ; Spanoghe, Janne 1 ; Sui, Yixing 1 ; Wambacq, Eva 2 ; Grunert, Oliver 3 ; Haesaert, Geert 2 ; Spiller, Marc 1 ; Vlaeminck, Siegfried E 1 1 Research Group of Sustainable Air, Energy and Water Technology, Department of Bioscience Engineering, University of Antwerp, Groenenborgerlaan 171, 2020, Antwerpen, Belgium, Belgium 2 Department of Plants and Crops, Faculty of Bioscience Engineering, Ghent University, V. Vaerwyckweg 1, 9000, Ghent, Belgium, Belgium 3 Greenyard Horticulture Belgium NV, Skaldenstraat 7a, 9042, Gent, Belgium, Belgium

Publication info: Microbial biotechnology 13.5 (Sep 2020): 1336-1365.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 247 of 266

High-performance thin-layer chromatography with atmospheric solids analysis probe mass spectrometry for analysis of gasoline polymeric additives

Author: Beaumesnil, Mathieu 1 ; Mendes Siqueira, Anna Luiza 1 ; Hubert-Roux, Marie 2 ; Loutelier-Bourhis, Corinne 2 ; Afonso, Carlos 2 ; Racaud, Amandine 3 ; Bai, Yang 3 1 Normandie Univ, COBRA, UMR6014 and FR3038, Université de Rouen, INSA de Rouen, CNRS, IRCOF, 1 rue Tesnière, 76821, Mont-Saint-Aignan Cedex, France, TOTAL Marketing Services, Research Center, 69360, Solaize, France, France 2 Normandie Univ, COBRA, UMR6014 and FR3038, Université de Rouen, INSA de Rouen, CNRS, IRCOF, 1 rue Tesnière, 76821, Mont-Saint-Aignan Cedex, France, France 3 TOTAL Marketing Services, Research Center, 69360, Solaize, France, France

Publication info: Rapid communications in mass spectrometry : RCM 34 2 (Aug 2020): e8755.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 248 of 266

Using inulin-based emulsion gels as fat substitute in salt reduced Bologna sausage

Author: de Souza Paglarini, Camila 1 ; Vidal, Vitor As 1 ; Ribeiro, Wanessa 1 ; Badan Ribeiro, Ana P 1 ; Bernardinelli, Oigres D 2 ; Herrero, Ana M 3 ; Ruiz-Capillas, Claudia 3 ; Sabadini, Edvaldo 2 ; Rodrigues Pollonio, Marise A 1 1 Department of Food Technology, School of Food Engineering, University of Campinas, Campinas, Brazil, Brazil 2 Institute of Chemistry, University of Campinas, Campinas, Brazil, Brazil 3 Institute of Food Science, Technology and Nutrition (ICTAN-CSIC), Madrid, Spain, Spain

Publication info: Journal of the science of food and agriculture (Jul 10, 2020).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 249 of 266

Effects of nonnutritive sweeteners on body weight and BMI in diverse clinical contexts: Systematic review and meta-analysis

Author: Laviada-Molina, Hugo 1 ; Molina-Segui, Fernanda 1 ; Pérez-Gaxiola, Giordano 2 ; Cuello-García, Carlos 3 ; Arjona-Villicaña, Ruy 4 ; Espinosa-Marrón, Alan 1 ; Martínez-Portilla, Raigam Jafet 5 1 Department of Human Nutrition and Metabolism Research, Health Sciences School, Universidad Marista de Merida, Mérida, Mexico, Mexico 2 Evidence-Based Medicine Department, Hospital Pediátrico de Sinaloa, Culiacán, Mexico, Mexico 3 Health Research Methods, Evidence, and Impact, McMaster University, Hamilton, Ontario, Canada, Canada 4 Endocrinology Service, Hospital Regional de Alta Especialidad de la Península de Yucatán, Merida, Mexico, Mexico 5 Nuffield Department of Primary Health Sciences, Radcliffe Observatory Quarter, University of Oxford Centre for Evidence-Based Medicine, Oxford, UK, UK

Publication info: Obesity reviews : an official journal of the International Association for the Study of Obesity 21.7 (Jul 2020): e13020.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 250 of 266

Tissue-specific study across the stem of *Taxus media* identifies a phloem-specific TmMYB3 involved in the transcriptional regulation of paclitaxel biosynthesis

Author: Yu, Chunna 1 ; Luo, Xiujun 1 ; Zhang, Chengchao 1 ; Xu, Xinyun 1 ; Huang, Jiefang 1 ; Chen, Yueyue 1 ; Feng, Shangguo 1 ; Zhan, Xiaori 1 ; Zhang, Lei 2 ; Yuan, Huwei 3 ; Zheng, Bingsong 3 ; Wang, Huizhong 1 ; Shen, Chenjia 1 1 College of Life and Environmental Sciences, Hangzhou Normal University, Hangzhou, 311121, China, Zhejiang Provincial Key Laboratory for Genetic Improvement and Quality Control of Medicinal Plants, Hangzhou Normal University, Hangzhou, 311121, China, China 2 Department of Plant Pathology, Washington State University, Pullman, WA, 99164-6430, USA, USA 3 State Key Laboratory of Subtropical Silviculture, Zhejiang A & F University, Hangzhou, 311300, China, Center for Cultivation of Subtropical Forest Resources (CCSFR), Zhejiang A & F University, Hangzhou, 311300, China, China

Publication info: The Plant journal : for cell and molecular biology 103.1 (Jul 2020): 95-110.

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Databases: MEDLINE® (1946 - current)

Document 251 of 266

Effect of sugarcane bagasse addition on physical, chemical, and sensory properties of oat flour and banana cake

Author: Silva, Amanda S 1 ; Correa, Luana G 1 ; Kanai, Rafaela S S 2 ; Shirai, Marianne A 3 1 Departamento Acadêmico de Alimentos, Universidade Tecnológica Federal do Paraná, Londrina, Brazil, Brazil 2 SL Cereais e Alimentos, Mauá da Serra, Brazil, Programa de Pós-graduação em Tecnologia de Alimentos, Universidade Tecnológica Federal do Paraná, Londrina, Brazil, Brazil 3 Programa de Pós-graduação em Tecnologia de Alimentos, Universidade Tecnológica Federal do Paraná, Londrina, Brazil, Brazil

Publication info: Journal of texture studies (Jun 9, 2020).

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 252 of 266

The effect of gelatin and thymol-loaded nanostructured lipid carrier on physicochemical, rheological, and sensory properties of sesame paste/date syrup blends as a snack bar

Author: Baqeri, Forod 1 ; Nejatian, Mohammad 1 ; Abbaszadeh, Sepideh 2 ; Taghdir, Maryam 1 1 Health Research Center, Life Style Institute, Baqiyatallah University of Medical Sciences, Tehran, Iran, Iran 2 Health Research Center, Life Style Institute, Baqiyatallah University of Medical Sciences, Tehran, Iran, Faculty of Health, Baqiyatallah University of Medical Sciences, Tehran, Iran, Iran

Publication info: Journal of texture studies 51.3 (Jun 2020): 501-510.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 253 of 266

The controversial effects of low-calorie sweeteners

Author: Pepin, Alexandra 1 ; Imbeault, Pascal 1 1 Unité de recherche sur le comportement et le métabolisme, École des Sciences de l'Activité Physique, Faculté des Sciences de la Santé, Université d'Ottawa, 125 rue Université, K1N 6N5 Ottawa, Ontario, Canada, Canada

Publication info: Medecine sciences : M/S 36.5 (May 2020): 472-478.

[ProQuest document link](#)

Databases: MEDLINE® (1946 - current)

Document 254 of 266

Recent advances in the bioanalytical methods of polyethylene glycols and PEGylated pharmaceuticals

Author: Zhang, Zhi 1 ; Zhang, Yuyao 1 ; Song, Shiwen 1 ; Yin, Lei 2 ; Sun, Dong 3 ; Gu, Jingkai 1 1 Research Center for Drug Metabolism, College of Life Science, Jilin University, Changchun, P. R. China, Beijing Institute of Drug Metabolism, Beijing, P. R. China, P. R. China 2 Research Center for Drug Metabolism, College of Life Science, Jilin University, Changchun, P. R. China, Research Institute of Translational Medicine, The First Bethune Hospital of Jilin University, Changchun, P. R. China, P. R. China 3 Department of Biopharmacy, College of Life Science, Jilin University, Changchun, P. R. China, Key Laboratory of Molecular Pharmacology and Drug Evaluation (Yantai University), Ministry of Education", Yantai University, Yantai, P. R. China, P. R. China

Publication info: Journal of separation science 43.9-10 (May 2020): 1978-1997.

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Databases: MEDLINE® (1946 - current)

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The expression of TwDXS in the MEP pathway specifically affects the accumulation of triptolide

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Trans-resveratrol extraction from peanut sprouts cultivated using fermented sawdust medium and its antioxidant activity

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Databases: MEDLINE® (1946 - current)

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Free radicals, antioxidants, nuclear factor-E2-related factor-2 and liver damage

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Induction of LOX by TGF- β 1/Smad/AP-1 signaling aggravates rat myocardial fibrosis and heart failure

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Publication info: IUBMB life 71.11 (Nov 2019): 1729-1739.

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A polyol-stevia blended sugar replacer exhibits low glycemic response among human subjects

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In ovo injection of flavanone on bone quality characteristics, biochemical parameters and antioxidant enzyme status of blood in daily chicks

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Publication info: Journal of animal physiology and animal nutrition 103.5 (Sep 2019): 1418-1426.

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Databases: MEDLINE® (1946 - current)

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Consumption of non-caloric sweeteners among pregnant Chileans: a cross-sectional study

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[ProQuest document link](#) **Databases:** MEDLINE® (1946 - current)

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4-Methylcatechol, a Flavonoid Metabolite with Potent Antiplatelet Effects

Author: Applová, Lenka 1 ; Karlíčková, Jana 2 ; Warncke, Paul 3 ; Macáková, Kateřina 2 ; Hrubša, Marcel 1 ; Macháček, Miloslav 4 ; Tvrdý, Václav 1 ; Fischer, Dagmar 3 ; Mladěnka, Přemysl 1 1 Department of Pharmacology and Toxicology, Faculty of Pharmacy in Hradec Králové, Charles University, Akademika Heyrovského 1203, 500 05, Hradec Králové, Czech Republic, Czech Republic 2 Department of Pharmaceutical Botany, Faculty of Pharmacy in Hradec Králové, Charles University, Akademika Heyrovského 1203, 500 05, Hradec Králové, Czech Republic, Czech Republic 3 Department of Pharmaceutical Technology and Biopharmacy, Institute of Pharmacy, Friedrich-Schiller-University Jena, Lessingstr. 8, 07743, Jena, Germany, Germany 4 Department of Biochemical Sciences, Faculty of Pharmacy in Hradec Králové, Charles University, Akademika Heyrovského 1203, 500 05, Hradec Králové, Czech Republic, Czech Republic

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Investigations into the performance of travelling wave enabled conventional and cyclic ion mobility systems to characterise protomers of fluoroquinolone antibiotic residues

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Publication info: Rapid communications in mass spectrometry : RCM 33 2 (Jul 2019): 11-21.

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Solubility and Permeability Improvement of Quercetin by an Interaction Between α -Glucosyl Stevia Nanoaggregates and Hydrophilic Polymer

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Publication info: Journal of pharmaceutical sciences 108.6 (Jun 2019): 2033-2040.

[ProQuest document link](#) **Databases:** ToxFile® (1900 - current)

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Stevia residue as new precursor of CO₂-activated carbon: Optimization of preparation condition and adsorption study of triclosan

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Publication info: Ecotoxicology and environmental safety 172 (May 15, 2019): 403-410.

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Prosaposin is a biomarker of mesenchymal glioblastoma and regulates mesenchymal transition through the TGF- β 1/Smad signaling pathway

Author: Jiang, Yang 1 ; Zhou, Jinpeng 2 ; Hou, Dianqi 3 ; Luo, Peng 2 ; Gao, Huiling 4 ; Ma, Yanju 5 ; Chen, Yin-Sheng 6 ; Li, Long 2 ; Zou, Dan 7 ; Zhang, Haiying 8 ; Zhang, Ye 7 ; Jing, Zhitao 2 1 Department of Neurosurgery, The First Hospital of China Medical University, Shenyang City, PR China, Department of Neurosurgery, Shanghai General Hospital, Shanghai Jiao Tong University School of Medicine, Shanghai, PR China, PR China 2 Department of Neurosurgery, The First Hospital of China Medical University, Shenyang City, PR China, PR China 3 Department of Neurosurgery, Shanghai General Hospital, Shanghai Jiao Tong University School of Medicine, Shanghai, PR China, PR China 4 College of Life and Health Sciences, Northeastern University, Shenyang, PR China, PR China 5 Department of Medical Oncology, Cancer Hospital of China Medical University, Shenyang, PR China, PR China 6 Department of Neurosurgery/Neuro-oncology, SunYat-sen University Cancer Center; State Key Laboratory of Oncology in South China; Collaborative Innovation Center for Cancer Medicine, Guangzhou, PR China, PR China 7 The First laboratory of cancer institute, the First Hospital of China Medical University, Shenyang City, PR China, PR China 8 International Education College, Liaoning University of Traditional Chinese Medicine, Shenyang City, PR China, PR China

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