



## Production of steviol glycosides in recombinant hosts

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*Publication date:*  
2016

*Document Version*  
Publisher's PDF, also known as Version of record

[Link back to DTU Orbit](#)

*Citation (APA):*  
Douchin, V., Mikkelsen, M. D., & Møller-Hansen, I. (2016). IPC No. C07H1/08; C12N15/81. Production of steviol glycosides in recombinant hosts (Patent No. WO2016023844.)

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- (51) **International Patent Classification:**  
A23L 1/236 (2006.01) C07H 1/08 (2006.01)  
C12N 15/81 (2006.01)
- (21) **International Application Number:**  
PCT/EP20 15/0683 14
- (22) **International Filing Date:**  
7 August 2015 (07.08.2015)
- (25) **Filing Language:** English
- (26) **Publication Language:** English
- (30) **Priority Data:**  
62/035,902 11 August 2014 (11.08.2014) US
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- (81) **Designated States** (unless otherwise indicated, for every kind of national protection available): AE, AG, AL, AM,

AO, AT, AU, AZ, BA, BB, BG, BH, BN, BR, BW, BY, BZ, CA, CH, CL, CN, CO, CR, CU, CZ, DE, DK, DM, DO, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IR, IS, JP, KE, KG, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LU, LY, MA, MD, ME, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PA, PE, PG, PH, PL, PT, QA, RO, RS, RU, RW, SA, SC, SD, SE, SG, SK, SL, SM, ST, SV, SY, TH, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW.

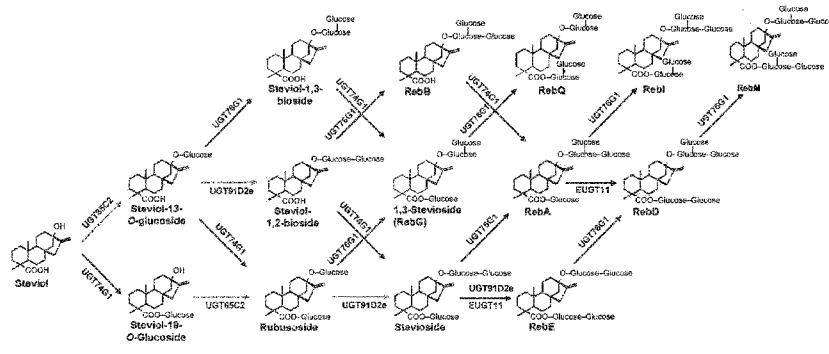
- (84) **Designated States** (unless otherwise indicated, for every kind of regional protection available): ARIPO (BW, GH, GM, KE, LR, LS, MW, MZ, NA, RW, SD, SL, ST, SZ, TZ, UG, ZM, ZW), Eurasian (AM, AZ, BY, KG, KZ, RU, TJ, TM), European (AL, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HR, HU, IE, IS, IT, LT, LU, LV, MC, MK, MT, NL, NO, PL, PT, RO, RS, SE, SI, SK, SM, TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, KM, ML, MR, NE, SN, TD, TG).

**Published:**

- with international search report (Art. 21(3))
- before the expiration of the time limit for amending the claims and to be republished in the event of receipt of amendments (Rule 48.2(h))
- with sequence listing part of description (Rule 5.2(a))

(54) **Title:** PRODUCTION OF STEVIOL GLYCOSIDES IN RECOMBINANT HOSTS

Figure 1



(57) **Abstract:** The invention relates to recombinant microorganisms and methods for producing steviol glycosides and steviol glycoside precursors.

WO 2016/023844 A1

## PRODUCTION OF STEVSOL GLYCOSIDES IN RECOMBINANT HOSTS

### BACKGROUND OF THE INVENTION

#### Field of the Invention

**[0001]** This disclosure relates generally to the recombinant production of steviol glycosides such as rebaudioside A (RebA), rebaudioside B (RebB), rebaudioside D (RebD), and rebaudioside M (RebM) by recombinant hosts such as recombinant microorganisms and isolation methods thereof. In particular, this disclosure relates to modifications to transport systems in a recombinant host to increase production of such steviol glycosides and/or transport of such steviol glycosides into the culture medium.

#### Description of Related Art

**[0002]** Sweeteners are well known as ingredients used most commonly in the food, beverage, or confectionary industries. The sweetener can either be incorporated into a final food product during production or for stand-alone use, when appropriately diluted, as a tabletop sweetener or an at-home replacement for sugars in baking. Sweeteners include natural sweeteners such as sucrose, high fructose corn syrup, molasses, maple syrup, and honey and artificial sweeteners such as aspartame, saccharine, and sucralose. Stevia extract is a natural sweetener that can be isolated and extracted from a perennial shrub, *Stevia rebaudiana*. Stevia is commonly grown in South America and Asia for commercial production of stevia extract. Stevia extract, purified to various degrees, is used commercially as a high intensity sweetener in foods and in blends or alone as a tabletop sweetener.

**[0003]** Chemical structures for several steviol glycosides are shown in Figure 1, including the diterpene steviol and various steviol glycosides. Extracts of the Stevia plant generally comprise rebaudiosides and other steviol glycosides that contribute to the sweet flavor, although the amount of each steviol glycoside often varies, *inter alia*, among different production batches.

**[0004]** As recovery and purification of steviol glycosides from the Stevia plant have proven to be labor intensive and inefficient, there remains a need for a recombinant production system that can produce high yields of desired steviol glycosides, such as RebD and RebM.

## SUMMARY OF THE INVENTION

[0005] It is against the above background that the present invention provides certain advantages and advancements over the prior art.

[0006] In particular, the invention provides a recombinant host capable of synthesizing a steviol glycoside, comprising a gene encoding a transporter polypeptide and/or a gene encoding a transcription factor polypeptide that regulates expression of at least one transporter gene; wherein expression of the gene encoding the transporter polypeptide and/or the gene encoding the transcription factor polypeptide that regulates expression of at least one transporter gene is modified and the recombinant host transports at least a portion of the synthesized steviol glycoside from the host into a culture medium.

[0007] In some aspects of the recombinant host disclosed herein, the gene encoding the transporter polypeptide is an endogenous gene.

[0008] In some aspects of the recombinant host disclosed herein, the transporter polypeptide comprises an ATP-binding cassette (ABC) transporter, a major facilitator superfamily (MFS) transporter, an amino acid/auxin permease (AAP) family transporter, ATPase transporter, a sulfate permease (SulP) family transporter, a lysosomal cystine transporter (LCT) family transporter, a Ca<sup>2+</sup>:cation antiporter (CaCA) family transporter, an amino acid-polyamine-organocation (APC) superfamily transporter, a multidrug/oligosaccharidyl-lipid/polysaccharide (MOP) transporter, a ZRT/IRT-like protein (ZIP) metal transporter family transporter, a mitochondrial protein translocase (MPT) family transporter, a voltage-gated ion channel (VIC) family transporter, a monovalent cation:proton antiporter-2 (CPA2) family transporter, a ThrE family of putative transmembrane amino acid efflux transporter, an oligopeptide transporter (OPT) family transporter, a K<sup>+</sup> transporter (Trk) family transporter, a bile acid:Na symporter (BASS) family transporter, a drug/metabolite transporter (DMT) superfamily transporter, a mitochondrial carrier (MC) family transporter, an auxin efflux carrier (AEC) family transporter, an ammonia channel transporter (Amt) family transporter, a metal ion (Mn<sup>2+</sup>-iron) transporter (Nramp) family transporter, a transient receptor potential Ca<sup>2+</sup> channel (TRP-CC) family transporter, an arsenical resistance-3 (ACR3) family transporter, a nucleobase:cation symporter-1 (NCS1) family transporter, an inorganic phosphate transporter (PiT) family transporter, an arsenite-antimonite (ArsAB) efflux family transporter, an IISP family of transporter, a glycerol uptake (GUP) family transporter, a metal ion transport (MIT) family transporter, a copper transport (Ctr) family or a cation diffusion facilitator (CDF) family transporter.

[0009] In some aspects of the recombinant host disclosed herein, the modified expression comprises modified expression comprises:

- (a) overexpressing the gene encoding the transporter polypeptide and/or the gene encoding the transcription factor polypeptide; or
- (b) deleting the gene encoding the transporter polypeptide and/or the gene encoding the transcription factor polypeptide.

**[0010]** In some aspects of the recombinant host disclosed herein, the gene encoding the transporter polypeptide and/or the gene encoding the transcription factor polypeptide has an activity that is increased.

**[0011]** In some aspects of the recombinant host disclosed herein, one or more of the genes encoding the transporter polypeptide and/or one or more of the genes encoding the transcription factor polypeptide are overexpressed.

**[0012]** In some aspects of the recombinant host disclosed herein, the transporter polypeptide and/or transcription polypeptide comprise YAL067C set forth in SEQ ID NO: 14, YBL089W set forth in SEQ ID NO: 15, YBL099W set forth in SEQ ID NO: 16, YBR008C set forth in SEQ ID NO: 86, YBR021W set forth in SEQ ID NO: 87, YBR043C set forth in SEQ ID NO: 88, YBR180W set forth in SEQ ID NO: 13, YBR241C set forth in SEQ ID NO: 17, YBR287W set forth in SEQ ID NO: 89, YBR294W set forth in SEQ ID NO: 18, YBR295W set forth in SEQ ID NO: 90, YBR296C set forth in SEQ ID NO: 91, YCL038C set forth in SEQ ID NO: 92, YCL069W set forth in SEQ ID NO: 19, YCR01 1C set forth in SEQ ID NO: 93, YCR028C set forth in SEQ ID NO: 20, YCR075C set forth in SEQ ID NO: 21, YDL054C set forth in SEQ ID NO: 94, YDL100C set forth in SEQ ID NO: 95, YDL128W set forth in SEQ ID NO: 22, YDL185W set forth in SEQ ID NO: 23, YDL194W set forth in SEQ ID NO: 24, YDL210W set forth in SEQ ID NO: 25, YDL245C set forth in SEQ ID NO: 96, YDL247W set forth in SEQ ID NO: 97, YDR01 1W set forth in SEQ ID NO: 98, YDR061W set forth in SEQ ID NO: 26, YDR093W set forth in SEQ ID NO: 27, YDR292C set forth in SEQ ID NO: 99, YDR338C set forth in SEQ ID NO: 28, YDR406W set forth in SEQ ID NO: 29, YDR497C set forth in SEQ ID NO: 100, YDR536W set forth in SEQ ID NO: 30, YEL006W set forth in SEQ ID NO: 101, YEL027W set forth in SEQ ID NO: 102, YEL031W set forth in SEQ ID NO: 31, YEL065W set forth in SEQ ID NO: 103, YER019C-A set forth in SEQ ID NO: 104, YER053C set forth in SEQ ID NO: 105, YER1 19C set forth in SEQ ID NO: 106, YER166W set forth in SEQ ID NO: 32, YFL01 1W set forth in SEQ ID NO: 33, YFL028C set forth in SEQ ID NO: 107, YFR045W set forth in SEQ ID NO: 108, YGL006W set forth in SEQ ID NO: 34, YGL013C set forth in SEQ ID NO: 35, YGL084C set forth in SEQ ID NO: 109, YGL104C set forth in SEQ ID NO: 110, YGL1 14W set forth in SEQ ID NO: 111, YGL167C set forth in SEQ ID NO: 112, YGL255W set forth in SEQ ID NO: 36, YGR125W set forth in SEQ ID NO: 37, YGR181W set forth in SEQ ID NO: 38, YGR217W set forth in SEQ ID NO: 39, YGR224W set forth in SEQ ID

NO:40, YGR257C set forth in SEQ ID NO:113, YGR281W set forth in SEQ ID NO:41, YHL016C set forth in SEQ ID NO:42, YHL035C set forth in SEQ ID NO:114, YHL036W set forth in SEQ ID NO:15, YHR002W set forth in SEQ ID NO:16, YHR096C set forth in SEQ ID NO:17, YIL006W set forth in SEQ ID NO:18, YIL088C set forth in SEQ ID NO:43, YIL120W set forth in SEQ ID NO:119, YIL121W set forth in SEQ ID NO:120, YIL166C set forth in SEQ ID NO:121, YJL093C set forth in SEQ ID NO:44, YJL094C set forth in SEQ ID NO:45, YJL108C set forth in SEQ ID NO:46, YJL133W set forth in SEQ ID NO:122, YJL212C set forth in SEQ ID NO:47, YJL219W set forth in SEQ ID NO:123, YJR106W set forth in SEQ ID NO:48, YJR160C set forth in SEQ ID NO:49, YKL016C set forth in SEQ ID NO:124, YKL050C set forth in SEQ ID NO:125, YKL064W set forth in SEQ ID NO:50, YKL120W set forth in SEQ ID NO:126, YKL146W set forth in SEQ ID NO:127, YKL209C set forth in SEQ ID NO:128, YKR039W set forth in SEQ ID NO:129, YKR050W set forth in SEQ ID NO:51, YKR105C set forth in SEQ ID NO:52, YKR106W set forth in SEQ ID NO:53, YLR41 1W set forth in SEQ ID NO:130, YLR447C set forth in SEQ ID NO:54, YML038C set forth in SEQ ID NO:131, YML1 16W set forth in SEQ ID NO:55, YMR034C set forth in SEQ ID NO:56, YMR056C set forth in SEQ ID NO:57, YMR166C set forth in SEQ ID NO:132, YMR253C set forth in SEQ ID NO:58, YMR279C set forth in SEQ ID NO:133, YNL003C set forth in SEQ ID NO:134, YNL065W set forth in SEQ ID NO:59, YNL070W set forth in SEQ ID NO:60, YNL083W set forth in SEQ ID NO:61, YNL095C set forth in SEQ ID NO:62, YNL121C set forth in SEQ ID NO:63, YNL142W set forth in SEQ ID NO:64, YNL268W set forth in SEQ ID NO:135, YNR055C set forth in SEQ ID NO:136, YOL020W set forth in SEQ ID NO:65, YOL075C set forth in SEQ ID NO:66, YOL077W-A set forth in SEQ ID NO:67, YOL122C set forth in SEQ ID NO:68, YOL158C set forth in SEQ ID NO:137, YOR079C set forth in SEQ ID NO:69, YOR087W set forth in SEQ ID NO:70, YOR092W set forth in SEQ ID NO:71, YOR100C set forth in SEQ ID NO:138, YOR130C set forth in SEQ ID NO:72, YOR153W set forth in SEQ ID NO:139, YOR222W set forth in SEQ ID NO:73, YOR271C set forth in SEQ ID NO:140, YOR273C set forth in SEQ ID NO:141, YOR291W set forth in SEQ ID NO:74, YOR306C set forth in SEQ ID NO:75, YOR307C set forth in SEQ ID NO:142, YOR316C set forth in SEQ ID NO:76, YOR332W set forth in SEQ ID NO:143, YOR334W set forth in SEQ ID NO:77, YOR348C set forth in SEQ ID NO:144, YPL036W set forth in SEQ ID NO:145, YPL078C set forth in SEQ ID NO:78, YPL270W set forth in SEQ ID NO:79, YPL274W set forth in SEQ ID NO:80, YPR003C set forth in SEQ ID NO:81, YPR01 1C set forth in SEQ ID NO:82, YPR058W set forth in SEQ ID NO:83, YPR128C set forth in SEQ ID NO:84, or YPR201W set forth in SEQ ID NO:85.

**[0013]** In some aspects of the recombinant host disclosed herein, YBR043C set forth in SEQ ID NO:88, YDL100C set forth in SEQ ID NO:95, YDL054C set forth in SEQ ID NO:94,

YDL128W set forth in SEQ ID NO:22, YDL198C set forth in SEQ ID NO:146, YDR061W set forth in SEQ ID NO:26, YDR536W set forth in SEQ ID NO:30, YEL027W set forth in SEQ ID NO:102, YFL054C set forth in SEQ ID NO:147, YGL167C set forth in SEQ ID NO:112, YGR181W set forth in SEQ ID NO:38, YHL016C set forth in SEQ ID NO:42, YIL166C set forth in SEQ ID NO:121, YJL093C set forth in SEQ ID NO:44, YJR106W set forth in SEQ ID NO:48, YKL120W set forth in SEQ ID NO:126, YKL146W set forth in SEQ ID NO:127, YKR039W set forth in SEQ ID NO:129, YMR034C set forth in SEQ ID NO:56, YMR166C set forth in SEQ ID NO:132, YOL122C set forth in SEQ ID NO:68, YOR079C set forth in SEQ ID NO:69, YPL270W set forth in SEQ ID NO:79, and/or YPR01 1C set forth in SEQ ID NO:82 are overexpressed.

**[0014]** In some aspects, the recombinant host further comprises:

- (a) one or more genes encoding a sucrose transporter and a sucrose synthase;
- (b) a gene encoding a geranylgeranyl diphosphate synthase (GGPPS) polypeptide;
- (c) a gene encoding an ent-copalyl diphosphate synthase (CDPS) polypeptide;
- (d) a gene encoding a kaurene synthase (KS) polypeptide;
- (e) a gene encoding a kaurene oxidase (KO) polypeptide;
- (f) a gene encoding a steviol synthase (KAH) polypeptide;
- (g) a gene encoding a cytochrome P450 reductase (CPR) polypeptide;
- (h) a gene encoding a UGT85C2 polypeptide;
- (i) a gene encoding a UGT76G1 polypeptide;
- (k) a gene encoding a UGT91 D2 functional homolog; and/or
- (l) a gene encoding a EUGT1 1 polypeptide;

wherein at least one of the genes is a recombinant gene; and

wherein the host is capable of producing one or more of RebA, RebB, RebD and/or RebM.

**[0015]** In some aspects of the recombinant host disclosed herein, at least one of the genes is codon optimized for expression in the host.

[0016] In some aspects of the recombinant host disclosed herein, at least one of the genes is codon optimized for expression in *Saccharomyces cerevisiae*.

**[0017]** In some aspects of the recombinant host disclosed herein,

- (a) the GGPPS polypeptide comprises a polypeptide having at least 70% identity to an amino acid sequence set forth in SEQ ID NO: 149;
- (b) the CDPS polypeptide comprises a polypeptide having at least 70% identity to an amino acid sequence set forth in SEQ ID NO: 150;
- (c) the KO polypeptide comprises a polypeptide having at least 70% identity to an amino acid sequence set forth in SEQ ID NO: 152;
- (d) the KS polypeptide comprises a polypeptide having at least 40% identity to an amino acid sequence set forth in SEQ ID NO: 151;
- (e) the KAH polypeptide comprises a polypeptide having at least 60% identity to an amino acid sequence set forth in SEQ ID NO: 154;
- (f) the CPR polypeptide comprises a polypeptide having at least 70% identity to an amino acid sequence set forth in SEQ ID NO: 153 and/or a polypeptide having at least 65% identity to an amino acid sequence set forth in SEQ ID NO: 155;
- (g) the UGT85C2 polypeptide comprises a polypeptide having at least 55% identity to an amino acid sequence set forth in SEQ ID NO: 156;
- (h) the UGT76G1 polypeptide comprises a polypeptide having at least 50% identity to an amino acid sequence set forth in SEQ ID NO: 158;
- (i) the UGT74G1 polypeptide comprises a polypeptide having at least 55% identity to an amino acid sequence set forth in SEQ ID NO: 157;
- (j) the a UGT91D2 functional homolog comprises a UGT91D2e-b polypeptide having at least 90% identity to the amino acid sequence set forth in SEQ ID NO: 159; and
- (k) the EUGT1 1 polypeptide comprises a polypeptide having at least 65% identity to an amino acid sequence set forth in SEQ ID NO: 148.

**[0018]** In some aspects, the recombinant host disclosed herein comprises a microorganism that is a plant cell, a mammalian cell, an insect cell, a fungal cell, or a bacterial cell.



[0019] In some aspects, the bacterial cell comprises *Escherichia* bacteria cells, *Lactobacillus* bacteria cells, *Lactococcus* bacteria cells, *Cornebacterium* bacteria cells, *Acetobacter* bacteria cells, *Acinetobacter* bacteria cells, or *Pseudomonas* bacterial cells.

[0020] In some aspects, the fungal cell is a yeast cell.

[0021] In some aspects, the yeast cell is a cell from *Saccharomyces cerevisiae*, *Schizosaccharomyces pombe*, *Yarrowia lipolytica*, *Candida glabrata*, *Ashbya gossypii*, *Cyberlindnera jadinii*, *Pichia pastoris*, *Kluyveromyces lactis*, *Hansenula polymorpha*, *Candida boidinii*, *Arxula adenivorans*, *Xanthophyllomyces dendrorhous*, or *Candida albicans* species.

[0022] In some aspects, the yeast cell is a *Saccharomycete*.

[0023] In some aspects, the yeast cell is a cell from the *Saccharomyces cerevisiae* species.

[0024] The invention further provides a method of producing a steviol glycoside, comprising:

- (a) growing the recombinant host disclosed herein in a culture medium, under conditions in which the genes comprising recombinant host disclosed herein are expressed,

wherein the steviol glycoside is synthesized by the host; and

- (b) optionally isolating the steviol glycoside.

[0025] In some aspects of the methods disclosed herein, the steviol glycoside is RebA, RebB, RebD, and/or RebM, and wherein:

- (a) RebA is capable of being synthesized in the recombinant host disclosed herein expressing UGT85C2, UGT76G1, UGT74G1, and UGT91D2;
- (b) RebB is capable of being synthesized in the recombinant host disclosed herein expressing UGT85C2, UGT76G1, and UGT91D2;
- (c) RebD is capable of being synthesized in the recombinant host disclosed herein expressing UGT85C2, UGT76G1, UGT74G1, and UGT91D2 and/or EUGT1 1; and
- (d) RebM is capable of being synthesized in the recombinant host disclosed herein expressing UGT85C2, UGT76G1, UGT74G1, and UGT91D2 and/or EUGT1 1.

**[0026]** In some aspects of the methods disclosed herein a gene encoding YBR043C set forth in SEQ ID NO:88, YDL100C set forth in SEQ ID NO:95, YDL054C set forth in SEQ ID NO:94, YDL128W set forth in SEQ ID NO:22, YDL198C set forth in SEQ ID NO:146, YDR061W set forth in SEQ ID NO:26, YDR536W set forth in SEQ ID NO:30, YEL027W set forth in SEQ ID NO:102, YFL054C set forth in SEQ ID NO:147, YGL167C set forth in SEQ ID NO:112, YGR181W set forth in SEQ ID NO:38, YHL016C set forth in SEQ ID NO:42, YIL166C set forth in SEQ ID NO:121, YJL093C set forth in SEQ ID NO:44, YJR106W set forth in SEQ ID NO:48, YKL120W set forth in SEQ ID NO:126, YKL146W set forth in SEQ ID NO:127, YKR039W set forth in SEQ ID NO:129, YMR034C set forth in SEQ ID NO:56, YMR166C set forth in SEQ ID NO:132, YOL122C set forth in SEQ ID NO:68, YOR079C set forth in SEQ ID NO:69, YPL270W set forth in SEQ ID NO:79, and/or YPR011C set forth in SEQ ID NO:82 is overexpressed.

**[0027]** In some aspects of the methods disclosed herein the steviol glycoside is produced at a concentration of between **about** 500 mg/L to about 10,000 mg/L.

**[0028]** The invention further provides a method of increasing production or transport of a steviol glycoside into a culture medium, comprising:

- (a) growing the recombinant host disclosed herein in a culture medium, under conditions in which the genes comprising the host disclosed herein are expressed,

wherein the steviol glycoside is synthesized by the host; and

- (b) optionally isolating the steviol glycoside.

**[0029]** In some aspects of the methods disclosed herein, the steviol glycoside is RebA, RebB, RebD, and/or RebM.

**[0030]** The invention further provides a method increasing production of steviol or a steviol glycoside in a recombinant host, comprising modifying expression of a gene encoding a transporter polypeptide and/or a gene encoding a transcription that regulates expression of at least one transporter gene, wherein the host is capable of transporting at least a portion of the produced steviol or a steviol glycoside from the host into a culture medium.

**[0031]** These and other features and advantages of the present invention will be more fully understood from the following detailed description of the invention taken together with the accompanying claims. It is noted that the scope of the claims is defined by the recitations therein and not by the specific discussion of features and advantages set forth in the present description.

## DESCRIPTION OF DRAWINGS

**[0032]** Figure 1 shows the chemical structures and synthesis pathways for various steviol glycosides.

**[0033]** Figure 2 is a bar graph of the amount ( $\mu\text{M}$ ) of RebA, RebB, RebD, or RebM in the supernatant of a steviol glycoside-producing strain overexpressing transporter genes YGR181W (SEQ ID NO:38) or YDR061W (SEQ ID NO:26), compared to a control steviol glycoside-producing strain. See Example 4.

**[0034]** Figure 3A and Figure 3B are bar graphs of the amount (mg/L) of RebA, RebD, or RebM in the supernatant (Figure 3A) or total culture (Figure 3B) of a YGR181W (SEQ ID NO:38) or YDR061W (SEQ ID NO:26) overexpressing strain, compared to a control steviol glycoside-producing strain. See Example 4.

**[0035]** Figure 4A shows levels of 13-SMG (total levels and supernatant levels;  $\mu\text{M}/\text{OD}_{600}$ ), Figure 4B shows levels of RebA (total levels and supernatant levels;  $\mu\text{M}/\text{OD}_{600}$ ), Figure 4C shows levels of RebB (total levels and supernatant levels;  $\mu\text{M}/\text{OD}_{600}$ ), Figure 4D shows levels of RebD (total levels and supernatant levels;  $\mu\text{M}/\text{OD}_{600}$ ), and Figure 4E shows levels of RebM (total levels and supernatant levels;  $\mu\text{M}/\text{OD}_{600}$ ) in a steviol glycoside-producing *S. cerevisiae* strain with a genomically integrated transporter gene. The genomically integrated transporter genes of Figures 4A-E are YBR043C (SEQ ID NO:88), YEL027W (SEQ ID NO:102), YJL093C (SEQ ID NO:44), YJR106W (SEQ ID NO:48), YMR166C (SEQ ID NO:132), YIL166C (SEQ ID NO:121), YKL120W (SEQ ID NO:126), YDL054C (SEQ ID NO:94), YDL128W (SEQ ID NO:22), YDR536W (SEQ ID NO:30), YGL167C (SEQ ID NO:112), YKL146W (SEQ ID NO:127), YKR039W (SEQ ID NO:129), YOL122C (SEQ ID NO:68), and YPR01 1C (SEQ ID NO:82). See Example 6.

**[0036]** Figure 5A shows supernatant levels of RebA, RebB, RebD, and RebM (in  $\mu\text{M}/\text{OD}_{600}$ ) of a steviol glycoside-producing strain overexpressing YMR166C (SEQ ID NO:132), YEL027W (SEQ ID NO:102), YKL120W (SEQ ID NO:126), YIL166C (SEQ ID NO:121), YJR106W (SEQ ID NO:48), YJL093C (SEQ ID NO:44), and YBR043C (SEQ ID NO:88) by the USER cloning system. Figure 5B shows total levels of RebA, RebB, RebD, and RebM (in  $\mu\text{M}/\text{OD}_{600}$ ) of a steviol glycoside-producing strain overexpressing YMR166C (SEQ ID NO:132), YEL027W (SEQ ID NO:102), YKL120W (SEQ ID NO:126), YIL166C (SEQ ID NO:121), YJR106W (SEQ ID NO:48), YJL093C (SEQ ID NO:44), and YBR043C (SEQ ID NO:88) by the USER cloning system.

**DETAILED DESCRIPTION**

**[0037]** All publications, patents and patent applications cited herein are hereby expressly incorporated by reference in their entirety for all purposes.

**[0038]** Before describing the present invention in detail, a number of terms will be defined. As used herein, the singular forms "a," "an," and "the" include plural referents unless the context clearly dictates otherwise. For example, reference to "a nucleic acid" means one or more nucleic acids.

**[0039]** It is noted that terms like "preferably," "commonly," and "typically" are not utilized herein to limit the scope of the claimed invention or to imply that certain features are critical, essential, or even important to the structure or function of the claimed invention. Rather, these terms are merely intended to highlight alternative or additional features that can or cannot be utilized in a particular embodiment of the present invention.

**[0040]** For the purposes of describing and defining the present invention it is noted that the term "substantially" is utilized herein to represent the inherent degree of uncertainty that can be attributed to any quantitative comparison, value, measurement, or other representation. The term "substantially" is also utilized herein to represent the degree by which a quantitative representation can vary from a stated reference without resulting in a change in the basic function of the subject matter at issue.

**[0001]** Methods well known to those skilled in the art can be used to construct genetic expression constructs and recombinant cells according to this invention. These methods include in vitro recombinant DNA techniques, synthetic techniques, in vivo recombination techniques, and polymerase chain reaction (PCR) techniques. See, for example, techniques as described in Green & Sambrook, 2012, MOLECULAR CLONING: A LABORATORY MANUAL, Fourth Edition, Cold Spring Harbor Laboratory, New York; Ausubel *et al.*, 1989, CURRENT PROTOCOLS IN MOLECULAR BIOLOGY, Greene Publishing Associates and Wiley Interscience, New York, and PCR Protocols: A Guide to Methods and Applications (Innis *et al.*, 1990, Academic Press, San Diego, CA).

**[0041]** As used herein, the terms "polynucleotide," "nucleotide," "oligonucleotide," and "nucleic acid" can be used interchangeably to refer to nucleic acid comprising DNA, RNA, derivatives thereof, or combinations thereof.

**[0042]** As used herein, the terms "microorganism," "microorganism host," "microorganism host cell," "host cell," "recombinant host," "recombinant microorganism host," and "recombinant host cell" can be used interchangeably. As used herein, the term "recombinant host" is intended to refer to a host, the genome of which has been augmented

by at least one DNA sequence. Such DNA sequences include but are not limited to genes that are not naturally present, DNA sequences that are not normally transcribed into RNA or translated into a protein ("expressed"), and other genes or DNA sequences which one desires to introduce into the non-recombinant host. It will be appreciated that typically the genome of a recombinant host described herein is augmented through stable introduction of one or more recombinant genes. Generally, introduced DNA is not originally resident in the host that is the recipient of the DNA, but it is within the scope of this disclosure to isolate a DNA segment from a given host, and to subsequently introduce one or more additional copies of that DNA into the same host, *e.g.*, to enhance production of the product of a gene or alter the expression pattern of a gene. In some instances, the introduced DNA will modify or even replace an endogenous gene or DNA sequence by, *e.g.*, homologous recombination or site-directed mutagenesis. Suitable recombinant hosts include microorganisms.

[0043] As used herein, the term "recombinant gene" refers to a gene or DNA sequence that is introduced into a recipient host, regardless of whether the same or a similar gene or DNA sequence may already be present in such a host. "Introduced," or "augmented" in this context, is known in the art to mean introduced or augmented by the hand of man. Thus, a recombinant gene can be a DNA sequence from another species or can be a DNA sequence that originated from or is present in the same species but has been incorporated into a host by recombinant methods to form a recombinant host. It will be appreciated that a recombinant gene that is introduced into a host can be identical to a DNA sequence that is normally present in the host being transformed and is introduced to provide one or more additional copies of the DNA to thereby permit overexpression or modified expression of the gene product of that DNA. Said recombinant genes are particularly encoded by cDNA.

[0044] As used herein, the term "engineered biosynthetic pathway" refers to a biosynthetic pathway that occurs in a recombinant host, as described herein, and does not naturally occur in the host.

[0045] As used herein, the term "endogenous" gene refers to a gene that originates from and is produced or synthesized within a particular organism, tissue, or cell. In some embodiments, the endogenous gene is a yeast transporter. In some embodiments, the transporter is endogenous to *S. cerevisiae*, including, but not limited to *S. cerevisiae* strain S288C. In some embodiments, an endogenous yeast transporter gene is overexpressed. As used herein, the term "overexpress" is used to refer to the expression of a gene in an organism at levels higher than the level of gene expression in a wild type organism. See, *e.g.*, Prelich, 2012, *Genetics* 190:841-54. In some embodiments, an endogenous yeast transporter gene is deleted. See, *e.g.*, Giaever & Nislow, 2014, *Genetics* 197(2):451-65. As used herein, the terms "deletion," "deleted," "knockout," and "knocked out" can be used

interchangeably to refer to an endogenous gene that has been manipulated to no longer be expressed in an organism, including, but not limited to, *S. cerevisiae*. In some embodiments, a deleted/knocked out gene is a transporter gene or a transcription factor gene that regulates expression of a transporter gene.

[0046] As used herein, the terms "heterologous sequence" and "heterologous coding sequence" are used to describe a sequence derived from a species other than the recombinant host. In some embodiments, the recombinant host is an *S. cerevisiae* cell, and a heterologous sequence is derived from an organism other than *S. cerevisiae*. A heterologous coding sequence, for example, can be from a prokaryotic microorganism, a eukaryotic microorganism, a plant, an animal, an insect, or a fungus different than the recombinant host expressing the heterologous sequence. In some embodiments, a coding sequence is a sequence that is native to the host.

[0047] A "selectable marker" can be one of any number of genes that complement host cell auxotrophy, provide antibiotic resistance, or result in a color change. Linearized DNA fragments of the gene replacement vector then are introduced into the cells using methods well known in the art (see below). Integration of the linear fragments into the genome and the disruption of the gene can be determined based on the selection marker and can be verified by, for example, PGR or Southern blot analysis. Subsequent to its use in selection, a selectable marker can be removed from the genome of the host cell by, e.g., Cre-LoxP systems (see, e.g., Gossen *et al.*, 2002, *Ann. Rev. Genetics* 36:153-173 and U.S. 2006/0014264). Alternatively, a gene replacement vector can be constructed in such a way as to include a portion of the gene to be disrupted, where the portion is devoid of any endogenous gene promoter sequence and encodes none, or an inactive fragment of, the coding sequence of the gene.

[0048] As used herein, the terms "variant" and "mutant" are used to describe a protein sequence that has been modified at one or more amino acids, compared to the wild type sequence of a particular protein.

[0049] As used herein, the term "inactive fragment" is a fragment of the gene that encodes a protein having, e.g., less than about 10% (e.g., less than about 9%, less than about 8%, less than about 7%, less than about 6%, less than about 5%, less than about 4%, less than about 3%, less than about 2%, less than about 1%, or 0%) of the activity of the protein produced from the full-length coding sequence of the gene. Such a portion of a gene is inserted in a vector in such a way that no known promoter sequence is operably linked to the gene sequence, but that a stop codon and a transcription termination sequence are operably linked to the portion of the gene sequence. This vector can be subsequently

linearized in the portion of the gene sequence and transformed into a cell. By way of single homologous recombination, this linearized vector is then integrated in the endogenous counterpart of the gene with inactivation thereof.

[0050] As used herein, the term "steviol glycoside" refers to Rebaudioside A (RebA) (CAS # 58543-16-1), Rebaudioside B (RebB) (CAS # 58543-17-2), Rebaudioside C (RebC) (CAS # 63550-99-2), Rebaudioside D (RebD) (CAS # 63279-13-0), Rebaudioside E (RebE) (CAS # 63279-14-1), Rebaudioside F (RebF) (CAS # 438045-89-7), Rebaudioside M (RebM) (CAS # 1220616-44-3), Rubusoside (CAS # 63849-39-4), Duicoside A (CAS # 64432-06-0), Rebaudioside I (RebI) (MassBank Record: FU000332), Rebaudioside Q (RebQ), 1,2-Stevioside (CAS # 57817-89-7), 1,3-Stevioside (RebG), 1,2-Bioside (MassBank Record: FU000299), 1,3-Bioside, Steviol-13-O-glucoside (13-SMG), Steviol-19-O-glucoside (19-SMG), a tri-glucosylated steviol glycoside, a tetra-glycosylated steviol glycoside, a penta-glucosylated steviol glycoside, a hexa-glucosylated steviol glycoside, a hepta-glucosylated steviol glycoside, di-glucosylated kaurenoic acid, tri-glucosylated kaurenoic acid, di-glucosylated kaurenol, tri-glucosylated kaurenol, and isomers thereof.

[0051] Recombinant steviol glycoside-producing *Saccharomyces cerevisiae* (*S. cerevisiae*) strains are described in WO 2011/153378, WO 2013/022989, WO 2014/122227, and WO 2014/122328, each of which has been incorporated by reference herein in its entirety. See, also, Example 2. Methods of producing steviol glycosides in recombinant hosts, by whole cell bio-conversion, and *in vitro* are also described in WO 2011/153378, WO 2013/022989, WO 2014/122227, and WO 2014/122328.

[0052] In some embodiments, steviol glycosides and/or steviol glycoside precursors are produced *in vivo* through expression of one or more enzymes involved in the steviol glycoside biosynthetic pathway in a recombinant host. For example, a steviol-producing recombinant host expressing one or more of a gene encoding a geranylgeranyl diphosphate synthase (GGPPS) polypeptide, a gene encoding an ent-copalyl diphosphate synthase (CDPS) polypeptide, a gene encoding a kaurene synthase (KS) polypeptide, a gene encoding a kaurene oxidase polypeptide (KO), a gene encoding a steviol synthase (KAH) polypeptide, a gene encoding a cytochrome P450 reductase (CPR) polypeptide, and a gene encoding a UGT polypeptide can produce a steviol glycoside and/or steviol glycoside precursors *in vivo*. See Example 2.

[0053] In some embodiments, a recombinant host comprises a nucleic acid encoding a UGT85C2 polypeptide, a nucleic acid encoding a UGT76G1 polypeptide, a nucleic acid encoding a UGT74G1 polypeptide, a nucleic acid encoding a UGT91D2 polypeptide, and/or a nucleic acid encoding a EUGT1 1 polypeptide. The skilled worker will appreciate that

expression of these genes may be necessary to produce a particular steviol glycoside but that one or more of these genes can be endogenous to the host provided that at least one (and in some embodiments, all) of these genes is a recombinant gene introduced into the microorganism. In a particular embodiment, a steviol-producing recombinant microorganism comprises exogenous nucleic acids encoding UGT85C2, UGT76G1, or UGT91D2 polypeptides. In another particular embodiment, a steviol-producing recombinant microorganism comprises exogenous nucleic acids encoding UGT85C2, UGT76G1, UGT74G1, and UGT91D2 polypeptides. In yet another particular embodiment, a steviol-producing recombinant microorganism comprises exogenous nucleic acids encoding UGT85C2, UGT76G1, UGT74G1, and EUGT11 polypeptides. In yet another particular embodiment, a steviol-producing recombinant microorganism comprises the exogenous nucleic acids encoding UGT85C2, UGT76G1, UGT74G1, UGT91D2 (including *inter alia* 91D2e, 91D2m, 91D2e-b, and functional homologs thereof), and EUGT11 polypeptides. See Example 2.

[0054] In certain embodiments, the steviol glycoside is RebA, RebB, RebD, and/or RebM. RebA can be synthesized in a steviol-producing recombinant microorganism expressing UGT85C2, UGT76G1, UGT74G1, and UGT91D2. RebB can be synthesized in a steviol-producing recombinant microorganism expressing UGT85C2, UGT76G1, and UGT91D2. RebD can be synthesized in a steviol-producing recombinant microorganism expressing UGT85C2, UGT76G1, UGT74G1, and UGT91D2 and/or EUGT11. RebM can be synthesized in a steviol-producing recombinant microorganism expressing UGT85C2, UGT76G1, UGT74G1, and UGT91D2 and/or EUGT11 (see Figure 1, Example 2).

[0055] In some embodiments, steviol glycosides and/or steviol glycoside precursors are produced through contact of a steviol glycoside precursor with one or more enzymes involved in the steviol glycoside pathway *in vitro*. For example, contacting steviol with a UGT polypeptide can result in production of a steviol glycoside *in vitro*. In some embodiments, a steviol glycoside precursor is produced through contact of an upstream steviol glycoside precursor with one or more enzymes involved in the steviol glycoside pathway *in vitro*. For example, contacting ent-kaurenoic acid with a KAH enzyme can result in production of steviol *in vitro*.

[0056] In some embodiments, a steviol glycoside or steviol glycoside precursor is produced by whole cell bioconversion. For whole cell bioconversion to occur, a host cell expressing one or more enzymes involved in the steviol glycoside pathway takes up and modifies a steviol glycoside precursor in the cell; following modification *in vivo*, a steviol glycoside remains in the cell and/or is excreted into the culture medium. For example, a host cell expressing a gene encoding a UGT polypeptide can take up steviol and glycosylate



steviol in the cell; following glycosylation *in vivo*, a steviol glycoside can be excreted into the culture medium. In some embodiments, the cell is permeabilized to take up a substrate to be modified or to excrete a modified product.

[0057] In some embodiments, a steviol glycoside or steviol glycoside precursor composition produced *in vivo*, *in vitro*, or by whole cell bioconversion comprises less contaminants than a stevia extract from, *inter alia*, a stevia plant. Contaminants include plant-derived compounds that contribute to off-flavors. Potential contaminants include pigments, lipids, proteins, phenolics, saccharides, spathulenol and other sesquiterpenes, labdane diterpenes, monoterpenes, decanoic acid, 8,11,14-eicosatrienoic acid, 2-methyloctadecane, pentacosane, octacosane, tetracosane, octadecanol, stigmasterol,  $\beta$ -sitosterol,  $\alpha$ - and  $\beta$ -amyrin, lupeol,  $\beta$ -amyrin acetate, pentacyclic triterpenes, centaureidin, quercetin, epi- $\alpha$ -cadinol, carophyllenes and derivatives, beta-pinene, beta-sitosterol, and gibberellin.

[0058] As used herein, the terms "detectable amount," "detectable concentration," "measurable amount," and "measurable concentration" refer to a level of steviol glycosides measured in AUC,  $\mu\text{M}/\text{OD}_{600}$ , mg/L,  $\mu\text{M}$ , or mM. Steviol glycoside production (*i.e.*, total, supernatant, and/or intracellular steviol glycoside levels) can be detected and/or analyzed by techniques generally available to one skilled in the art, for example, but not limited to, liquid chromatography-mass spectrometry (LC-MS), thin layer chromatography (TLC), high-performance liquid chromatography (HPLC), ultraviolet-visible spectroscopy/spectrophotometry (UV-Vis), mass spectrometry (MS), and nuclear magnetic resonance spectroscopy (NMR).

[0059] As used herein, the terms "or" and "and/or" is utilized to describe multiple components in combination or exclusive of one another. For example, "x, y, and/or z" can refer to "x" alone, "y" alone, "z" alone, "x, y, and z," "(x and y) or z," "x or (y and z)," or "x or y or z." In some embodiments, "and/or" is used to refer to the exogenous nucleic acids that a recombinant cell comprises, wherein a recombinant cell comprises one or more exogenous nucleic acids selected from a group. In some embodiments, "and/or" is used to refer to production of steviol glycosides and/or steviol glycoside precursors. In some embodiments, "and/or" is used to refer to production of steviol glycosides, wherein one or more steviol glycosides are produced. In some embodiments, "and/or" is used to refer to production of steviol glycosides, wherein one or more steviol glycosides are produced through one or more of the following steps: culturing a recombinant microorganism, synthesizing one or more steviol glycosides in a recombinant microorganism, and/or isolating one or more steviol glycosides.

### **Transporters and Transcription Factor Expression**

**[0060]** This document describes reagents and methods that can be used to efficiently produce steviol glycoside compositions. Modification of transport systems in a recombinant host that are involved in transport of steviol glycosides into culture medium can **allow** more effective production of steviol glycosides in recombinant hosts.

**[0061]** As set forth herein, recombinant cells having modifications to cellular transport are capable of producing steviol. Recombinant hosts described herein can produce steviol and have altered expression of at least one endogenous transporter gene. Recombinant hosts described herein can produce steviol and have altered expression of a transcription factor that regulates expression of at least one endogenous transporter gene. Altering expression of endogenous transporter genes can be useful for increasing production of steviol and/or excretion of steviol into the culture medium.

**[0062]** As set forth herein, recombinant cells having modifications to cellular transport are capable of producing at least one steviol glycoside, including, but not limited to, RebA, RebB, RebD, and/or RebM. Recombinant hosts described herein can produce at least one steviol glycoside such as RebA, RebB, RebD, and/or RebM and have altered expression of at least one endogenous transporter gene. Recombinant hosts described herein can produce at least one steviol glycoside such as RebA, RebB, RebD, and/or RebM and have altered expression of a transcription factor that regulates expression of at least one endogenous transporter gene. Recombinant hosts described herein can produce at least one steviol glycoside such as RebA, RebB, RebD, and/or RebM and have altered expression of a plurality of endogenous transporter genes and/or of a plurality of transcription factor genes that regulate expression of a plurality of endogenous transporter genes. Altering expression of endogenous transporter genes and/or transcription factors regulating expression of at least one transporter gene can be useful for increasing production of steviol glycosides and/or excretion of steviol glycosides into the culture medium.

**[0063]** Recombinant hosts disclosed herein can include one or more biosynthesis genes, such as one or more genes encoding a sucrose transporter and a sucrose synthase; a gene encoding a geranylgeranyl diphosphate synthase (GGPPS) polypeptide; a gene encoding an ent-copalyl diphosphate synthase (CDPS) polypeptide; a gene encoding a kaurene synthase (KS) polypeptide; a gene encoding a kaurene oxidase (KO) polypeptide; a gene encoding a steviol synthase (KAH) polypeptide; a gene encoding a cytochrome P450 reductase (CPR) polypeptide; a gene encoding a UGT85C2 polypeptide; a gene encoding a

UGT76G1 polypeptide; a gene encoding a UGT74G1 polypeptide; a gene encoding a UGT91D2 functional homolog; and/or a gene encoding a EUGT11 polypeptide; wherein expression of one or more of these genes results in production of steviol glycosides such as RebA, RebB, RebD, and/or RebM.

[0064] As used herein, the terms "transport of a steviol glycoside," "steviol glycoside transport," "excretion of a steviol glycoside," and "steviol glycoside excretion" can be used interchangeably.

[0065] As used herein, the term "transporter" (also referred to as a membrane transport protein) refers to a membrane protein involved in the movement of small molecules, macromolecules (such as carbohydrates), and ions across a biological membrane. Transporters span the membrane in which they are localized and across which they transport substances. Transporter proteins can assist in the movement (*i.e.*, transport or excretion) of a substance from the intracellular space to the culture medium. Transporters are known to function as passive transport systems, carrying molecules down their concentration gradient, or as active transport systems, using energy to carry molecules uphill against their concentration gradient. Active transport is mediated by carriers which couple transport directly to the use of energy derived from hydrolysis of an ATP molecule or by carriers which make use of a pre-established electrochemical ion gradient to drive co-transport of the nutrient molecule and a co-transported ion. The latter category comprises symporters and antiporters, which carry the ion in the same or opposite direction, respectively, as the transported substrate.

[0066] Transport proteins have been classified according to various criteria at the Transporter Classification Database (on the world wide web at [tcdb.org](http://tcdb.org)). See, Saier Jr. *et al.*, Nucl. Acids Res., 42(1):D251-258 (2014). Non-limiting examples thereof include, among others, the family of Multiple Drug Resistance (MDR) plasma membrane transporters that is thought to be ubiquitous among living organisms. The MDR transporter superfamily can be further subdivided according to the mode of operation by which the substrate is transported from one side of the membrane to the other. Transporters can operate to move substances across membranes in response to chemiosmotic ion gradients or by active transport. ATP-binding cassette transporters (ABC transporters) are transmembrane proteins that utilize the energy of adenosine triphosphate (ATP) hydrolysis to carry out translocation of various substrates across membranes. They can transport a wide variety of substrates across the plasma membrane and intracellular membranes, including metabolic products, lipids and sterols, and drugs. Particular non-limiting examples of endogenous ABC transporter genes include PDR5, YDR061W, PDR15, SNQ2, YOR1, YOL075C, MDL2, ADP1, CAF16, VMR1

and STE6 (or a functional homolog thereof). In some aspects, ABC transporters transport steviol glycosides.

[0067] A second group of MDRs is further subdivided based on the nature of the chemiosmotic gradient that facilitates the transport. Saier, Jr. *et al.*, J. Mol. Microbiol. Biotechnol. 1:257-279 (1999). In some aspects, MDR transporters transport steviol glycosides.

[0068] Another transporter family, the Major Facilitator Superfamily (MFS) transporters are monomeric polypeptides that can transport small solutes in response to proton gradients. The MFS transporter family is sometimes referred to as the uniporter-symporter-antiporter family. MFS transporters function in, *inter alia*, in sugar uptake and drug efflux systems. MFS transporters typically comprise conserved MFS-specific motifs. Non-limiting examples of endogenous MFS transporter genes include DTR1, SE01, YBR241C, VBA3, FEN2, SNF3, STL1, HXT10, AZR1, MPH3, VBA5, GEX2, SNQ1, AQR1, MCH1, MCH5, ATG22, HXT15, MPH2, ITR1, SIT1, VPS73, HXT5, QDR1, QDR2, QDR3, SOA1, HXT9, YMR279C, YIL166C, HOL1, ENB1, TP04 and FLR1 (or a functional homolog thereof). In some aspects, MFS transporters transport steviol glycosides.

[0069] Other transporter families include the SMR (small multidrug resistant) family, RND (Resistance-Nodulation-Cell Division) family, and the MATE (multidrug and toxic compound extrusion) family. The SMR family members are integral membrane proteins characterized by four alpha-helical transmembrane strands that confer resistance to a broad range of antiseptics, lipophilic quaternary ammonium compounds (QAC), and aminoglycoside resistance in bacteria. See, Bay & Turner, 2009, BMC Evol Biol., 9:140. In some aspects, SMR transporters transport steviol glycosides.

[0070] The MATE family members comprise 12 transmembrane (TM) domains. Members of the MATE family have been identified in prokaryotes, yeast such as *S. cerevisiae* and *Schizosaccharomyces pombe*, and plants. See Diener *et al.*, 2001, Plant Cell. 13(7):1625-8. The MATE family members are sodium or proton antiporters. In some aspects, MATE transporters transport steviol glycosides.

[0071] Additional transporter families include the amino acid/auxin permease (AAP) family (for example, YKL146W/AVT3, YBL089W/AVT5, YER119C/AVT6 and YIL088C/AVT7), the ATPase family (for example, YBL099W/ATP1, YDL185W/VMA1, YLR447C/VMA6, YOL077W/ATP19, YPL078C/ATP4, YEL027W/VMA3, YKL016C/ATP7, and YOR332W/VMA4), the sulfate permease (SuIP) family (for example, YBR294W/SUL1, YGR125W and YPR003C), the lysosomal cystine transporter (LCT) family (for example, YCR075C/ERS1), the Ca<sup>2+</sup>:cation antiporter (CaCA) family (for example, YDL128W/VCX1

and YJR106W/ECM27), the amino acid-polyamine-organocation (APC) superfamily (for example, YDL210W/UGA4, YOL020W/TAT2, YPL274W/SAM3, YNL268W/LYP1, YHL036W/MUP3, YKR039W/GAP1 and YOR348C/PUT4), multidrug/oligosaccharidylipid/polysaccharide (MOP) (for example, YDR338C), the ZRT/IRT-like protein (ZIP) metal transporter family (for example, YGL225W/ZRT1 and YOR079C/ATX2), the mitochondrial protein translocase (MPT) family (for example, YGR181W/TIM13, YNL070W/TOM7, YNL121C/TOM70, the voltage-gated ion channel (VIC) family (for example, YGR217W/CCH1 and YJL093C/TOK1), the monovalent cation:proton antiporter-2 (CPA2) family (for example, YJL094C/KHA1), the ThrE family of putative transmembrane amino acid efflux transporters (for example, YJL108C/PRM10), the oligopeptide transporter (OPT) family (for example, YJL212C/OPT1 and YGL114W), the K<sup>+</sup> transporter (Trk) family (for example, TKR050W/TRK2), the bile acid:Na symporter (BASS) family (for example, YMR034C), the drug/metabolite transporter (DMT) superfamily (for example, YMR253C, YML038C/ YMD8, and YOR307C/SLY41), the mitochondrial carrier (MC) family (for example, YMR056C/AAC1, YNL083W/SAL1, YOR130C/ORT1, YOR222W/ODC2, YPR011C, YPR058W/YMC1, YPR128C/ANT1, YEL006W/YEA6, YER053C/PIC2, YFR045W, YGR257C/MTM1, YHR002W/LEU5, YIL006W/YIA6, YJL133W/MRS3, YKL120W/OAC1, YMR166C, YNL003C/PET8 and YOR100C/CRC1), the auxin efflux carrier (AEC) family (for example, YNL095C, YOR092W/ECM3 and YBR287W), the ammonia channel transporter (Amt) family (for example, YNL142W/MEP2), the metal ion (Mn<sup>2+</sup>-iron) transporter (Nramp) family (for example, YOL122C/SMF1), the transient receptor potential Ca<sup>2+</sup> channel (TRP-CC) family (for example, YOR087W/YVC1), the arsenical resistance-3 (ACR3) family (for example, YPR201W/ARR3), the nucleobasexation symporter-1 (NCS1) family (for example, YBR021W/FUR4), the inorganic phosphate transporter (PiT) family (for example, YBR296C/PH089), the arsenite-antimonite (ArsAB) efflux family (for example, YDL100C/GET3), the MSP family of transporters, the glycerol uptake (GUP) family (for example, YGL084C/GUP1), the metal ion transport (MIT) family (for example, YKL064W/MNR2, YKL050C and YOR334W/MRS2), the copper transport (Ctr) family (for example, YLR411W/CTR3) and the cation diffusion facilitator (CDF) family (for example, YOR316C/COT1). Particular members of any of these transporter families are included within the scope of the disclosed invention to the extent that altered expression in a cell capable of producing steviol glycoside increases production of said steviol glycoside from the cell; exemplary members are disclosed above and in Tables 5, 6, and 14.

**[0072]** As used herein, the term "transcription factor" refers to a DNA-binding protein that regulates gene expression. Preferably, the transcription factor regulates expression of at least one transporter gene.

[0073] Methods for identifying a gene affecting production or transport of steviol glycosides and steviol glycoside pathway intermediates are disclosed herein. Such methods can involve inactivating at least one endogenous transporter gene or modifying expression of at least one transporter gene. Typically, a library of mutant microorganisms is prepared, each mutant in the library having a different endogenous transporter gene inactivated. Methods of inactivating genes and determining their effect in a microorganism are known to a person having ordinary skill in the art; additional methods are disclosed in WO 2014/122328, the disclosure of which is incorporated by reference in its entirety. The mutant microorganisms comprising one or more steviol glycoside pathway genes are cultured in a medium under conditions in which steviol or a steviol glycoside is synthesized, and the amount of total, supernatant, and/or intracellular steviol glycosides produced by the microorganism is measured (e.g., using LC-MS) as described herein.

[0074] The disclosure is directed to recombinant host cells in which expression of endogenous transporter or transcription factor genes is modified. In some embodiments, the transporter or transcription factor gene is endogenous to *S. cerevisiae*, including, but not limited to *S. cerevisiae* strain S288C. In some embodiments, expression of an endogenous transporter or transcription factor can be modified by replacing the endogenous promoter with a different promoter that results in increased expression of the transporter protein (e.g., at least a 5% increase in expression, such as at least a 10%, 15%, 20%, 25%, 30%, 35%, 40%, 45%, or 50%, 100%, 200% increase or more in expression). For example, an endogenous promoter can be replaced with a constitutive or inducible promoter that results in increased expression of the transporter. Homologous recombination can be used to replace the promoter of an endogenous gene with a different promoter that results in increased expression of the transporter. In other embodiments, the inducible or constitutive promoter and endogenous transporter or transcription factor can be integrated into another locus of the genome using homologous recombination. In other embodiments, the transporter or transcription factor gene can be introduced into a microorganism using exogenous plasmids with a promoter that results in overexpression of the transporter or transcription factor in the microorganism. In yet another embodiment, the exogenous plasmids may also comprise multiple copies of the transporter or transcription factor gene. In a further embodiment, the endogenous transporter or transcription factor can be induced to be overexpressed using native mechanisms to the recombinant microorganism (e.g. heat shock, stress, heavy metal, or antibiotic exposure). In yet a further embodiment, the activity of an endogenous gene product is enhanced or increased (for example, by mutation). In yet another embodiment, a homologous or orthologous gene of an endogenous yeast transporter or transcription factor gene is overexpressed.

**[0075]** In certain other embodiments, modified expression of a target gene in a recombinant microorganism comprises overexpressing a transporter gene and/or a transcription factor gene involved in expression of said transporter gene. In yet other embodiments, a plurality of endogenous transporter genes or transcription factor genes is overexpressed in said recombinant microorganism.

**[0076]** Modification of transcription factor expression can be used to increase transporter expression. For example, yeast transcription factor PDR1 regulates expression of the genes encoding ABC transporters PDR5, SNQ2 and YOR1. Therefore, in some embodiments, promoters for the endogenous PDR1 locus can be replaced with a different promoter that results in increased expression of the transcription factors, which can increase production of endogenous transporters.

**[0077]** In some embodiments, the transporter gene or transcription factor gene is (using Uniprot Ordered Locus Name for each): YAL067C, YBL089W, YBL099W, YBR008C, YBR021W, YBR043C, YBR180W, YBR241C, YBR287W, YBR294W, YBR295W, YBR296C, YCL038C, YCL069W, YCR01 1C, YCR028C, YCR075C, YDL054C, YDL100C, YDL128W, YDL185W, YDL194W, YDL210W, YDL245C, YDL247W, YDR01 1W, YDR061W, YDR093W, YDR292C, YDR338C, YDR406W, YDR497C, YDR536W, YEL006W, YEL027W, YEL031W, YEL065W, YER019C-A, YER053C, YER1 19C, YER166W, YFL01 1W, YFL028C, YFR045W, YGL006W, YGL013C, YGL084C, YGL104C, YGL1 14W, YGL167C, YGL255W, YGR125W, YGR181W, YGR217W, YGR224W, YGR257C, YGR281W, YHL016C, YHL035C, YHL036W, YHR002W, YHR096C, YIL006W, YIL088C, YIL120W, YIL121W, YIL166C, YJL093C, YJL094C, YJL108C, YJL133W, YJL212C, YJL219W, YJR106W, YJR160C, YKL016C, YKL050C, YKL064W, YKL120W, YKL146W, YKL209C, YKR039W, YKR050W, YKR105C, YKR106W, YLR41 1W, YLR447C, YML038C, YML1 16W, YMR034C, YMR056C, YMR166C, YMR253C, YMR279C, YNL003C, YNL065W, YNL070W, YNL083W, YNL095C, YNL121C, YNL142W, YNL268W, YNR055C, YOL020W, YOL075C, YOL077W-A, YOL122C, YOL158C, YOR079C, YOR087W, YOR092W, YOR100C, YOR130C, YOR153W, YOR222W, YOR271C, YOR273C, YOR291W, YOR306C, YOR307C, YOR316C, YOR332W, YOR334W, YOR348C, YPL036W, YPL078C, YPL270W, YPL274W, YPR003C, YPR01 1C, YPR058W, YPR128C, and/or YPR201W. SEQ ID NOs, Uniprot Accession Numbers, and gene names for each Ordered Locus can be found in Tables 5, 6, and 14. In some embodiments, the above transporter genes and transcription factor genes regulate excretion of steviol glycosides.

**[0078]** In some embodiments, deletion in a steviol glycoside-producing strain of YDL128W (SEQ ID NO:22), YDL194W (SEQ ID NO:24), YDL210W (SEQ ID NO:25), YDR536W (SEQ ID NO:30), YFL01 1W (SEQ ID NO:33), YGL006W (SEQ ID NO:34),

YGL013C (SEQ ID NO:35), YGL255W (SEQ ID NO:36), YGR181W (SEQ ID NO:38), YGR217W (SEQ ID NO:39), YHL016C (SEQ ID NO:42), Y1L088C (SEQ ID NO:43), YJL094C (SEQ ID NO:45), YJR106W (SEQ ID NO:48), YKR050W (SEQ ID NO:51), YNL065W (SEQ ID NO:59), YNL083W (SEQ ID NO:61), YNL121C (SEQ ID NO:63), YNL142W (SEQ ID NO:64), YOR291W (SEQ ID NO:74), YOR306C (SEQ ID NO:75), YOR334W (SEQ ID NO:77), YPL270W (SEQ ID NO:79), YPR01 1C (SEQ ID NO:82), YPR128C (SEQ ID NO:84) results in a measurable decrease of RebD excreted into the culture medium, indicating that each plays a role in RebD excretion. See Example 3 and Tables 7-10.

**[0079]** In some embodiments, deletion in a steviol glycoside-producing strain of YBR180W (SEQ ID NO:13), YAL067C (SEQ ID NO:14), YBR241C (SEQ ID NO:17), YCL069W (SEQ ID NO:19), YCR075C (SEQ ID NO:21), YDL128W (SEQ ID NO:22), YDL194W (SEQ ID NO:24), YDR093W (SEQ ID NO:27), YDR338C (SEQ ID NO:28), YDR406W (SEQ ID NO:29), YER166W (SEQ ID NO:32), YFL01 1W (SEQ ID NO:33), YGL006W (SEQ ID NO:34), YGL013C (SEQ ID NO:35), YGL255W (SEQ ID NO:36), YGR217W (SEQ ID NO:39), YHL016C (SEQ ID NO:42), YJL094C (SEQ ID NO:45), YJL212C (SEQ ID NO:47), YJR106W (SEQ ID NO:48), YJR160C (SEQ ID NO:49), YKR050W (SEQ ID NO:51), YKR106W (SEQ ID NO:53), YML1 16W (SEQ ID NO:55), YMR034C (SEQ ID NO:56), YMR056C (SEQ ID NO:57), YMR253C (SEQ ID NO:58), YNL070W (SEQ ID NO:60), YNL083W (SEQ ID NO:61), YNL095C (SEQ ID NO:62), YNL121C (SEQ ID NO:63), YOL075C (SEQ ID NO:66), YOL122C (SEQ ID NO:68), YOR087W (SEQ ID NO:70), YOR222W (SEQ ID NO:73), YOR291W (SEQ ID NO:74), YOR306C (SEQ ID NO:75), YPL274W (SEQ ID NO:80), YPR003C (SEQ ID NO:81), YPR01 1C (SEQ ID NO:82), or YPR201W (SEQ ID NO:85) results in a measurable decrease of RebM, indicating that each plays a role in RebM excretion. See Example 3 and Tables 7-10.

**[0080]** In some embodiments, overexpression of YGR181W (SEQ ID NO:38) or YDR061W (SEQ ID NO:26) improves RebD and RebM transport into the culture medium by approximately 2-fold (-400-500 mg/L of supernatant RebD and RebM in YGR181W (SEQ ID NO:38) and YDR061W (SEQ ID NO:26) overexpression strains versus -250 mg/L of supernatant RebD and RebM in a control steviol glycoside-producing strain). See Example 4, Figure 2, and Figure 3.

**[0081]** In some embodiments, overexpression of a transporter of Table 11 increases excretion of RebA, RebB, RebD, and/or RebM by at least 20%. In some embodiments, overexpression of a transporter of Table 12 increases production of RebA, RebB, RebD, and/or RebM by at least 40%. See Example 5.



[0082] In some embodiments, a transporter gene is integrated into the genome of a steviol glycoside-producing host. In some embodiments, the integrated transporter is YBR043C (SEQ ID NO:88), YEL027W (SEQ ID NO:102), YJL093C (SEQ ID NO:44), YJR106W (SEQ ID NO:48), YMR166C (SEQ ID NO:132), YIL166C (SEQ ID NO:121), YKL120W (SEQ ID NO:126), YDL054C (SEQ ID NO:94), YDL128W (SEQ ID NO:22), YDR536W (SEQ ID NO:30), YGL167C (SEQ ID NO:112), YKL146W (SEQ ID NO:127), YKR039W (SEQ ID NO:129), YOL122C (SEQ ID NO:68), or YPR011C (SEQ ID NO:82). In some embodiments, integration of YBR043C (SEQ ID NO:88), YEL027W (SEQ ID NO:102), YJL093C (SEQ ID NO:44), YJR106W (SEQ ID NO:48), YKL120W (SEQ ID NO:126), or YMR166C (SEQ ID NO:132) improves excretion and/or total production of 13-SMG. In some embodiments, integration of YBR043C (SEQ ID NO:88), YEL027W (SEQ ID NO:102), or YMR166C (SEQ ID NO:132) improves excretion and/or total production of RebA. In some embodiments, integration of YBR043C (SEQ ID NO:88), YEL027W (SEQ ID NO:102), or YMR166C (SEQ ID NO:132) improves excretion and/or total production of RebB. In some embodiments, integration of YBR043C of SEQ ID NO:88, YEL027W of SEQ ID NO:102, YJL093C of SEQ ID NO:44, YJR106W of SEQ ID NO:48, and YMR166C of SEQ ID NO:132 improves excretion and/or total production of RebD, and YBR043C of SEQ ID NO:88, YEL027W of SEQ ID NO:102, YIL166C (SEQ ID NO:121), YJL093C of SEQ ID NO:44, YJR106W of SEQ ID NO:48, and YMR166C of SEQ ID NO:132 improves excretion and/or total production of RebM, as measured by an increase in RebD and RebM levels in the supernatant compared to a control steviol glycoside-producing strain. See Example 6.

[0083] In some embodiments, steviol glycoside-producing *S. cerevisiae* strains overexpressing YJL093C (SEQ ID NO:44) or YBR043C (SEQ ID NO:88) produce higher levels of RebD + RebM, compared to a steviol glycoside-producing *S. cerevisiae* strain that does not overexpress YJL093C or YBR043C. See Example 7.

[0084] In some embodiments, a transporter that is knocked out can also have specificity for transport of larger molecular weight steviol glycosides (for example, RebD and the knockout of YGR181W of SEQ ID NO:38 or YOR291W of SEQ ID NO:74), and therefore, can be useful to overexpress in strains where transport of RebD into the culture medium is desired. With appropriate balancing of the rate of glycosylation activity through expression of pathway UGTs, smaller molecular weight steviol glycosides are further glycosylated before they are transported into the culture medium. For example, higher expression levels of a UGT76G1 and UGT91D2e and/or EUGT11, as compared to the UGT74G1 and UGT85C2 enzymes, can prevent accumulation of the steviol monoglucosides that are transported more readily. If the UGT activity level is higher (so the glycosylation rate is

faster) than the rate of transport, then greater amounts of larger molecular weight steviol glycosides will be produced.

### Steviol and Steviol Glycoside Biosynthesis Nucleic Acids

[0085] A recombinant gene encoding a polypeptide described herein comprises the coding sequence for that polypeptide, operably linked in sense orientation to one or more regulatory regions suitable for expressing the polypeptide. Because many microorganisms are capable of expressing multiple gene products from a polycistronic mRNA, multiple polypeptides can be expressed under the control of a single regulatory region for those microorganisms, if desired. A coding sequence and a regulatory region are considered to be operably linked when the regulatory region and coding sequence are positioned so that the regulatory region is effective for regulating transcription or translation of the sequence. Typically, the translation initiation site of the translational reading frame of the coding sequence is positioned between one and about fifty nucleotides downstream of the regulatory region for a monocistronic gene.

[0086] In many cases, the coding sequence for a polypeptide described herein is identified in a species other than the recombinant host, *i.e.*, is a heterologous nucleic acid. Thus, if the recombinant host is a microorganism, the coding sequence can be from other prokaryotic or eukaryotic microorganisms, from plants or from animals. In some case, however, the coding sequence is a sequence that is native to the host and is being reintroduced into that organism. A native sequence can often be distinguished from the naturally occurring sequence by the presence of non-natural sequences linked to the exogenous nucleic acid, e.g., non-native regulatory sequences flanking a native sequence in a recombinant nucleic acid construct. In addition, stably transformed exogenous nucleic acids typically are integrated at positions other than the position where the native sequence is found. "Regulatory region" refers to a nucleic acid having nucleotide sequences that influence transcription or translation initiation and rate, and stability and/or mobility of a transcription or translation product. Regulatory regions include, without limitation, promoter sequences, enhancer sequences, response elements, protein recognition sites, inducible elements, protein binding sequences, 5' and 3' untranslated regions (UTRs), transcriptional start sites, termination sequences, polyadenylation sequences, introns, and combinations thereof. A regulatory region typically comprises at least a core (basal) promoter. A regulatory region also may include at least one control element, such as an enhancer sequence, an upstream element or an upstream activation region (UAR). A regulatory region is operably linked to a coding sequence by positioning the regulatory region and the coding

sequence so that the regulatory region is effective for regulating transcription or translation of the sequence. For example, to operably link a coding sequence and a promoter sequence, the translation initiation site of the translational reading frame of the coding sequence is typically positioned between one and about fifty nucleotides downstream of the promoter. A regulatory region can, however, be positioned as much as about 5,000 nucleotides upstream of the translation initiation site, or about 2,000 nucleotides upstream of the transcription start site.

[0087] The choice of regulatory regions to be included depends upon several factors, including, but not limited to, efficiency, selectability, inducibility, desired expression level, and preferential expression during certain culture stages. It is a routine matter for one of skill in the art to modulate the expression of a coding sequence by appropriately selecting and positioning regulatory regions relative to the coding sequence. It will be understood that more than one regulatory region may be present, *e.g.*, introns, enhancers, upstream activation regions, transcription terminators, and inducible elements.

[0088] One or more genes can be combined in a recombinant nucleic acid construct in "modules" useful for a discrete aspect of steviol and/or steviol glycoside production. Combining a plurality of genes in a module, particularly a polycistronic module, facilitates the use of the module in a variety of species. For example, a steviol biosynthesis gene cluster, or a UGT gene cluster, can be combined in a polycistronic module such that, after insertion of a suitable regulatory region, the module can be introduced into a wide variety of species. As another example, a UGT gene cluster can be combined such that each UGT coding sequence is operably linked to a separate regulatory region, to form a UGT module. Such a module can be used in those species for which monocistronic expression is necessary or desirable. In addition to genes useful for steviol or steviol glycoside production, a recombinant construct typically also comprises an origin of replication, and one or more selectable markers for maintenance of the construct in appropriate species.

[0089] It will be appreciated that because of the degeneracy of the genetic code, a number of nucleic acids can encode a particular polypeptide; *i.e.*, for many amino acids, there is more than one nucleotide triplet that serves as the codon for the amino acid. Thus, codons in the coding sequence for a given polypeptide can be modified such that optimal expression in a particular host is obtained, using appropriate codon bias tables for that host (*e.g.*, microorganism). As isolated nucleic acids, these modified sequences can exist as purified molecules and can be incorporated into a vector or a virus for use in constructing modules for recombinant nucleic acid constructs.

**[0090]** In some cases, it is desirable to inhibit one or more functions of an endogenous polypeptide in order to divert metabolic intermediates towards steviol or steviol glycoside biosynthesis. For example, it may be desirable to downregulate synthesis of sterols in a strain in order to further increase steviol or steviol glycoside production, e.g., by downregulating squalene epoxidase. As another example, it may be desirable to inhibit degradative functions of certain endogenous gene products, e.g., glycohydrolases that remove glucose moieties from secondary metabolites or phosphatases as discussed herein. As another example, expression of membrane transporters involved in transport of steviol glycosides can be activated, such that transportation of steviol glycosides is increased. Such regulation can be beneficial in that transportation of steviol glycosides can be increased for a desired period of time during culture of the microorganism, thereby increasing the yield of glycoside product(s) at harvest. In such cases, a nucleic acid that overexpresses the polypeptide or gene product may be included in a recombinant construct that is transformed into the strain. Alternatively, mutagenesis can be used to generate mutants in genes for which it is desired to increase or enhance function.

### **Recombinant Hosts**

**[0091]** Recombinant hosts can be used to express polypeptides for the producing steviol glycosides, including mammalian, insect, plant, and algal cells. A number of prokaryotes and eukaryotes are also suitable for use in constructing the recombinant microorganisms described herein, e.g., gram-negative bacteria, yeast, and fungi. A species and strain selected for use as a steviol glycoside production strain is first analyzed to determine which production genes are endogenous to the strain and which genes are not present. Genes for which an endogenous counterpart is not present in the strain are advantageously assembled in **one** or more recombinant constructs, which are then transformed into the strain in order to supply the missing function(s).

**[0092]** Typically, the recombinant microorganism is grown in a fermenter at a defined temperature(s) for a desired period of time. The constructed and genetically engineered microorganisms provided by the invention can be cultivated using conventional fermentation processes, including, *inter alia*, chemostat, batch, fed-batch cultivations, semi-continuous fermentations such as draw and fill, continuous perfusion fermentation, and continuous perfusion cell culture. Depending on the particular microorganism used in the method, other recombinant genes such as isopentenyl biosynthesis genes and terpene synthase and cyclase genes may also be present and expressed. Levels of substrates and intermediates, e.g., isopentenyl diphosphate, dimethylallyl diphosphate, GGPP, kaurene and kaurenoic

acid, can be determined by extracting samples from culture media for analysis according to published methods.

[0093] Carbon sources of use in the instant method include any molecule that can be metabolized by the recombinant host cell to facilitate growth and/or production of the steviol glycosides. Examples of suitable carbon sources include, but are not limited to, sucrose (e.g., as found in molasses), fructose, xylose, ethanol, glycerol, glucose, cellulose, starch, cellobiose or other glucose-comprising polymer. In embodiments employing yeast as a host, for example, carbon sources such as sucrose, fructose, xylose, ethanol, glycerol, and glucose are suitable. The carbon source can be provided to the host organism throughout the cultivation period or alternatively, the organism can be grown for a period of time in the presence of another energy source, e.g., protein, and then provided with a source of carbon only during the fed-batch phase.

[0094] After the recombinant microorganism has been grown in culture for the desired period of time, steviol and/or one or more steviol glycosides can then be recovered from the culture using various techniques known in the art. In some embodiments, a permeabilizing agent can be added to aid the feedstock entering into the host and product getting out. For example, a crude lysate of the cultured microorganism can be centrifuged to obtain a supernatant. The resulting supernatant can then be applied to a chromatography column, e.g., a C-18 column, and washed with water to remove hydrophilic compounds, followed by elution of the compound(s) of interest with a solvent such as methanol. The compound(s) can then be further purified by preparative HPLC. See *a/so*, WO 2009/140394.

[0095] it will be appreciated that the various genes and modules discussed herein can be present in two or more recombinant hosts rather than a single host. When a plurality of recombinant hosts is used, they can be grown in a mixed culture to produce steviol and/or steviol glycosides.

[0096] Alternatively, the two or more hosts each can be grown in a separate culture medium and the product of the first culture medium, e.g., steviol, can be introduced into second culture medium to be converted into a subsequent intermediate, or into an end product such as, for example, RebA. The product produced by the second, or final host is then recovered. It will also be appreciated that in some embodiments, a recombinant host is grown using nutrient sources other than a culture medium and utilizing a system other than a fermenter.

[0097] Exemplary prokaryotic and eukaryotic species are described in more detail below. However, it will be appreciated that other species can be suitable. For example, suitable species can be in a genus such as *Agaricus*, *Aspergillus*, *Bacillus*, *Candida*,

*Corynebacterium*, *Eremothecium*, *Escherichia*, *Fusarium/Gibberella*, *Kluyveromyces*, *Laetiporus*, *Lentinus*, *Phaffia*, *Phanerochaete*, *Pichia*, *Physcomitreila*, *Rhodoturula*, *Saccharomyces*, *Schizosaccharomyces*, *Sphaceloma*, *Xanthophylomyces* or *Yarrowia*. Exemplary species from such genera include *Lentinus tigrinus*, *Laetiporus sulphureus*, *Phanerochaete chrysosporium*, *Pichia pastoris*, *Cyberlindnera jadinii*, *Physcomitreila patens*, *Rhodoturula giutinis*, *Rhodoturula mucilaginoso*, *Phaffia rhodozyma*, *Xanthophylomyces dendrorhous*, *Fusarium fujikuroi/Gibberella fujikuroi*, *Candida utilis*, *Candida glabrata*, *Candida albicans*, and *Yarrowia lipolytica*.

**[0098]** In some embodiments, a microorganism can be a prokaryote such as *Escherichia coli*.

**[0099]** In some embodiments, a microorganism can be an Ascomycete such as *Gibberella fujikuroi*, *Kluyveromyces lactis*, *Schizosaccharomyces pombe*, *Aspergillus niger*, *Yarrowia lipolytica*, *Ashbya gossypii*, or *S. cerevisiae*.

**[00100]** In some embodiments, a microorganism can be an algal cell such as *Blakeslea trispora*, *Dunaliella salina*, *Haematococcus pluvialis*, *Chlorella sp.*, *Undaria pinnatifida*, *Sargassum*, *Laminaria japonica*, *Scenedesmus almeriensis* species.

**[001 01]** In some embodiments, a microorganism can be a cyanobacterial cell such as *Blakeslea trispora*, *Dunaliella salina*, *Haematococcus pluvialis*, *Chlorella sp.*, *Undaria pinnatifida*, *Sargassum*, *Laminaria japonica*, *Scenedesmus almeriensis*.

#### *Saccharomyces spp.*

**[001 02]** *Saccharomyces* is a widely used chassis organism in synthetic biology, and can be used as the recombinant microorganism platform. For example, there are libraries of mutants, plasmids, detailed computer models of metabolism and other information available for *S. cerevisiae*, allowing for rational design of various modules to enhance product yield. Methods are known for making recombinant microorganisms.

#### *Aspergillus spp.*

**[00103]** *Aspergillus* species such as *A. oryzae*, *A. niger* and *A. sojae* are widely used microorganisms in food production and can also be used as the recombinant microorganism platform. Nucleotide sequences are available for genomes of *A. nidulans*, *A. fumigatus*, *A. oryzae*, *A. clavatus*, *A. flavus*, *A. niger*, and *A. terreus*, allowing rational design and modification of endogenous pathways to enhance flux and increase product yield. Metabolic models have been developed for *Aspergillus*, as well as transcriptomic studies and proteomics studies. *A. niger* is cultured for the industrial production of a number of food

ingredients such as citric acid and gluconic acid, and thus species such as *A. niger* are generally suitable for producing steviol glycosides.

*E. coli*

**[00104]** *E. coli*, another widely used platform organism in synthetic biology, can also be used as the recombinant microorganism platform. Similar to *Saccharomyces*, there are libraries of mutants, plasmids, detailed computer models of metabolism and other information available for *E. coli*, allowing for rational design of various modules to enhance product yield. Methods similar to those described above for *Saccharomyces* can be used to make recombinant *E. coli* microorganisms.

**[00105]** *Agaricus, Gibberella, and Phanerochaete spp.*

**[00106]** *Agaricus, Gibberella, and Phanerochaete* spp. can be useful because they are known to produce large amounts of isoprenoids in culture. Thus, the terpene precursors for producing large amounts of steviol glycosides are already produced by endogenous genes. Thus, modules comprising recombinant genes for steviol glycoside biosynthesis polypeptides can be introduced into species from such genera without the necessity of introducing mevalonate or MEP pathway genes.

*Arxula adenivorans (Blastobotrys adenivorans)*

**[00107]** *Arxula adenivorans* is dimorphic yeast (it grows as budding yeast like the baker's yeast up to a temperature of 42°C, above this threshold it grows in a filamentous form) with unusual biochemical characteristics. It can grow on a wide range of substrates and can assimilate nitrate. It has successfully been applied to the generation of strains that can produce natural plastics or the development of a biosensor for estrogens in environmental samples.

*Yarrowia lipolytica*

**[00108]** *Yarrowia lipolytica* is dimorphic yeast (see *Arxula adenivorans*) and belongs to the family Hemiascomycetes. The entire genome of *Yarrowia lipolytica* is known. *Yarrowia* species is aerobic and considered to be non-pathogenic. *Yarrowia* is efficient in using hydrophobic substrates (e.g. alkanes, fatty acids, oils) and can grow on sugars. It has a high potential for industrial applications and is an oleaginous microorganism. *Yarrowia lipolytica* can accumulate lipid content to approximately 40% of its dry cell weight and is a model organism for lipid accumulation and remobilization. See e.g., Nicaud, 2012, *Yeast* 29(10):409-18; Beopoulos *et al.*, 2009, *Biochimie* 91(6):692-6; Bankar *et al.*, 2009, *Appl Microbiol Biotechnol.* 84(5):847-65.

*Rhodotorula sp.*

**[00109]** *Rhodotorula* is unicellular, pigmented yeast. The oleaginous red yeast, *Rhodotorula glutinis*, has been shown to produce lipids and carotenoids from crude glycerol (Saenge *et al.*, 2011, *Process Biochemistry* 46(1):210-8). *Rhodotorula toruloides* strains have been shown to be an efficient fed-batch fermentation system for improved biomass and lipid productivity (Li *et al.*, 2007, *Enzyme and Microbial Technology* 41:312-7).

*Rhodospiridium toruloides*

**[00110]** *Rhodospiridium toruloides* is oleaginous yeast and useful for engineering lipid-production pathways (See e.g. Zhu *et al.*, 2013, *Nature Commun.* 3:1112; Ageitos *et al.*, 2011, *Applied Microbiology and Biotechnology* 90(4):1219-27).

*Candida boidinii*

**[00111]** *Candida boidinii* is methylotrophic yeast (it can grow on methanol). Like other methylotrophic species such as *Hansenula polymorpha* and *Pichia pastoris*, it provides an excellent platform for producing heterologous proteins. Yields in a multigram range of a secreted foreign protein have been reported. A computational method, IPRO, recently predicted mutations that experimentally switched the cofactor specificity of *Candida boidinii* xylose reductase from NADPH to NADH. See, e.g., Mattanovich *et al.*, 2012, *Methods Mol Biol.* 824:329-58; Khoury *et al.*, 2009, *Protein Sci.* 18(10):2125-38.

*Hansenula polymorpha* (*Pichia angusta*)

**[00112]** *Hansenula polymorpha* is methylotrophic yeast (see *Candida boidinii*). It can furthermore grow on a wide range of other substrates; it is thermo-tolerant and can assimilate nitrate (see also *Kluyveromyces lactis*). It has been applied to producing hepatitis B vaccines, insulin and interferon alpha-2a for the treatment of hepatitis C, furthermore to a range of technical enzymes. See, e.g., Xu *et al.*, 2014, *Virol Sin.* 29(6):403-9.

*Kluyveromyces lactis*

**[00113]** *Kluyveromyces lactis* is yeast regularly applied to the production of kefir. It can grow on several sugars, most importantly on lactose which is present in milk and whey. It has successfully been applied among others for producing chymosin (an enzyme that is usually present in the stomach of calves) for producing cheese. Production takes place in fermenters on a 40,000 L scale. See, e.g., van Ooyen *et al.*, 2006, *FEMS Yeast Res.* 6(3):381-92.

*Pichia pastoris*

**[00114]** *Pichia pastoris* is methylotrophic yeast (see *Candida boidinii* and *Hansenula polymorpha*). It provides an efficient platform for producing foreign proteins. Platform



elements are available as a kit and it is worldwide used in academia for producing proteins. Strains have been engineered that can produce complex human N-glycan (yeast glycans are similar but not identical to those found in humans). See, e.g., Piirainen *et al.*, 2014, *N Biotechnol.* 31(6):532-7.

*Physcomitrella* spp.

**[00115]** *Physcomitrella* mosses, when grown in suspension culture, have characteristics similar to yeast or other fungal cultures. This genera can be used for producing plant secondary metabolites, which can be difficult to produce in other types of cells.

**Steviol Glycoside Compositions**

**[00116]** Steviol glycosides do not necessarily have equivalent performance in different food systems. It is therefore desirable to have the ability to direct the synthesis to steviol glycoside compositions of choice. Recombinant hosts described herein can produce compositions that are selectively enriched for specific steviol glycosides (e.g., RebD) and have a consistent taste profile. Thus, the recombinant hosts described herein can facilitate the production of compositions that are tailored to meet the sweetening profile desired for a given food product and that have a proportion of each steviol glycoside that is consistent from batch to batch. Hosts described herein do not produce the undesired plant by-products found in *Stevia* extracts. Thus, steviol glycoside compositions produced by the recombinant hosts described herein are distinguishable from compositions derived from *Stevia* plants.

**[00117]** The amount of an individual steviol glycoside (e.g., RebA, RebB, RebD, or RebM) produced can be from about 1 mg/L to about 2,800 mg/L, e.g., about 1 to about 10 mg/L, about 3 to about 10 mg/L, about 5 to about 20 mg/L, about 10 to about 50 mg/L, about 10 to about 100 mg/L, about 25 to about 500 mg/L, about 100 to about 1,500 mg/L, or about 200 to about 1,000 mg/L, at least about 1,000 mg/L, at least about 1,200 mg/L, at least about at least 1,400 mg/L, at least about 1,600 mg/L, at least about 1,800 mg/L, or at least about 2,800 mg/L. In some aspects, the amount of an individual steviol glycoside can exceed 2,800 mg/L. The amount of a combination of steviol glycosides (e.g., RebA, RebB, RebD, or RebM) produced can be from about 1 mg/L to about 6,000 mg/L, e.g., about 200 to about 1,500, at least about 2,000 mg/L, at least about 3,000 mg/L, at least about 4,000 mg/L, at least about 5,000 mg/L, or at least about 6,000 mg/L. In some aspects, the amount of a combination of steviol glycosides can exceed 6,000 mg/L. In general, longer culture times will lead to greater amounts of product. Thus, the recombinant microorganism can be cultured for from 1 day to 7 days, from 1 day to 5 days, from 3 days to 5 days, about 3 days, about 4 days, or about 5 days.

**[00118]** It will be appreciated that the various genes and modules discussed herein can be present in two or more recombinant microorganisms rather than a single microorganism. When a plurality of recombinant microorganisms is used, they can be grown in a mixed culture to produce stevioi and/or stevioi glycosides. For example, a first microorganism can comprise one or more biosynthesis genes for producing stevioi and null mutations in a first group of endogenous transporters, while a second microorganism comprises stevioi glycoside biosynthesis genes and null mutations in a second group of endogenous transporters. The product produced by the second, or final microorganism is then recovered, it will also be appreciated that in some embodiments, a recombinant microorganism is grown using nutrient sources other than a culture medium and utilizing a system other than a fermenter.

**[00119]** Alternatively, the two or more microorganisms each can be grown in a separate culture medium and the product of the first culture medium, e.g., stevioi, can be introduced into second culture medium to be converted into a subsequent intermediate, or into an end product such as RebA. The product produced by the second, or final microorganism is then recovered. The microorganisms can have the same or a different group of mutations in endogenous transporters. It will also be appreciated that in some embodiments, a recombinant microorganism is grown using nutrient sources other than a culture medium and utilizing a system other than a fermenter.

**[00120]** Stevioi glycosides do not necessarily have equivalent performance in different food systems. It is therefore desirable to have the ability to direct the synthesis to stevioi glycoside compositions of choice. Recombinant hosts described herein can produce compositions that are selectively enriched for specific stevioi glycosides (e.g., RebD) and have a consistent taste profile. Thus, the recombinant microorganisms described herein can facilitate the production of compositions that are tailored to meet the sweetening profile desired for a given food product and that have a proportion of each stevioi glycoside that is consistent from batch to batch. Microorganisms described herein do not produce the undesired plant byproducts found in *Stevia* extracts. Thus, stevioi glycoside compositions produced by the recombinant microorganisms described herein are distinguishable from compositions derived from *Stevia* plants.

**[00121]** Stevioi glycosides and compositions obtained by the methods disclosed herein can be used to make food products, dietary supplements and sweetener compositions. See, e.g., WO 2011/153378, WO 2013/022989, WO 2014/122227, and WO 2014/122328, each of which has been incorporated by reference in its entirety.

[00122] For example, substantially pure steviol or steviol glycoside such as RebM or RebD can be included in food products such as ice cream, carbonated beverages, fruit juices, yogurts, baked goods, chewing gums, hard and soft candies, and sauces. Substantially pure steviol or steviol glycoside can also be included in non-food products such as pharmaceutical products, medicinal products, dietary supplements and nutritional supplements. Substantially pure steviol or steviol glycosides may also be included in animal feed products for both the agriculture industry and the companion animal industry. Alternatively, a mixture of steviol and/or steviol glycosides can be made by culturing recombinant microorganisms separately, each producing a specific steviol or steviol glycoside, recovering the steviol or steviol glycoside in substantially pure form from each microorganism and then combining the compounds to obtain a mixture comprising each compound in the desired proportion. The recombinant microorganisms described herein permit more precise and consistent mixtures to be obtained compared to current Stevia products. For example, recombinant microorganisms described herein can express transporters specific for transport of a particular rebaudioside into the culture medium. When a transporter is specific for a particular rebaudioside it will enrich the concentration of that compound in the fermentation broth, preventing it from being further reacted to a different compound, and by selectively transporting the rebaudioside into the fermentation broth it will make it easier to recover from the other rebaudiosides and therefore making the process more efficient.

[00123] In another alternative, a substantially pure steviol or steviol glycoside can be incorporated into a food product along with other sweeteners, *e.g.* saccharin, dextrose, sucrose, fructose, erythritol, aspartame, sucralose, monatin, or acesulfame potassium. The weight ratio of steviol or steviol glycoside relative to other sweeteners can be varied as desired to achieve a satisfactory taste in the final food product. See, *e.g.*, U.S. 2007/01 2831 1. In some embodiments, the steviol or steviol glycoside may be provided with a flavor (*e.g.*, citrus) as a flavor modulator.

[00124] Compositions produced by a recombinant microorganism described herein can be incorporated into food products. For example, a steviol glycoside composition produced by a recombinant microorganism can be incorporated into a food product in an amount ranging from about 20 mg steviol glycoside/kg food product to about 1800 mg steviol glycoside/kg food product on a dry weight basis, depending on the type of steviol glycoside and food product. For example, a steviol glycoside composition produced by a recombinant microorganism can be incorporated into a dessert, cold confectionary (*e.g.*, ice cream), dairy product (*e.g.*, yogurt), or beverage (*e.g.*, a carbonated beverage) such that the food product has a maximum of 500 mg steviol glycoside/kg food on a dry weight basis. A steviol

glycoside composition produced by a recombinant microorganism can be incorporated into a baked good (e.g., a biscuit) such that the food product has a maximum of 300 mg steviol glycoside/kg food on a dry weight basis. A steviol glycoside composition produced by a recombinant microorganism can be incorporated into a sauce (e.g., chocolate syrup) or vegetable product (e.g., pickles) such that the food product has a maximum of 1000 mg steviol glycoside/kg food on a dry weight basis. A steviol glycoside composition produced by a recombinant microorganism can be incorporated into a bread such that the food product has a maximum of 160 mg steviol glycoside/kg food on a dry weight basis. A steviol glycoside composition produced by a recombinant microorganism, plant, or plant cell can be incorporated into a hard or soft candy such that the food product has a maximum of 1600 mg steviol glycoside/kg food on a dry weight basis. A steviol glycoside composition produced by a recombinant microorganism, plant, or plant cell can be incorporated into a processed fruit product (e.g., fruit juices, fruit filling, jams, and jellies) such that the food product has a maximum of 1000 mg steviol glycoside/kg food on a dry weight basis.

**[00125]** For example, such a steviol glycoside composition can have from 90-99% RebA and an undetectable amount of stevia plant-derived contaminants, and be incorporated into a food product at from 25-1600 mg/kg, e.g., 100-500 mg/kg, 25-100 mg/kg, 250-1000 mg/kg, 50-500 mg/kg or 500-1000 mg/kg on a dry weight basis.

**[00126]** Such a steviol glycoside composition can be a RebB-enriched composition having greater than 3% RebB and be incorporated into the food product such that the amount of RebB in the product is from 25-1600 mg/kg, e.g., 100-500 mg/kg, 25-100 mg/kg, 250-1000 mg/kg, 50-500 mg/kg or 500-1000 mg/kg on a dry weight basis. Typically, the RebB-enriched composition has an undetectable amount of stevia plant-derived contaminants.

**[00127]** Such a steviol glycoside composition can be a RebD-enriched composition having greater than 3% RebD and be incorporated into the food product such that the amount of RebD in the product is from 25-1600 mg/kg, e.g., 100-500 mg/kg, 25-100 mg/kg, 250-1000 mg/kg, 50-500 mg/kg or 500-1000 mg/kg on a dry weight basis. Typically, the RebD-enriched composition has an undetectable amount of stevia plant-derived contaminants.

**[00128]** Such a steviol glycoside composition can be a RebE-enriched composition having greater than 3% RebE and be incorporated into the food product such that the amount of RebE in the product is from 25-1600 mg/kg, e.g., 100-500 mg/kg, 25-100 mg/kg, 250-1000 mg/kg, 50-500 mg/kg or 500-1000 mg/kg on a dry weight basis. Typically, the

RebE-enriched composition has an undetectable amount of stevia plant-derived contaminants.

**[00129]** Such a steviol glycoside composition can be a RebM-enriched composition having greater than 3% RebM and be incorporated into the food product such that the amount of RebM in the product is from 25-1600 mg/kg, e.g., 100-500 mg/kg, 25-100 mg/kg, 250-1000 mg/kg, 50-500 mg/kg or 500-1000 mg/kg on a dry weight basis. Typically, the RebM-enriched composition has an undetectable amount of stevia plant-derived contaminants.

**[00130]** In some embodiments, a substantially pure steviol or steviol glycoside is incorporated into a tabletop sweetener or "cup-for-cup" product. Such products typically are diluted to the appropriate sweetness level with one or more bulking agents, e.g., maltodextrins, known to those skilled in the art. Steviol glycoside compositions enriched for RebA, RebB, RebD, RebE, or RebM, can be packaged in a sachet, for example, at from 10,000 to 30,000 mg steviol glycoside/kg product on a dry weight basis, for tabletop use.

**[00131]** The invention will be further described in the following examples, which do not limit the scope of the invention described in the claims.

## EXAMPLES

**[00132]** The Examples that follow are illustrative of specific embodiments of the invention, and various uses thereof. They are set forth for explanatory purposes only, and are not to be taken as limiting the invention.

### Example 1. LC-MS Analytical Procedures

**[00133]** The LC-MS methods described here are oriented towards the separation, general detection and potential identification of chemicals of particular masses (*i.e.* steviol glycosides) in the presence of a mixture (*i.e.* culture media). LC-MS analyses were performed on: (A) an UltiMate® 3000-TSQ (Thermo Fisher Scientific); (B) a 1290 Infinity - 6130SQ (Agilent); or (C) an Acquity -XevoTQD (Waters) system. Specific methods used for each system are described below.

**[00134]** Method A: LC-MS analyses were performed using an UltiMate® 3000 UPLC system (Dionex) fitted with a waters ACQUITY UPLC® BEH shield RP18 column (2.1 x 50 mm, 1.7 µm particles, 130 Å pore size) connected to a TSQ Quantum® Access (ThermoFisher Scientific) triple quadrupole mass spectrometer with a heated electrospray ion (HESI) source, unless otherwise indicated. Elution was carried out using a mobile phase of eluent B (MeCN with 0.1% Formic acid) and eluent A (water with 0.1% Formic acid) by

increasing the gradient from 25% to 47 % B from min. 0.0 to 4.0, increasing 47% to 100% B in min. 4.0 to 5.0, holding 100% B from min. 5.0 to 6.5 re-equilibration. The flow rate was 0.4 mL/min and the column temperature 35°C. The steviol glycosides were detected using SIM (Single Ion Monitoring) with the following m/z-traces.

**Table 1. MS analytical information for Steviol Glycosides**

Description	Exact Mass	m/z trace	compound (typical $t_R$ in min)
Steviol + 1 Glucose	[M+H] <sup>+</sup> 481.2796 [M+Na] <sup>+</sup> 503.2615	481.2± 0.5 503.1± 0.5	19-SMG (2.29), 13-SMG (3.5)
Steviol + 2 Glucose	[M+Na] <sup>+</sup> 665.3149	665± 0.5	Rubusoside (2.52) Steviol-1,2-bioside (2.92) Steviol-1,3-bioside (2.28)
Steviol + 3 Glucose	[M+Na] <sup>+</sup> 827.3677	827.4 ± 0.5	1,2-Stevioside (2.01) 1,3-Stevioside (2.39) RebB (2.88)
Steviol + 4 Glucose	[M+Na] <sup>+</sup> 989.4200	989.4 ± 0.5	RebA (2.0)
Steviol + 5 Glucose	[M+Na] <sup>+</sup> 1151.4728	1151.4 ± 0.5	RebD (1.1)
Steviol + 6 Glucose	[M+Na] <sup>+</sup> 1313.5257	1313.5 ± 0.5	RebM (1.3)

**[00135]** The levels of steviol glycosides were quantified by comparing with calibration curves obtained with authentic standards from LGC Standards. For example, standard solutions of 0.5 to 100 µM RebA were typically utilized to construct a calibration curve.

**[00136]** Method B: A second analytical method was performed on the Agilent system 1290 Infinity fitted with a waters ACQUITY UPLC® BEH shield RP18 column (2.1 x 50 mm, 1.7 µm particles, 130 Å pore size, Waters) was connected to a 6130 single quadrupol mass detector (Agilent) with a APCI ion source. Elution was carried out using a mobile phase of eluent B (MeCN with 0.1% Formic acid) and eluent A (water with 0.1% Formic acid) by increasing the gradient from 23% to 47 % B from min. 0.0 to 4.0, increasing 47% to 100% B in min. 4.0 to 5.0, holding 100% B from min. 5.0 to 6.5 re-equilibration. The flow rate was 0.6 mL/min and the column temperature 50°C. The steviol glycosides were detected using SIM (Single Ion Monitoring) with the following m/z-traces.

**Table 2. MS analytical information for Steviol Glycosides**

SIM trace No	time window	m/z trace	Exact Mass	Description	compound (typical $t_R$ in min)
1	0.0-1.51min	1289.5	[M-H] <sup>-</sup>	Steviol	RebM (0.91)

			1289.5281	+ 6 Glucose	
	1.51-1.90 min	687.3	[M+HCOOH-H]- 687.3217	Steviol + 2 Glucose	Rubusoside
	1.90-5.0 min	641.0	[M-H] <sup>-</sup> 641.3168	Steviol + 2 Glucose	1.2-Stevioside (1.44) 1.3-stevioside (1.74)
2	0.0-1.0 min	1127.4	[M-H] <sup>-</sup> 1127.4752	Steviol + 5 Glucose	RebD (0.81)
	1.0-5.0 min	525.3	[M-HCOOH-H]- 525.2689	Steviol + 1 Glucose	19SMG (2.49) 13SMG (2.65)
3	0.0-2.8 min	965.4	[M-H] <sup>-</sup> 965.4224	Steviol + 4 Glucose	RebA (1.42)
4	0.0-3.2 min	803.4	[M-H] <sup>-</sup> 803.3696	Steviol + 2 Glucose	1.2-Stevioside (2.16) 1.3-Stevioside (2.34) RebB (2.13)

**[00137]** The levels of steviol glycosides were quantified by comparing with calibration curves obtained with authentic standards from LGC Standards. For example, standard solutions of 0.3 to 25  $\mu$ M RebA were typically utilized to construct a calibration curve.

**[00138]** Method C: A third analytical method used was LC-MS analyses performed using a Waters ACQUITY UPLC (Waters Corporation, Milford, MA) with Waters ACQUITY UPLC® BEH C18 column (2.1 x 50 mm, 1.7  $\mu$ m particles, 130 Å pore size) coupled to a Waters ACQUITY TQD triple quadrupole mass spectrometer with electrospray ionization (ESI) in negative mode. Compound separation was achieved by a gradient of the two mobile phases A (water with 0.1% formic acid) and B (MeCN with 0.1% formic acid) by increasing from 20% to 50% B between 0.3 to 2.0 min, increasing to 100% B at 2.01 min, holding 100% B for 0.6 min and re-equilibrate for another 0.6 min. The flow rate was 0.6 mL/min and the column temperature 55°C. RebD (m/z 1127.5), RebM (m/z 1289.5), rebaudioside A (m/z 965.4) and RebB (m/z 803.4) were monitored using SIM (Single Ion Monitoring) and quantified by comparing with authentic standards.

### Example 2. Construction of a Steviol Glycoside-Producing Yeast Strain

**[00139]** Steviol glycoside-producing *S. cerevisiae* strains were constructed as described in WO 2011/153378, WO 2013/022989, WO 2014/122227, and WO 2014/122328, each of which is incorporated by reference in its entirety. For example, a yeast strain comprising a recombinant gene encoding a *Synechococcus* sp. GGPPS polypeptide (SEQ ID NO:1, SEQ ID NO:149), a recombinant gene encoding a truncated *Zea mays* CDPS polypeptide (SEQ ID NO:2, SEQ ID NO:150), a recombinant gene encoding an *A. thaliana* KS polypeptide (SEQ ID NO:3, SEQ ID NO:151), a recombinant gene encoding a recombinant *S. rebaudiana* K01 polypeptide (SEQ ID NO:4, SEQ ID NO:152), a recombinant gene encoding

an *A. thaliana* ATR2 polypeptide (SEQ ID NO:5, SEQ ID NO:153), a recombinant gene encoding an *O. sativa* EUGT1 1 polypeptide (SEQ ID NO:12; SEQ ID NO:148), a recombinant gene encoding an SrKAHel polypeptide (SEQ ID NO:6, SEQ ID NO:154), a recombinant gene encoding an *S. rebaudiana* CPR8 polypeptide (SEQ ID NO:7, SEQ ID NO:155), a recombinant gene encoding an *S. rebaudiana* UGT85C2 polypeptide (SEQ ID NO:8, SEQ ID NO:156), a recombinant gene encoding an *S. rebaudiana* UGT74G1 polypeptide (SEQ ID NO:9, SEQ ID NO:157), a recombinant gene encoding an *S. rebaudiana* UGT76G1 polypeptide (SEQ ID NO:10, SEQ ID NO:158), and a recombinant gene encoding an *S. rebaudiana* UGT91D2 variant (or functional homolog), UGT91D2e-b (SEQ ID NO:11, SEQ ID NO:159) polypeptide produced steviol glycosides. As analyzed by LC-MS (Method C) following DMSO-extraction of total steviol glycosides from the whole cell and broth mixture (total production), the strain produced between 18-21  $\mu\text{g/mL}$  or 1-1.5  $\mu\text{g/mL}/\text{OD}_{600}$  RebM after growth for five days in 1 mL SC (Synthetic Complete) media at 30°C with 400 rpm shaking in deep-well plates. See Table 3.

**Table 3. Steviol glycoside production in a representative *S. cerevisiae* strain comprising genes encoding GGPPS, truncated CDPS, KS, KO, ATR2, EUGT11, SrKAHel, CPR8, UGT85C2, UGT74G1, UGT76G1, and EUGT11 polypeptides.**

RebB ( $\mu\text{g/mL}/\text{OD}_{600}$ )	RebA ( $\mu\text{g/mL}/\text{OD}_{600}$ )	RebD ( $\mu\text{g/mL}/\text{OD}_{600}$ )	RebM ( $\mu\text{g/mL}/\text{OD}_{600}$ )	Normalized by $\text{OD}_{600}$
0.21	0.33	0.33	1.3	<b>Average</b>
0.028	0.054	0.032	0.14	<b>Std Deviation</b>
RebB ( $\mu\text{g/mL}$ )	RebA ( $\mu\text{g/mL}$ )	RebD ( $\mu\text{g/mL}$ )	RebM ( $\mu\text{g/mL}$ )	
3.1	4.9	5.0	19.0	<b>Average</b>
0.42	0.81	0.48	2.1	<b>Std Deviation</b>

**[00140]** A second strain, which comprised additional copies of the genes of the first strain, was analyzed for steviol glycoside production. The second strain produced RebD and RebM as primary steviol glycosides, although at higher levels than the first strain.

**[00141]** As analyzed by LC-MS (Method C) following DMSO-extraction of total steviol glycosides from the whole cell and broth mixture (total production), the second strain produced between 60-80  $\text{pg/mL}$  or 4-6  $\mu\text{g/mL}/\text{OD}_{600}$  RebM, after growth for five days in 1 mL SC media at 30°C with 400 rpm shaking in deep-well plates. Production of RebA, RebB, RebD and RebM by the second strain is shown in Table 4.



**Table 4. Steviol glycoside production in an *S. cerevisiae* strain comprising additional copies of genes encoding GGPPS, truncated CDPS, KS, KO, ATR2, EUGT11, SrKAHe-I, CPR8, UGT85C2, UGT74G1, UGT76G1, and EUGT11 polypeptides.**

RebA ( $\mu\text{g/mL}/\text{OD}_{600}$ )	RebB ( $\mu\text{g/mL}/\text{OD}_{600}$ )	RebD ( $\mu\text{g/mL}/\text{OD}_{600}$ )	RebM ( $\mu\text{g/mL}/\text{OD}_{600}$ )	Normalized by $\text{OD}_{600}$
2.1	0.67	1.6	4.8	<b>Average</b>
0.66	0.21	0.75	2.3	<b>Std Deviation</b>
RebA ( $\mu\text{g/mL}$ )	RebB ( $\mu\text{g/mL}$ )	RebD ( $\mu\text{g/mL}$ )	RebM ( $\mu\text{g/mL}$ )	
31.0	10.1	23.7	72.5	<b>Average</b>
9.9	3.1	11.3	34.4	<b>Std Deviation</b>

### **Example 3. Knockout of Yeast Endogenous Transport Genes and Transport-Related Genes**

**[00142]** Observations from deep-well studies of Example 2 and similar strains indicated that the fraction of RebA, RebB, RebD or RebM in the supernatant changes with time, and the effect was determined not to be the result of cell lysis. To determine the effect of various transporters on steviol glycoside excretion in *S. cerevisiae*, deletion cassettes for homologous recombination were obtained by designing primers annealing approximately 200 bp upstream and downstream of the open reading frame (ORF) and then amplifying the ORF-specific deletion cassette from the *S. cerevisiae* deletion collection. The candidate genes selected include identified ORFs with relation to transport or comprising membrane spanning domains, regardless of subcellular localization. In the resulting colonies, the presence of the deletion cassette at the correct locus was verified by colony **PGR**. A maximum of 6 clones of each deletion was frozen down as freezer stock. All samples for analysis were initiated from the freezer stock and grown in SC medium for 5 days (30°C, shaking 400 rpm) prior to harvest and extraction of samples for **LC-MS**. Samples were analyzed for the presence of RebA, RebB, RebD and RebM in the culture broth lacking cells (Supernatant) as well as in the whole cell and broth mixture (Total production).

**[00143]** Concentrations of total and supernatant RebA, RebB, RebD and RebM were compared to the levels in a control steviol glycoside-producing strain. The amounts of RebA, RebB, RebD and RebM in each sample were normalized to the control strain by dividing the value of a particular steviol glycoside with the corresponding value for the control strain, thereby calculating a percentage to the control strain, where 1 equals 100 percent. The "ideal candidate" would exhibit a decrease in RebA, RebB, RebD and/or RebM levels in the supernatant, as compared to the control steviol glycoside-producing strain, without decreasing RebA, RebB, RebD, and/or RebM total production.

**[00144]** The effect of yeast gene knockouts on transport of higher molecular weight steviol glycosides into the culture medium was tested in a strain that produces steviol glycosides, such as the strains described in Example 2. Disruption of each specific transporter gene was performed by homologous recombination. After 5 days of growth in 1 mL SC medium at 30°C and 400 rpm, cells were harvested. A 50  $\mu$ L aliquot of the culture was mixed with an equal volume of 100% DMSO, vortexed, and heated to 80°C for 10 min. The suspension was then centrifuged to remove cell debris. 60  $\mu$ L of the mixture were analyzed by LC-MS as the "Total" sample. The remaining culture was then centrifuged to pellet cells. An aliquot of 50  $\mu$ L was removed from the supernatant (*i.e.*, the culture medium) and mixed with an equal volume of 100% DMSO. The suspension was heated to 80°C for 10 min and centrifuged. 60  $\mu$ L of the mixture were analysed by LC-MS as the "Supernatant" sample. The amounts of higher molecular weight steviol glycosides (including RebA, RebB, RebD, RebM) were measured by LC-MS (Method C), as described in Example 1.

**[00145]** The data demonstrate that disruption of a single endogenous yeast transporter gene in a steviol glycoside-producing strain resulted in a decrease in the level of various steviol glycosides in the supernatant of the culture media, as evaluated by the normalized amount transported into the supernatant (see Tables 5-10). Tables 5-10 comprise lists of transport related genes that were knocked out in a steviol glycoside-producing strain. More specifically, Table 5 comprises a compiled list of genes by ordered locus name found to affect steviol glycoside excretion in steviol glycoside-producing strains and are therefore identified as having a role in steviol glycoside excretion. When the specified genes were knocked out, a more than 40% decrease in either the supernatant alone or in the ratio of supernatant/total production of RebA, RebB, RebD, and/or RebM was observed. This corresponded approximately to more than 2 standard deviations removed from the mean of a control steviol glycoside-producing strain (a value of 1 equals 100 percent of the control strain, whereas a value of 0.5 indicates a 50% decrease).

**[00146]** Table 6 comprises a compiled list of genes by ordered locus name found to affect steviol glycoside excretion in steviol glycoside-producing strains and are therefore identified as having a role in steviol glycoside excretion. When knocked out, these genes caused a mean of between 20-40% decrease in either the supernatant alone or in the ratio of supernatant/total production. This corresponded to approximately between 1 and 2 standard deviations removed from the mean of the control strain (a value of 1 equals 100 percent of the control strain, whereas a value of 0.5 indicates a 50% decrease).

**Table 5A. Transport related genes with over a 40% decrease in Reb A, RebB, RebD or RebM levels compared to a control steviol glycoside-producing strain.**

SEQ ID No.	Ordered Locus Name	Family	Description	Gene name	Uniprot Accession No.
13	YBR180W	MFS	Secondary Transporter	DTR1	P38125
14	YAL067C	MFS	Secondary Transporter	SEO1	P39709
15	YBL089W	AAAP	Secondary Transporter	AVT5	P38176
16	YBL099W	F-ATPase	ATP-Dependent	ATP1	P07251
17	YBR241C	MFS	Secondary Transporter		P38142
18	YBR294W	SulP	Secondary Transporter	SUL1	P38359
19	YCL069W	MFS	Secondary Transporter	VBA3	P25594
20	YCR028C	MFS	Secondary Transporter	FEN2	P25621
21	YCR075C	LCT	Secondary Transporter	ERS1	P17261
22	YDL128W	CaCA	Secondary Transporter	VCX1	Q99385
23	YDL185W	F-ATPase	ATP-Dependent	VMA1	P17255
24	YDL194W	MFS	Secondary Transporter	SNF3	P10870
25	YDL210W	APC	Secondary Transporter	UGA4	P32837
26	YDR061W	ABC	ATP-Dependent		Q12298
27	YDR093W	P-ATPase	ATP-Dependent	DNF2	Q12675
28	YDR338C	MOP/MATE	Secondary Transporter		Q05497
29	YDR406W	ABC	ATP-Dependent	PDR15	Q04182
30	YDR536W	MFS	Secondary Transporter	STL1	P39932
31	YEL031W	P-ATPase	ATP-Dependent	SPF1	P39986
32	YER166W	P-ATPase	ATP-Dependent	DNF1	P32660
33	YFL011W	MFS	Secondary Transporter	HXT10	P43581
34	YGL006W	P-ATPase	ATP-Dependent	PMC1	P38929
35	YGL013C		Transcription factor	PDR1	P12383
36	YGL255W	ZIP	Secondary Transporter	ZRT1	P32804
37	YGR125W	SulP	Secondary Transporter		P53273
38	YGR181W	MPT	ATP-Dependent	TIM13	P53299
39	YGR217W	VIC	Ion Channels	CCH1	P50077
40	YGR224W	MFS	Secondary Transporter	AZR1	P50080
41	YGR281W	ABC	ATP-Dependent	YOR1	P53049
42	YHL016C	SSS	Secondary Transporter	DUR3	P33413
43	YIL088C	AAAP	Secondary Transporter	AVT7	P40501
44	YJL093C	VIC	Ion Channels	TOK1	P40310
45	YJL094C	CPA2	Secondary Transporter	KHA1	P40309
46	YJL108C	ThrE	Secondary Transporter	PRM10	P42946
47	YJL212C	OPT	Secondary Transporter	OPT1	P40897
48	YJR106W	CaCA	Secondary Transporter	ECM27	P47144

**Table 5B. Continued list of Transport related genes with over a 40% decrease in Reb A, RebB, RebD or RebM levels compared to a control steviol glycoside-producing strain.**

No.	Ordered Locus Name	Family	Description	Gene name	Uniprot Accession No.
49	YJR160C	MFS	Secondary Transporter	MPH3	P0CE00
50	YKL064W	MIT	Ion Channels	MNR2	P35724
51	YKR050W	Trk	Secondary Transporter	TRK2	P28584
52	YKR105C	MFS	Secondary Transporter	VBA5	P36172
53	YKR106W	MFS	Secondary Transporter	GEX2	P36173
54	YLR447C	F-ATPase	ATP-Dependent	VMA6	P32366
55	YML116W	MFS	Secondary Transporter	SNQ1/ATR1	P13090
56	YMR034C	BASS	Secondary Transporter		Q05131
57	YMR056C	MC	Secondary Transporter	AAC1	P04710
58	YMR253C	DMT	Secondary Transporter		Q04835
59	YNL065W	MFS	Secondary Transporter	AQR1	P53943
60	YNL070W	MPT	ATP-Dependent	TOM7	P53507
61	YNL083W	MC	Secondary Transporter	SAL1	D6W196
62	YNL095C	AEC	Secondary Transporter		P53932
63	YNL121C	MPT	ATP-Dependent	TOM70	P07213
64	YNL142W	Amt	Ion Channels	MEP2	P41948
65	YOL020W	APC	Secondary Transporter	TAT2	P38967
66	YOL075C	ABC	ATP-Dependent		Q08234
67	YOL077W-A	F-ATPase	ATP-Dependent	ATP19	P81451
68	YOL122C	Nramp	Secondary Transporter	SMF1	P38925
69	YOR079C	ZIP	Secondary Transporter	ATX2	Q12067
70	YOR087W	TRP-CC	Ion Channels	YVC1	Q12324
71	YOR092W	AEC	Secondary Transporter	ECM3	Q99252
72	YOR130C	MC	Secondary Transporter	ORT1	Q12375
73	YOR222W	MC	Secondary Transporter	ODC2	Q99297
74	YOR291W	P-ATPase	ATP-Dependent	YPK9	Q12697
75	YOR306C	MFS	Secondary Transporter	MCH5	Q08777
76	YOR316C	CDF	Secondary Transporter	COT1	P32798
77	YOR334W	MIT	Ion Channels	MRS2	Q01926
78	YPL078C	F-ATPase	ATP-Dependent	ATP4	P05626
79	YPL270W	ABC	ATP-Dependent	MDL2	P33311
80	YPL274W	APC	Secondary Transporter	SAM3	Q08986
81	YPR003C	SulP	Secondary Transporter		P53394
82	YPR011C	MC	Secondary Transporter		Q12251
83	YPR058W	MC	Secondary Transporter	YMC1	P32331
84	YPR128C	MC	Secondary Transporter	ANT1	Q06497
85	YPR201W	ACR3	Secondary Transporter	ARR3	Q06598

**Table 6A. Transport related genes with a 20-40% decrease in Reb A, RebB, RebD or RebM levels compared to a control steviol glycoside-producing strain.**

SEQ ID No.	Ordered Locus Name	Family	Description	Gene name	Uniprot Accession No.
86	YBR008C	MFS	Secondary Transporter	FLR1	P38124
87	YBR021W	NCS1	Secondary Transporter	FUR4	P05316
88	YBR043C	MFS	Secondary Transporter	QDR3	P38227
89	YBR287W	AEC	Secondary Transporter		P38355
90	YBR295W	P-ATPase	ATP-Dependent	PCA1	P38360
91	YBR296C	PIT	Secondary Transporter	PHO89	P38361
92	YCL038C	MFS	Secondary Transporter	ATG22	P25568
93	YCR011C	ABC	ATP-Dependent	ADP1	P25371
94	YDL054C	MFS	Secondary Transporter	MCH1	Q07376
95	YDL100C	ArsAB	ATP-Dependent	GET3	Q12154
96	YDL245C	MFS	Secondary Transporter	HXT15	P54854
97	YDL247W	MFS	Secondary Transporter	MPH2	P0CD99
98	YDR011W	ABC	ATP-Dependent	SNQ2	P32568
99	YDR292C	IISP	ATP-Dependent	SRP101	P32916
100	YDR497C	MFS	Secondary Transporter	ITR1	P30605
101	YEL006W	MC	Secondary Transporter	YEA6	P39953
102	YEL027W	F-ATPase	ATP-Dependent	VMA3	P25515
103	YEL065W	MFS	Secondary Transporter	SIT1	P39980
104	YER019C-A	IISP	ATP-Dependent	SBH2	P52871
105	YER053C	MC	Secondary Transporter	PIC2	P40035
106	YER119C	AAAP	Secondary Transporter	AVT6	P40074
107	YFL028C	ABC	ATP-Dependent	CAF16	P43569
108	YFR045W	MC	Secondary Transporter		P43617
109	YGL084C	GUP	Secondary Transporter	GUP1	P53154
110	YGL104C	MFS	Secondary Transporter	VPS73	P53142
111	YGL114W	OPT	Secondary Transporter		P53134
112	YGL167C	P-ATPase	ATP-Dependent	PMR1	P13586
113	YGR257C	MC	Secondary Transporter	MTM1	P53320
114	YHL035C	ABC	ATP-Dependent	VMR1	P38735
115	YHL036W	APC	Secondary Transporter	MUP3	P38734

**Table 6B. Continued list of Transport related genes with a 20-40% decrease in Reb A, RebB, RebD or RebM levels compared to a control steviol glycoside-producing strain.**

No.	Ordered Locus Name	Family	Description	Gene name	Accession No.
116	YHR002W	MC	Secondary Transporter	LEU5	P38702
117	YHR096C	MFS	Secondary Transporter	HXT5	P38695
118	YIL006W	MC	Secondary Transporter	YIA6	P40556
119	YIL120W	MFS	Secondary Transporter	QDR1	P40475
120	YIL121W	MFS	Secondary Transporter	QDR2	P40474
121	YIL166C	MFS	Secondary Transporter	SOA1	P40445
122	YJL133W	MC	Secondary Transporter	MRS3	P10566
123	YJL219W	MFS	Secondary Transporter	HXT9	P40885
124	YKL016C	F-ATPase	ATP-Dependent	ATP7	P30902
125	YKL050C	MIT	Ion Channels		P35736
126	YKL120W	MC	Secondary Transporter	OAC1	P32332
127	YKL146W	AAAP	Secondary Transporter	AVT3	P36062
128	YKL209C	ABC	ATP-Dependent	STE6	P12866
129	YKR039W	APC	Secondary Transporter	GAP1	P19145
130	YLR411W	Ctr	Ion Channels	CTR3	Q06686
131	YML038C	DMT	Secondary Transporter	YMD8	Q03697
132	YMR166C	MC	Secondary Transporter		Q03829
133	YMR279C	MFS	Secondary Transporter		Q03263
134	YNL003C	MC	Secondary Transporter	PET8	P38921
135	YNL268W	APC	Secondary Transporter	LYP1	P32487
136	YNR055C	MFS	Secondary Transporter	HOL1	P53389
137	YOL158C	MFS	Secondary Transporter	ENB1	Q08299
138	YOR100C	MC	Secondary Transporter	CRC1	Q12289
139	YOR153W	ABC	ATP-Dependent	PDR5	P33302
140	YOR271C	MTC	Secondary Transporter	FSF1	Q12029
141	YOR273C	MFS	Secondary Transporter	TPO4	Q12256
142	YOR307C	DMT	Secondary Transporter	SLY41	P22215
143	YOR332W	F-ATPase	ATP-Dependent	VMA4	P22203
144	YOR348C	APC	Secondary Transporter	PUT4	P15380
145	YPL036W	P-ATPase	ATP-Dependent	PMA2	P19657

**[00147]** Steviol glycoside exporter candidates were selected from the data based on two selection criteria for each steviol glycoside measured (*i.e.*, two methods of normalizing expression).

**[00148]** Transporter selection criterion 1 corresponded to selection based on the level of high molecular weight steviol glycosides (RebA, RebB, RebD, or RebM) available in the supernatant, as well as the total production of the said steviol glycoside. Both values were normalized to the value of the corresponding steviol glycoside-producing control strain. The control level was set to 1, and the corresponding steviol glycoside level was calculated as a

percentage of the control. For Ordered Locus Names (*i.e.*, genes) of interest, the steviol glycoside available in the supernatant should be below 0.6 (below 60% of the control) or between 0.8-0.6 (80-60% of the control). To avoid false positives or a bias towards transporters that decrease the production in general, the calculation had an additional requirement that the total production had to be similar to the control. In the current calculation, production was set to be between 0.85 and 1.15 of the control, when the control is set to 1. In this regard, steviol glycoside production levels did not affect results. Table 7 shows the supernatant/total ratio for each candidate that fulfills the selection criteria.

[00149] Transporter selection criterion 2 corresponded to selection based on the ratio of high molecular weight steviol glycosides (RebA, RebB, RebD, or RebM) in the supernatant relative to total production of the said steviol glycoside. The supernatant-to-total production ratio was normalized to the ratio of the corresponding steviol glycoside-producing strain control. The control level was set to 1, and the corresponding steviol glycoside ratio was calculated as a percentage of the control. For Ordered Locus Names (*i.e.*, genes) of interest, the supernatant-to-total production ratio for a given steviol glycoside should be below 0.6 (below 60% of the control) or between 0.8-0.6 (80-60% of the control). To avoid false positives or a bias towards transporters that decrease the production in general, the calculation had an additional requirement that the total production had to be similar to the control. In the current calculation, production was set to be between 0.85 and 1.15 of the control, when the control is set to 1. In this regard, steviol glycoside production levels did not affect results. Table 8 shows the supernatant/total ratio for each candidate that fulfills the selection criteria.

[00150] The data demonstrate that disruption of a single endogenous yeast transporter gene in a steviol glycoside-producing strain resulted in a decrease in the level of various steviol glycosides in the supernatant of the culture media, as evaluated by the normalized amount transported into the supernatant (see Tables 5-10), and are therefore identified as having a role in steviol glycoside excretion.

[00151] For example, deletion in a steviol glycoside-producing strain of YDL128W (SEQ ID NO:22), YDL194W (SEQ ID NO:24), YDL210W (SEQ ID NO:25), YFL01 1W (SEQ ID NO:33), YGL006W (SEQ ID NO:34), YGL013C (SEQ ID NO:35), YGL255W (SEQ ID NO:36), YGR181W (SEQ ID NO:38), YGR217W (SEQ ID NO:39), YIL088C (SEQ ID NO:43), YJL094C (SEQ ID NO:45), YJR106W (SEQ ID NO:48), YNL065W (SEQ ID NO:59), YNL083W (SEQ ID NO:61), YNL121C (SEQ ID NO:63), YNL142W (SEQ ID NO:64), YOR306C (SEQ ID NO:75), or YPR01 1C (SEQ ID NO:82) led to a measurable decrease of RebD excreted into the culture medium, indicating that each plays a role in RebD excretion.

This was confirmed by transporter selection criteria 1 and 2 (see Tables 7 and 8, RebD column).

**[00152]** Furthermore, for example, deletion in a steviol glycoside-producing strain of YBR180W (SEQ ID NO:13), YBR241C (SEQ ID NO:17), YCL069W (SEQ ID NO:19), YCR075C (SEQ ID NO:21), YDL128W (SEQ ID NO:22), YDL194W (SEQ ID NO:24), YDR093W (SEQ ID NO:27), YDR338C (SEQ ID NO:28), YER166W (SEQ ID NO:32), YFL011W (SEQ ID NO:33), YGL006W (SEQ ID NO:34), YGL013C (SEQ ID NO:35), YGL255W (SEQ ID NO:36), YGR217W (SEQ ID NO:39), YJL094C (SEQ ID NO:45), YJR106W (SEQ ID NO:48), YJR160C (SEQ ID NO:49), YKR106W (SEQ ID NO:53), YML116W (SEQ ID NO:55), YMR056C (SEQ ID NO:57), YNL070W (SEQ ID NO:60), YNL083W (SEQ ID NO:61), YNL095C (SEQ ID NO:62), YNL121C (SEQ ID NO:63), YOR087W (SEQ ID NO:70), YOR291W (SEQ ID NO:74), YOR306C (SEQ ID NO:75), YPL274W (SEQ ID NO:80), or YPR011C (SEQ ID NO:82) led to a measurable decrease of RebM, indicating that each plays a role in RebM excretion. This was confirmed by transporter selection criteria 1 and 2 (see Tables 7 and 8, RebM column).

**[00153]** Table 7 represents the calculated ratio, normalized to a steviol glycoside-producing strain comprising genes encoding GGPPS, truncated CDPS, KS, KO, ATR2, EUGT1 1, SrKAHe1, CPR8, UGT85C2, UGT74G1, UGT76G1, and EUGT1 1 polypeptides, of supernatant/total production for each gene (by ordered locus name) deleted in the steviol glycoside-producing strain. The supernatant or supernatant/total ratio of less than 0.6 represented a more than 40% decrease in either the supernatant alone or in the ratio of supernatant/total production of RebA, RebB, RebD, or RebM, which corresponded approximately to more than 2 standard deviations removed from the mean of the control steviol glycoside-producing strain and indicates the gene as having a role in steviol glycoside transportation (Table 7). The supernatant or ratio supernatant/total of between 0.6 and 0.8 represents a 40-20% decrease in either the supernatant alone or in the ratio of supernatant/total production of RebA, RebB, RebD, or RebM, which corresponds to approximately between 1 and 2 standard deviations removed from the mean of the control steviol glycoside-producing strain, and indicates the gene as having a role in steviol glycoside transportation and/or production (Table 8). Total production of each steviol glycoside was between 0.85 and 1.15 compared to the steviol glycoside-producing strain. Table 8 shows the supernatant/total ratio for each candidate that fulfills the selection criteria.

**Table 7. Transport related genes with over a 40% decrease in RebA, RebB, RebD or RebM compared to a control steviol glycoside-producing strain comprising genes**



encoding **GGPPS**, truncated **CDPS**, **KS**, **KO**, **ATR2**, **EUGT11**, **SrKAHel**, **CPR8**, **UGT85C2**, **UGT74G1**, **UGT76G1**, and **EUGT11** polypeptides.

	Transporter selection criterion 1 Total vs. Supernatant				Transporter selection criterion 2 Ratio Sup/Total vs. Total			
	RebA	RebB	RebD	RebM	RebA	RebB	RebD	RebM
YBR180W				0.486				0.486
YBR241C				0.529				0.529
YCL069W				0.519				0.519
YCR075C				0.448				0.448
YDL128W			0.459	0.405			0.459	0.405
YDL194W			0.652	0.482				0.482
YDL210W			0.000				0.000	
YDR093W				0.569				0.569
YDR338C				0.451				0.451
YEL031W		0.488				0.488		
YER166W				0.495				0.495
YFL011W			0.581	0.547			0.581	0.547
YGL006W							0.410	0.424
YGL013C			0.673	0.507				0.507
YGL255W			0.669	0.632				
YGR181W			0.419				0.419	
YGR217W			0.598	0.429			0.598	0.429
YIL088C			0.135				0.135	
YJL094C			0.568	0.525			0.568	0.525
YJR106W			0.470	0.432			0.470	0.432
YJR160C				0.689				
YKL064W		0.337				0.337		
YKR106W				0.509				0.509
YML116W				0.706				
YMR056C								0.591
YNL065W							0.571	
YNL070W				0.633				
YNL083W				0.481			0.592	0.481
YNL095C				0.610				
YNL121C			0.620	0.456				0.456
YNL142W	0.561		0.369		0.561		0.369	
YOR087W				0.611				
YOR291W				0.681				
YOR306C			0.596	0.559			0.596	0.559
YOR334W		0.520				0.520		
YPL078C		0.590				0.590		
YPL270W		0.665						
YPL274W				0.561				0.561
YPR011C			0.542	0.611			0.542	

**Table 8.** Transport related genes with a 20-40% decrease in Reb A, RebB, RebD or RebM compared to a control steviol glycoside-producing strain comprising genes

encoding GGPPS, truncated CDPS, KS, KO, ATR2, EUGT11, SrKAHel, CPR8, UGT85C2, UGT74G1, UGT76G1, and EUGT11 polypeptides.

	Transports cal 1; total vs sup				Transports cal 2; ratio sup/total vs total			
	RebA	RebB	RebD	RebM	RebA	RebB	RebD	RebM
YBL089W				0.739				0.739
YBR008C	0.784			0.640	0.784			0.640
YBR021W		0.731				0.731		
YBR043C	0.755			0.796	0.755			0.796
YBR180W			0.747				0.747	
YBR241C			0.688		0.798		0.688	
YBR287W	0.781	0.823	0.768		0.781		0.768	
YBR295W			0.885	0.876				
YBR296C		0.724	0.799	0.790		0.724	0.799	0.790
YCL038C		0.709		0.752		0.709		0.752
YCL069W			0.785				0.785	
YCR075C			0.634				0.634	
YDL054C			0.920					
YDL100C			0.867					
YDL194W							0.652	
YDL210W				0.834				
YDL245C	0.852							
YDL247W				0.682				0.682
YDR011W			0.852					
YDR093W	0.792	0.775	0.704		0.792	0.775	0.704	
YDR338C	0.711	0.695	0.680		0.711	0.695	0.680	
YDR497C				0.694				0.694
YEL006W				0.657			0.774	0.657
YEL065W			0.635				0.635	
YER119C				0.872				
YER166W	0.771	0.843	0.687		0.771		0.687	
YFL011W		0.787				0.787		
YFL028C			0.641				0.641	
YFR045W			0.779				0.779	
YGL006W			0.410	0.424				
YGL013C							0.673	
YGL084C		0.804						
YGL104C	0.628	0.731		0.683	0.628	0.731		0.683
YGL114W						0.796		
YGL167C	0.829							
YGL255W							0.669	0.632
YGR217W		0.801						
YGR257C	0.842							
YHL035C			0.900	0.792				0.792

	Transports cal 1; total vs sup				Transports cal 2; ratio sup/total vs total			
	RebA	RebB	RebD	RebM	RebA	RebB	RebD	RebM
YHL036W				0.798				0.798
YHR096C			0.879	0.798				0.798
YIL006W	0.763			0.689	0.763		0.791	0.689
YIL120W				0.814				
YIL121W			0.903					
YIL166C			0.844					
YJL212C			0.817	0.682				0.682
YJR106W	0.719				0.719			
YJR160C		0.781	0.985			0.781		0.689
YKL050C				0.896				
YKL120W				0.706				0.706
YKL146W		0.890						
YKR039W	0.763				0.763			
YKR106W		0.785	0.738			0.785	0.738	
YLR41 1W	0.852		0.782				0.782	
YML038C			0.724				0.724	
YML116W			0.898					0.706
YMR056C			0.675	0.591		0.786	0.675	
YMR279C				0.885				
YNL065W	0.710	0.792	0.571		0.710	0.792		
YNL070W	0.893		0.892					0.633
YNL083W			0.592					
YNL095C			0.726				0.726	0.610
YNL121C							0.620	
YNL268W		0.920						
YNR055C			0.643				0.643	
YOL122C				0.935				
YOL158C			0.848	0.728				0.728
YOR087W								0.61 1
YOR100C		0.916						
YOR271C		0.889	0.758	0.608			0.758	0.608
YOR273C	0.726	0.916	0.635		0.726		0.635	
YOR291W								0.681
YOR307C								0.765
YOR348C				0.644				0.644
YPL036W	0.763		0.698		0.763		0.698	
YPL078C			0.798				0.798	
YPL270W			0.746			0.665	0.746	
YPL274W	0.817	0.807	0.721				0.721	
YPR01 1C	0.763				0.763			0.61 1

[00154] The effect of yeast gene knockouts on transport of higher molecular weight steviol glycosides into the culture medium (*i.e.*, supernatant) also was tested in a steviol glycoside-producing strain comprising additional copies of genes encoding GGPPS, truncated CDPS, KS, KO, ATR2, EUGT1 1, SrKAHef, CPR8, UGT85C2, UGT74G1, UGT76G1, and EUGT1 1 polypeptides, which was described in Example 2. The data demonstrated that disruption of a single endogenous yeast transporter gene in the steviol glycoside-producing strain resulted in a decrease in the level of various steviol glycosides in the supernatant of the culture media, as evaluated by the normalized amount transported or by the supernatant-to-total-production ratio (see Tables 9 and 10, RebD column). For example, deletion in the steviol glycoside-producing strain of YDR536W (SEQ ID NO:30), YHL016C (SEQ ID NO:42), YKR050W (SEQ ID NO:51), YOR291W (SEQ ID NO:74), YOR334W (SEQ ID NO:77), YPL270W (SEQ ID NO:79), YPR058W (SEQ ID NO:83), or YPR128C (SEQ ID NO:84) led to a measurable decrease of RebD transported into the supernatant, indicating that they play a role in RebD excretion. This was confirmed by transporter selection criteria 1 and 2 (see Tables 9 and 10, RebD column).

**[00155]** Furthermore, for example, deletion of YAL067C (SEQ ID NO:14), YDR406W (SEQ ID NO:29), YHL016C (SEQ ID NO:42), YJL212C (SEQ ID NO:47), YKR050W (SEQ ID NO:51), YMR034C (SEQ ID NO:56), YMR253C (SEQ ID NO:58), YOL075C (SEQ ID NO:66), YOL122C (SEQ ID NO:68), YOR222W (SEQ ID NO:73), YPR003C (SEQ ID NO:81), or YPR201W (SEQ ID NO:85) led to a measurable decrease of RebM transported into the supernatant, indicating that they play a role in RebM excretion. This was confirmed by transporter selection criteria 1 and 2 (see Tables 9 and 10, RebM column).

**[00156]** Table 9 represents the calculated ratio, normalized to a steviol glycoside-producing strain comprising additional copies of genes encoding GGPPS, truncated CDPS, KS, KO, ATR2, EUGT1 1, SrKAHef, CPR8, UGT85C2, UGT74G1, UGT76G1, and EUGT1 1 polypeptides, of supernatant/total production for each gene (by ordered locus name) deleted in the steviol glycoside-producing strain. The supernatant or ratio supernatant/total of less than 0.6 represents a more than 40% decrease in either the supernatant alone or in the ratio of supernatant/total production of RebA, RebB, RebD, or RebM, which corresponds approximately to more than 2 standard deviations removed from the mean of a control steviol glycoside-producing strain, and indicates the gene as having a role in steviol glycoside transportation and/or production (Table 9). The supernatant or ratio supernatant/total of between 0.6 and 0.8 represents a 40-20% decrease in either the supernatant alone or in the ratio of supernatant/total production of RebA, RebB, RebD, or RebM, which corresponds to approximately between 1 and 2 standard deviations removed from the mean of the control strain, and indicates the gene as having a role in steviol

glycoside transportation and/or production, and indicates the gene as having a role in steviol glycoside transportation and/or production (Table 10). Total production of each steviol glycoside was between 0.85 and 1.15 compared to the control steviol glycoside-producing strain. Table 10 shows the supernatant/total ratio for each candidate that fulfills the selection criteria.

**Table 9. Transport related genes with over a 40% decrease in Reb A, RebB, RebD or RebM compared to a control steviol glycoside-producing strain comprising additional copies of genes encoding GGPPS, truncated CDPS, KS, KO, ATR2, EUGT11, SrKAHel, CPR8, UGT85C2, UGT74G1, UGT76G1, and EUGT11 polypeptides.**

	Transporter selection criterion 1 total vs sup				Transporter selection criterion 2 ratio sup/total vs total			
	RebA	RebB	RebD	RebM	RebA	RebB	RebD	RebM
YAL067C				0.541				0.541
YBL089W	0.433	0.416			0.433	0.416		
YBL099W	0.523				0.523			
YBR294W	0.495				0.495			
YCR028C		0.419				0.419		
YDL185W	0.551				0.551			
YDL210W	0.626	0.469				0.469		
YDR061W	0.482		0.471		0.482		0.471	
YDR406W				0.288				0.288
YDR536W	0.715		0.365				0.365	
YFL011W		0.444				0.444		
YGR125W		0.400				0.400		
YGR224W		0.361				0.361		
YGR281W		0.596				0.596		
YHL016C			0.427	0.296			0.427	0.296
YJL093C		0.449				0.449		
YJL108C	0.589				0.589			
YJL212C	0.442			0.461	0.442			0.461
YKR050W	0.554		0.378	0.304	0.554		0.378	0.304
YLR447C	0.512				0.512			
YMR034C	0.331			0.316	0.331			0.316
YMR253C	0.389			0.375	0.389			0.375
YOL020W	0.371				0.371			
YOL075C	0.494			0.471	0.494			0.471
YOL077W-A	0.531				0.531			
YOL122C				0.457				0.457
YOR079C	0.552				0.552			
YOR092W	0.407				0.407			
YOR130C	0.588				0.588			
YOR222W	0.469			0.457	0.469			0.457
YOR291W			0.428				0.428	
YOR334W			0.327				0.327	

	Transporter selection criterion 1 total vs sup				Transporter selection criterion 2 ratio sup/total vs total			
	RebA	RebB	RebD	RebM	RebA	RebB	RebD	RebM
YPL270W			0.375				0.375	
YPR003C	0.400			0.418	0.400			0.418
YPR058W			0.461				0.461	
YPR128C			0.342				0.342	
YPR201W	0.376			0.353	0.376			0.353

**Table 10. Transport related genes with a 20-40% decrease in Reb A, RebB, RebD or RebM compared to a control steviol glycoside-producing strain comprising additional copies of genes encoding GGPPS, truncated CDPS, KS, KO, ATR2, EUGT11, SrKAHel, CPR8, UGT85C2, UGT74G1, UGT76G1, and EUGT11 polypeptides.**

	Transports cal 1; total vs sup				Transports cal 2; ratio sup/total vs total			
	RebA	RebB	RebD	RebM	RebA	RebB	RebD	RebM
YCR011C				0.654				0.654
YDL210W			0.729		0.626		0.729	
YDR292C	0.724				0.724			
YDR536W					0.715			
YEL027W		0.799				0.799		
YER019C-A	0.789				0.789			
YER053C	0.651				0.651			
YGR256W	0.744				0.744			
YHR002W	0.795				0.795			
YJL133W	0.691				0.691			
YJL219W	0.674				0.674			
YKL016C	0.627				0.627			
YKL209C	0.721				0.721			
YKR105C					0.646			
YMR166C		0.924						
YNL003C		0.814						
YOR153W	0.801							
YOR316C					0.640			
YOR332W	0.700				0.700			

**[00157]** Knockouts of YDL210W (SEQ ID NO:25) and YPL270W (SEQ ID NO:79) resulted in decreased RebD excretion in the steviol glycoside-producing strain comprising genes encoding GGPPS, truncated CDPS, KS, KO, **ATR2**, EUGT11, SrKAHel, CPR8, UGT85C2, UGT74G1, UGT76G1, and EUGT11 polypeptides and the steviol glycoside-producing strain comprising additional copies of genes encoding GGPPS, truncated CDPS, KS, KO, ATR2, EUGT11, SrKAHel, CPR8, UGT85C2, UGT74G1, UGT76G1, and EUGT11 polypeptides. As well, knockouts of YJL212C (SEQ ID NO:47) and YOL122C (SEQ ID NO:68) resulted in decreased RebM transport in both strains.

**Example 4. Confirmation of Knockout of Yeast Endogenous Transport Genes by overexpression in a RebD/M-producing strain**

**[00158]** Overexpression of a subset of the initial candidate transporters from Example 3 was performed using both plasmid-based expression and an integration cassette. First, deep-well microtiter plate culture experiments were carried out. Two transport genes were overexpressed using a plasmid in a RebD/M-producing strain in order to confirm the results from the knockout experiments. YGR181W (SEQ ID NO:38), a TIM complex, helper protein for insertion of mitochondrial inner membrane proteins, and YDR061W (SEQ ID NO:26) an ABC-like transporter were overexpressed. The data shown in Figure 2 demonstrate that the phenotype based on the knockout studies was confirmed with a plasmid based overexpression phenotype for YGR181W (SEQ ID NO:38) and YDR061W (SEQ ID NO:26) in deep-well plates.

**[00159]** Next, confirmation of the phenotype in fermenters was performed in additional steviol glycoside-producing strains, which were characterized by intergration of YGR181W (SEQ ID NO:38) or YDR061W (SEQ ID NO:26) on chromosome XII. The steviol glycoside-producing strains were grown on defined media at 30°C in a fed-batch fermentation for about 5 days under glucose-limited conditions, and the levels of RebA, RebB, RebD, and RebM were measured using LC-MS (Method B, Example 1). The graphs shown in Figure 3 illustrate an approximate 2-fold increase in RebD and RebM transported in the culture medium for the new integration constructs, and little change in RebA and RebB transport. Overexpression of YGR181W (SEQ ID NO:38) or YDR061W (SEQ ID NO:26) resulted in improved (~2-fold) RebD and RebM transport into the culture medium (~400-500 mg/L of supernatant RebD and RebM in YGR181W (SEQ ID NO:38) and YDR061W (SEQ ID NO:26) overexpression strains versus ~250 mg/L of supernatant RebD and RebM in a control steviol glycoside-producing strain). See Figure 3A. The ratio of transported RebD as compared to the total RebD increased from 0.158 in the control strain to 0.21-0.25 with the candidate genes overexpressed. RebM transport into the culture medium was also simultaneously improved. See Figure 3.

#### **Example 5. Overexpression of Selected Yeast Endogenous Transport Genes**

**[00160]** Overexpression in a steviol glycoside-producing strain (as described in Example 2) using a plasmid with a constitutive promoter of the transporter genes shown in Table 11 resulted in greater than a 20% increase in excretion of RebA, RebB, RebD, and/or RebM. Results were analyzed using criterion 2 described in Example 3. Additionally, overexpression of the transporter genes shown in Table 12 resulted in greater than a 40% improvement in production of RebA, RebB, RebD, and/or RebM. Table 11 shows the supernatant/total ratio for each candidate that fulfills the selection criteria.



Table 11. Transport related genes with over a 20% increase in RebA, RebB, RebD or RebM excretion, compared to a control steviol glycoside-producing strain.

	Ratio Supernatant/Total			
	RebB	RebA	RebD	RebM
YOR079C			1.21	
YMR166C		1.36	1.53	1.38
YEL027W		1.62	1.82	1.52
YDL054C		1.45	1.38	1.31
YKL120W		1.83	1.89	1.93
YDR536W		1.79	1.80	1.76
YBL099W				1.22
YML116W		1.32	1.31	1.42
YIL166C			1.27	1.22
YKR039W			1.26	1.41
YOR307C				1.23
YKL146W		1.36	1.47	1.66
YGL167C				1.33
YJL093C				1.29
YOR306C	1.67			
YDL128W	1.85		1.29	
YOR153W	1.42		1.21	
YKL050C	1.59	1.22		
YJL094C	1.71	1.24	1.24	
YCL069W	1.59			
YOL158C	1.52			
YFL011W	1.44			

	Ratio Supernatant/Total			
	RebB	RebA	RebD	RebM
YJR106W			1.38	1.33
YBR043C				1.20
YPR011C				1.27

**Table 12. Transport related genes with over a 40% increase in RebA, RebB, RebD or RebM production, compared to a control stevioi glycoside-producing strain.**

	Increases in Production			
	RebB	RebA	RebD	RebM
YMR166C				1.52
YIL166C		1.41	1.50	1.55
YKR039W			1.48	1.52
YKL146W				1.42
YJL093C			1.46	1.43
YOR306C				1.59
YDL128W				1.49
YOL122C			1.41	1.59
YIL006W			1.64	2.03
YFL028C				1.55
YBR021W			1.51	1.87
YHR002W			1.51	1.73
YEL031W			1.45	1.66
YCL069W				1.53
YOL158C			1.42	1.63
YKL064W			1.40	1.44
YHR096C				1.42

	Increases in Production			
	RebB	RebA	RebD	RebM
YOR332W				1.44
YDR338C			1.50	1.55
YJR106W			1.41	1.44
YBR043C			1.55	1.49
YPR011C				1.43
YFR045W			1.44	

#### Example 6. Genomic **Integration of** Transporter Genes

**[00161]** DNA of the transporter genes selected for integration into the genome of a RebD/M-producing *S. cerevisiae* strain (see Example 2) was amplified from an S288C background by PGR and cloned into a plasmid with homology regions for the integration site and a PGK1 promoter for overexpression, using the USER cloning system. See, e.g., Nour-Eidin *et al.*, 2010, *Methods Mol Biol.* 643:185-200. The USER cloning construct including the homology regions and the transporter was cut out from the plasmid using restriction enzymes, and the linear piece of DNA was integrated into the genome of the receiving RebD/M-producing strain by standard LiAc method. The genomically integrated transporters were tested in plates that release glucose from a polymer after addition of a growth medium. A polymer that releases 20 g/L glucose over 3 days was used to mimic the feed profile during fermentation. Steviol glycoside levels were measured by LC-MS (see Example 1), and OD<sub>600</sub> was measured on a Perkin Elmer 2104 Multilabel reader. YBR043C (SEQ ID NO:88), YEL027W (SEQ ID NO:102), YJL093C (SEQ ID NO:44), YJR106W (SEQ ID NO:48), YKL120W (SEQ ID NO:126), and YMR166C (SEQ ID NO:132) showed improved excretion of 13-SMG. (Figure 4A). YBR043C (SEQ ID NO:88), YEL027W (SEQ ID NO:102), and YMR166C (SEQ ID NO:132) showed improved excretion of RebA (Figure 4B). YBR043C (SEQ ID NO:88), YEL027W (SEQ ID NO:102), and YMR166C (SEQ ID NO:132) showed improved excretion of RebB (Figure 4C). YBR043C of SEQ ID NO:88, YEL027W of SEQ ID NO:102, YJL093C of SEQ ID NO:44, YJR106W of SEQ ID NO:48, and YMR166C of SEQ ID NO:132 showed improved production of RebD, and YBR043C of SEQ ID NO:88, YEL027W of SEQ ID NO:102, YJL093C (SEQ ID NO:121), YJL093C of SEQ ID NO:44, YJR106W of SEQ ID NO:48, and YMR166C of SEQ ID NO:132 showed improved production of RebM, as measured by an increase in RebD and RebM levels in the

supernatant compared to a control steviol glycoside-producing strain. See Figures 4D and 4E. Controls with a URA marker are also shown in Figure 4.

**[00162]** Figure 5A shows supernatant levels of RebA, RebB, RebD, and RebM of an additional steviol glycoside-producing strain overexpressing YMR166C (SEQ ID NO:132), YEL027W (SEQ ID NO:102), YKL120W (SEQ ID NO:126), YJR106W (SEQ ID NO:48), YJL093C (SEQ ID NO:44), and YBR043C (SEQ ID NO:88) by the USER cloning system. The strain of Figure 5 comprised a recombinant gene encoding a *Synechococcus sp.* GGPPS polypeptide (SEQ ID NO:1, SEQ ID NO:149), a recombinant gene encoding a truncated *Zea mays* CDPS polypeptide (SEQ ID NO:2, SEQ ID NO:150), a recombinant gene encoding an *A. thaliana* KS polypeptide (SEQ ID NO:3, SEQ ID NO:151), a recombinant gene encoding a recombinant *S. rebaudiana* K01 polypeptide (SEQ ID NO:4, SEQ ID NO:152), a recombinant gene encoding a KO polypeptide (SEQ ID NO:XX, SEQ ID NO:XX), a recombinant gene encoding an *A. thaliana* ATR2 polypeptide (SEQ ID NO:5, SEQ ID NO:153), a recombinant gene encoding an *O. sativa* EUGT1 1 polypeptide (SEQ ID NO:12; SEQ ID NO:148), a recombinant gene encoding an SrKAHel polypeptide (SEQ ID NO:6, SEQ ID NO:154), a recombinant gene encoding an *S. rebaudiana* CPR8 polypeptide (SEQ ID NO:7, SEQ ID NO:155), a recombinant gene encoding an *S. rebaudiana* UGT85C2 polypeptide (SEQ ID NO:8, SEQ ID NO:156), a recombinant gene encoding an *S. rebaudiana* UGT74G1 polypeptide (SEQ ID NO:9, SEQ ID NO:157), a recombinant gene encoding an *S. rebaudiana* UGT76G1 polypeptide (SEQ ID NO:10, SEQ ID NO:158), and a recombinant gene encoding an *S. rebaudiana* UGT91D2 variant (or functional homolog), UGT91D2e-b (SEQ ID NO:11, SEQ ID NO:159) polypeptide. Figure 5B shows total levels of RebA, RebB, RebD, and RebM of the above described steviol glycoside-producing strain overexpressing YMR166C (SEQ ID NO:132), YEL027W (SEQ ID NO:102), YKL120W (SEQ ID NO:126), YIL166C (SEQ ID NO:132), YJR106W (SEQ ID NO:48), YJL093C (SEQ ID NO:44), and YBR043C (SEQ ID NO:88) by the USER cloning system.

#### **Example 7. Production of RebD and RebM by Fermentation of Steviol Glycoside-Producing *S. cerevisiae* strains overexpressing YJL093C or YBR043C**

**[00163]** YJL093C (SEQ ID NO:44) and YBR043C (SEQ ID NO:88) were individually overexpressed in the steviol glycoside-producing strain described in Example 3. The strains were cultivated by fermentation (fed-batch, minimum medium, glucose-limiting) for approximately 130 h. Production of RebD and RebM was measured by LC-MS. As shown in Table 13, the strains overexpressing YJL093C or YBR043C produced higher levels of RebD and RebD + RebM, as compared to a control steviol glycoside-producing strain.

**Table 13. Production of RebD and RebM in *S. cerevisiae* strains overexpressing YJL093C and YBR043C.**

<b>Strain</b>	<b>Ferm. Length (h)</b>	<b>Final Cell Dry Weight</b>	<b>RebD Titer (g/L)</b>	<b>RebM Titer (g/L)</b>	<b>RebD + RebM</b>	<b>RebD/RebM Ratio (g/g)</b>
Control	126.83	104.53	1.38	4.47	5.85	0.31
YJL093C	130.10	114.40	3.42	2.80	6.22	1.22
YBR043C	129.17	112.00	3.56	2.72	6.28	1.31

Table 14. Sequences disclosed herein.

SEQ ID NO: 1

Synechococcus sp. GGPPS (GenBank ABC98596.1)

atggtcgcac	aaactttcaa	cctggatacc	<b>tacttatccc</b>	aaagacaaca	acaagttgaa	60
gaggccctaa	<b>gtgctgctct</b>	tgtgccagct	tatcctgaga	gaatatacga	<b>agctatgaga</b>	120
tactccctcc	<b>tgccaggtgg</b>	caaaagatta	<b>agacctatct</b>	tatgttttagc	<b>tgcttgcgaa</b>	180
ttggcaggtg	gttctgttga	<b>acaagccatg</b>	ccaactgcgt	gtgcacttga	aatgatccat	240
acaatgtcac	taattcatga	tgacctgcca	gccatggata	acgatgattt	<b>cagaagagga</b>	300
aagccaacta	<b>atcacaaggt</b>	gttcggggaa	<b>gatatagcca</b>	tcttagcggg	<b>tgatgcgctt</b>	360
ttagcttacg	cttttgaaca	tattgcttct	caaacaagag	gagtaccacc	tcaattggtg	420
ctacaagtta	ttgctagaat	cggacacgcc	gttgctgcaa	caggcctcgt	tggaggccaa	480
gtcgtagacc	ttgaatctga	aggtaaagct	atctccttag	aaacattgga	gtatattcac	540
tcacataaga	ctggagcctt	gctggaagca	tcagttgtct	caggcgggat	tctcgcaggg	600
gcagatgaag	agcttttggc	cagattgtct	cattacgcta	gagatatagg	<b>cttggctttt</b>	660
caaatcgtcg	atgatatcct	ggatgttact	gctacatctg	aacagttggg	gaaaaccgct	720
ggtaaagacc	aggcagccgc	aaaggcaact	tatccaagtc	<b>tattgggttt</b>	agaagcctct	780
agacagaaaag	<b>cggaagagtt</b>	gattcaatct	<b>gctaaggaag</b>	<b>ccttaagacc</b>	ttacggttca	840
caagcagagc	cactcctagc	gctggcagac	ttcatcacac	gtcgtcagca	ttaa	894

SEQ ID NO: 2

Zea mays truncated CDPS

atggcacagcaca	catcagaatc	cgagctgtc	gcaaaggca	gcagtttgac	ccctatagtg	60
agaactgacg	ctgagtcaag	gagaacaaga	tggccaaccg	atgacgatga	cgccgaacct	120
ttagtggtgag	agatcagggc	aatgcttact	<b>tccatgtctg</b>	atggtgacat	<b>ttccgtgagc</b>	180
gcacacgata	cagcctgggt	cggattgggt	<b>ccaagattag</b>	acggcgggtga	<b>aggtcctcaa</b>	240
tttccagcag	ctgtgagatg	gataagaaat	aaccagttgc	ctgacggaag	ttggggcgat	300
gccgcattat	tctctgcta	tgacaggctt	atcaataccc	ttgcctgcgt	<b>tgtaactttg</b>	360
acaagtggt	ccctagaacc	agagatgaga	ggtagaggac	tatctttttt	<b>gggtaggaac</b>	420
atgtggaat	tagcaactga	agatgaagag	tcaatgccta	ttggcttcga	attagcattt	480
ccatctttga	tagagcttgc	taagagccta	ggtgtccatg	acttccctta	tgatcaccag	540
gcctacaag	gaatctactc	ttcaagagag	atcaaaatga	agaggattcc	aaaagaagtg	600
atgcataccg	ttccaacatc	aatattgcac	agtttgagg	gtatgcctgg	cctagattgg	660
gctaaactac	ttaaactaca	gagcagcgac	ggaagttttt	tgttctcacc	agctgccact	720
gcatatgctt	<b>taatgaatac</b>	cggagatgac	aggtgtttta	gctacatcga.	tagaacagta	780
aagaaattca	acggcggcgt	ccctaattgtt	tatccagtg	atctatttga	acataatttg	840
gccgttgata	gacttgaaag	attaggaatc	tccaggtact	tccaaaagga	gatcgaacaa	900
tgcatggatt	atgtaaacag	gcattggact	gaggacggta	tttgttgggc	<b>aaggaactct</b>	960
gatgtcaaaag	aggtggacga	cacagctatg	gcctttagac	ttcttaggtt	gcacggctac	1020
agcgtcagtc	ctgatgtgtt	taaaaacttc	gaaaaggacg	gtgaattttt	cgcatctgtc	1080
ggacagtcta	atcaagctgt	taccggtatg	tacaacttaa	acagagcaag	ccagatatcc	1140
ttcccaggcg	aggatgtgct	tcatagagct	ggtgccttct	catatgagtt	cttgaggaga	1200
aaagaagcag	agggagcttt	gagggacaag	tggatcattt	ctaaagatct	acctggtgaa	1260
gttgtgtata	ctttggattt	tccatggtac	ggcaacttac	ctagagtcca	ggccagagac	1320
tacctagagc	aatacggagg	tggtgatgac	gtttgattg	gcaagacatt	<b>gtataggatg</b>	1380
ccacttgtaa	acaatgatgt	atatttgga	ttggcaagaa	tggatttcaa	ccactgccag	1440
gctttgcatc	<b>agttagagtg</b>	gcaaggacta	<b>aaaagatggt</b>	atactgaaaa	taggttgatg	1500
gactttgggtg	tcgcccaaga	agatgccctt	agagcttatt	ttcttgacgc	cgcatctggt	1560
tacgagcctt	gtagagctgc	cgagaggctt	gcatgggcta	gagccgcaat	actagctaac	1620
gccgtgagca	cccacttaag	aaatagccca	tcattcagag	aaaggttaga	gcattctctt	1680
aggtgtagac	ctagtgaaga	gacagatggc	tcttggttta	actcctcaag	tggctctgat	1740
gcagtttttag	taaaggctgt	cttaagactt	actgattcat	tagccagggg	agcacagcca	1800
atccatggag	gtgaccaga	agatattata	cacaagttgt	taagatctgc	ttgggcccag	1860

<b>tgggttaggg</b>	aaaaggcaga	cgctgccgat	agcgtgtgca	atggtagttc	tgcagtagaa	1920
caagagggat	caagaatggt	ccatgataaa	cagacctgtc	tattattggc	tagaatgatc	1980
gaaatttctg	ccggtagggc	agctggtgaa	gcagccagtg	aggacggcga	tagaagaata	2040
attcaattaa	caggctccat	ctgcgacagt	cttaagcaaa	aatgctagt	ttcacaggac	2100
cctgaaaaaa	atgaagagat	gatgtctcac	gtggatgacg	aattgaagtt	<b>gaggattaga</b>	2160
gagttcgttc	aatatttgct	tagactaggt	gaaaaaaaga	ctggatctag	cgaaaccagg	2220
caaacatttt	taagtatagt	gaaatcatgt	tactatgctg	ctcattgccc	acctcatgtc	2280
gttgatagac	acattagtag	agtgattttc	gagccagtaa	gtgccgcaaa	gtaaccgcgg	2340

SEQ ID NO: 3

Arabidopsis thaliana KS (similar to GenBank AEE36246. 1)

atgtctatta	atgtgagatc	ttccggttgt	agctcccaa	taagcgcaac	tttggaaagg	60
ggtctagact	ctgaagttca	aacaagagca	aacaatgtat	cttttgagca	gaccaaagag	120
aagatcagga	aatgcttga	gaaggtcgag	ttgagcgtga	<b>gtgcctatga</b>	cactagttgg	180
gtagctatgg	tcccatcacc	atccagtcaa	<b>aacgcacctc</b>	ttttcccaca	gtgcgtcaaa	240
tggctacttg	ataatcaaca	tgaggacggc	tcttggggat	tggataacca	cgaccatcag	300
agcttaaaga	aagatgtggt	gtcatccaca	<b>ttagcctcta</b>	tcctagctct	taagaaatgg	360
ggaataggcg	aaagacagat	caataagggt	ctacagttca	ttgaattaaa	ctctgcacta	420
gttaccgatg	aaactataca	aaaacctaca	ggtttcgaca	tcatttttcc	aggaatgatt	480
aagtacgcca	gggaccttaa	tttgaccata	cctcttggtc	cagaagtagt	cgacgatatg	540
atcaggaaaa	gagatctaga	cttaaagtgt	gatagcgaga	aattcagcaa	aggtagagag	600
gcttatcttg	cctatgttct	tgaaggaact	aggaacttga	aggactggga	cttaattgtg	660
aaatatcaga	gaaagaacgg	tagtctattt	gatagtccag	ctacaaccgc	cgcagctttc	720
actcaatttg	gcaatgacgg	ttgcttgagg	tacttatggt	cacttttaca	gaaattcgag	780
gccgcagtgc	ctagtgtata	tccatttgat	caatacgcta	gattaagcat	aatcgtcact	840
ttagaatcat	<b>tgggaattga</b>	cagagatttc	aagactgaga	taaaaagcat	attggatgag	900
acctataggt	actggcttag	aggtgacgaa	gaaatttgcc	tagatttggc	cacatgtgca	960
<b>cttgctttta</b>	ggttgctttt	agcccacggc	<b>tatgacgtgt</b>	catacgatcc	tctaaagcca	1020
tttgacagagg	aatctggttt	cagcgatacc	cttgagggat	atgttaaaaa	caccttttcc	1080
gtattagagc	ttttcaaggc	tgcccaaagt	taccctcatg	agagtgcctt	gaaaaagcag	1140
tgttgctgga	caaaacaata	tctagaaatg	gaactaagtt	catgggttaa	aacaagcgtt	1200
agggacaagt	acttgaaaaa	ggaagtggag	gatgctttgg	catttccatc	atatgcctct	1260
ttagaaagaa	gtgaccacag	aaggaaaatt	cttaatggct	cagcagttga	aaacacaaga	1320
gtaaccaaga	cctcttacag	gttgcataat	atatgtacat	cagatatcct	<b>aaaacttgct</b>	1380
gtcgacgatt	tcaacttttg	ccaatctatt	catagagagg	aaatggaaaag	attggataga	1440
tggatagtgg	agaatagact	acaggaatta	aagttcgcca	gacaaaaatt	ggcttactgt	1500
tacttttagtg	gcgctgccac	actattctct	ccagaattgt	ctgacgcaag	gatctcatgg	1560
gctaagggag	gtgttctaac	cacagtagtc	gatgactttt	<b>ttgatgttgg</b>	cggtagtaaa	1620
gaagagcttg	agaacttaat	tcacttgggtg	gaaaagtggg	<b>atcttaatgg</b>	agttcctgaa	1680
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ccagctacat	accttatagg	acctccacta	cctgaaaaga	ctgtcgactc	<b>ccaccaatat</b>	1980
aatcaattat	acaaattggg	tagtaccatg	ggtagactat	taaacgatat	ccagggcttt	2040
aagagggaa	cagccgaggg	aaaacttaat	gcagtgtctc	tacatatgaa	gcatgaaaga	2100
gacaacagaa	gcaaagaggt	tattatagaa	tccatgaaag	gattggctga	aaggaaaaga	2160
gaggaattac	acaaacttgt	actagaagag	aaaggtagtg	tcgttccaag	agaatgcaag	2220
gaagccttct	taaaaatgtc	aaaagtgttg	aacctttttt	ataggaagga	tgatggcttc	2280
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aaggagagtc	taacttga					2358

SEQ ID NO; 4

s. rebaudiana KOI (codon optimized)

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gctgtagcat	tgccggtagc	gctaactctt	tggtacctga	aatcctacac	atcagctaga	120
agatcccaat	caaatcatct	tccaagagtg	cctgaagtcc	caggtgttcc	attgttagga	180
aatctgttac	aattgaagga	gaaaaagcca	tacatgactt	ttacgagatg	ggcagcgaca	240
tatggaccta	tctatagtat	caaaactggg	gctacaagta	tggttgtggt	atcatcta	300
gagatagcca	aggaggcatt	ggtgaccaga	ttccaatcca	tatctacaag	gaacttatct	360
aaagccctga	aagtacttac	agcagataag	acaatggtcg	caatgtcaga	ttatgatgat	420
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aagcatagaa	ttcacagaga	tatcatgatg	gataacatat	ctactcaact	tcatgaattc	540
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ttcggcttag	ctatgagaca	agccttagga	aaggatggtg	aaagtttgta	cgttgaagac	660
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<b>ctaggcggct</b>	accatgttcc	tgctggcaca	gaacttgccg	ttaacatcta	cggttgcaac	1260
atggacaaaa	acgtttggga	aatccagag	gaatggaacc	cagaaagatt	catgaaagag	1320
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ggttccttgc	aagccctttt	aactgcactc	attgggattg	ggagaatggt	tcaagagttc	1440
gaatggaaac	tgaaggatat	gactcaagag	gaagtgaaca	cgataggcct	aactacacaa	1500
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SEQ ID NO: 5

A. thaliana ATR2 (codon optimized)

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gaattgtctt	caatgttgat	cgaaaacaga	caattcgcca	tgatcgtaac	tacatcaatc	180
<b>gctgttttga</b>	tcggttgat	tgctcatggt	gtatggagaa	gatccggtag	tggttaattct	240
aaaagagtcg	aacctttgaa	accattagta	attaagccaa	gagaagaaga	aatagatgac	300
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<b>ttggatgact</b>	atgccgctga	tgacgatgaa	tacgaagaaa	agttgaagaa	agaagatggt	480
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tacaaatggt	ttacagaggg	taatgatcgt	ggtgaatggt	tgaaaaactt	aaagtacggt	600
gttttcgggt	tggttaacag	acaatacgaa	catttcaaca	aagttgcaaa	ggttgcgac	660
gatattttgg	tcgaacaagg	tgctcaaaga	ttagtccaag	taggtttggg	tgacgatgac	720
caatgtatag	aagatgactt	tactgcctgg	agagaagctt	tgtggcctga	attagacaca	780
atcttgagag	aagaaggtga	caccgccggt	gctaccccat	atactgctgc	agtattagaa	840
tacagagttt	ccatccatga	tagtgaagac	gcaaagttta	atgatatcac	tttggccaat	900
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agagaattac	atacaccaga	atccgacaga	agttgtatac	acttgggaatt	tgatatcgct	1020
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gaaactgttg	atgaagcatt	gagattggtg	gatatgtccc	ctgacactta	ttttagtttg	1140
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ttagcatctc	cagccggtaa	agatgaatat	<b>tcaaagtggtg</b>	tagttgaatc	tcaaagatca	1380



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gaaactagaa	ttcatgttac	atgtgcatta	gtctacgaaa	agatgccaac	cggtagaatt	1560
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acaattgctc	aagaacaagg	ttccatggat	agtaccaaag	ctgaaggttt	cgtaaagaac	2100
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SEQ ID NO: 6

**Stevia** rebaudiana KAHel (codon-optimized)

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aaaaagagag	atgacttttt	ccagggtttg	attgaacagg	ttagaaaatc	tcgtgggtgct	780
aaagtaggca	aaggtagaaa	aacgatgatc	gaactcctat	tatctttgca	agagtcagaa	840
cctgagtact	atacagatgc	tatgataaga	tcttttgccc	taggtctgct	ggctgcaggt	900
agtgatactt	cagcgggcac	tatggaatgg	gcatgatgct	tactgggtcaa	tcaccacat	960
gtattgaaga	aagctcaagc	tgaaatcgat	agagttatcg	gtaataacag	attgattgac	1020
gagtcagaca	ttggaaatat	cccttacatc	gggtgtatta	tcaatgaaac	tctaagactc	1080
tatccagcag	ggcattgtt	gttcccacat	gaaagtctg	ccgactgctg	tatttccggt	1140
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taa						1503

SEQ ID NO: 7

**Stevia** rebaudiana CPR8

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ATCGTGAGCTGTTGATGATACTCACAACGTCGGTTGCTGTATTGATCGGATGCGTTGTCGTTTTGGTG  
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 CAACTTGGATGAAGAACGCAGTGCCTATGACCGAGAGTCAAGATTGCAGTTGGGCCCAATATACGTC  
 CGAACATCCAATTTTCAGACTACCATCTGACCCTAAGGTCCCGGTTATCATGATTGGACCTGGCACTGG  
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 CCGTGACGTTTGGTAA

SEQ ID NO: 8

*Stevia rebaudiana* UGT85C2 (codon optimized)

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ataactttcg	tgaataccga	cttcatccat	aatcaatttc	tggaaatctag	tggccctcat	180
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gatcgtttca	ttgacttggg	cacaaaactt	ccagaccac	caacttgcac	aatctctgat	360
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cacaagatga	gaaacaaggc	caaagattgg	aaggaaaaag	ccagaattgc	tattgctcct	138 0
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aactaa						1446

SEQ ID NO: 9

S, rebaudiana UGT74G1 {GenBank AAR06920.1)

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<b>gcaggagaa</b>	catatttgg	aacattcaaa	caagttgggt	ctaaatcact	agctgactta	300
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<b>aatctttcgg</b>	<b>aagtaataaa</b>	aaccggaaag	ggtttgattg	tagcatggtg	caaacaattg	1020
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SEQ ID NO: 10

S, rebaudiana UGT7 6G1 (codon optimized)

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caacaggaag	tttttagctca	tggcgcctatt	<b>ggggcattct</b>	ggactcattc	cggatggaat	1080

tcaacttttag	aatcagtatg	cgaaggggta	cctatgatct	tttcagattt	tggctctgat	1140
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ggctgggaaa	ggggtgaaat	agctaatagca	ataagacgtg	ttatggttga	tgaagagggg	1260
gagtatatca	gacaaaacgc	aagagtgcctg	aagcaaaagg	ccgacgtttc	tctaatagaag	1320
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SEQ ID NO: 11

s. rebaudiana **UGT91D2e-b** (codon optimized)

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SEQ ID NO: 12

Oryza sativa sequence encoding EUGT11 (codon optimized)

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agatcctgcg	tcgagttcga	acctgaaaca	gtacctttac	tatctacttt	gagaggcaaa	720
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ctaccagctg	ggttcgaaga	gagaacaaga	ggccgtgggtg	tcgttgctac	tagatgggtc	1020
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aatgatggtg atggttcctt tgatagagaa ggcggtgcag ctgccatcag agcagtcgcc      1260
gttgaggaag agtcatctaa agttttccaa gctaaggcca aaaaattaca agagattgtg      1320
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aaagactaa                                     1389
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SEQ ID NO: 13

YBR180W

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>sp|P38125|DTR1_YEAST Dityrosine transporter 1 OS=Saccharomyces
cerevisiae (strain ATCC 204508 / S288c) GN=DTR1 PE=1 SV=1
MGSEPFQKKNLGLQINSQESGTTTRSTFHSLEDLGDDVINESWDQVNQKRANIDHDFHEH
PDSSPSLSAQKAKTKEEEVAVKSSNSQSRDPSPTQAHIPYTYFSKDQRLI IFGII IFIG
FLGPMGSGNIYIPALPLLQREYDVSATTINATVSVFMAVFSVGPLFWGALADFGGRKFLYM
VLSLMLLIVNILLAAVPVNIAAL FVLRIFQAFASSSVISLGAGTVTDVVPKHRGKAIAY
FMMGPNMGPI IAPIVAGLILMKGNVWRWLFGFTSMTGIALILVTALLPETLRCIVGNGD
PKWGDKKDERENNESPFEGNKISHRRLFPDIGIRKPVNNDAAFFQENFPKPPKAGLTLYW
KMIKCPPI IITSVSTALLFSSYYAFSVTFSYYLEHDYRFTMLEIGAAYVCPGVAMLLGSQ
SGGHLSDYLRSRWIKSHPKKKFPAEFRLLLNIGILLTICGTIGYGWAIFFFHYHFVLLV
FSALTAFGMTWCSNTSMTYLTELFPKRAAGTVAVSSFFRNVGAAISSAI ILQLCNAMGIG
WCFTGLGLCSSISLIGILYLLI FORKYTAKE F
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SEQ ID NO: 14

YAL067C

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>sp|P39709|SE01_YEAST Probable transporter SE01 OS=Saccharomyces
cerevisiae (strain ATCC 204508 / S288c) GN=SE01 PE=1 SV=1
MYSIVKEIIVDPYKRLKWFIPVKRQVEDLPDDLNSTEIVTISNSIQSHETAENFITTTT
EKDQLHFETSSYSEHKDNVNVTRSIEYRDEADRPWRRFFDEQEYRINEKERSHKNWYSWF
KQGTSFKEKLLIKLDVLLAFYSCIAYWVKYLDVTNINNAVYVSGMKEDLGFQGNLHVHTQ
VMYTVGNI IFQLPFLI YLNKLPLNYVLPDLCLWVSLTLVGAAYVNSVPHLKAIRFFIGAF
EAPSYLAYQYLFGSFYKHDEMVRSAFYLLGQYIGILSAGGIQSAVYSSLNGVNGLEGWR
WNFI IDAIVSVVVGILGFYSLPGDPYNCYSIFLTDDEIRLARKRLKENQTKGSDFFETKVF
DIKLWKTIFSDWKIYILTWNIFCWNSNVSSGAYLLWLKSLKRYISIPKLNQLSMITPGL
GMVYLMGTGI IADKLHSRWF AI IFTQVFNI IGNS ILAAWDVAEGAKWFAFMLQCFGWAMA
PVLVSWQNDICRRDAQTRAITLVTMNIMAQSSTAWISVLVWKTEEAPRYLKGFTFTACSA
FCLSIWTFVVLYFYKRDERNNAKNGIVLYNSKHGVEKPTSKDVETLSVSDEK
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SEQ ID NO: 15

YBL089W

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>sp|P38176|AVT5_YEAST Vacuolar amino acid transporter 5
OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=AVT5
PE=3 SV=2
MPSNVRSGVLTLLHTACGAGVLAMPFAFKPFGLMPGLITLTFCGICSLCGLLLQTRIAKY
VPKSENASFAKLTQLINPSISVVFDAIAVKCFGVGSYLIIVGDLVPQI VQSIFYRND
NMSGSQEHHMFLDRRLYITLI IVFVISPLCFKRSLNSLRYASMIIVSVAYLSGLIYHF
VNRHQLEERGQYVFMVPHGDSQSHSPLTTLPI FVFAYTCHHNMFSVINEQVDKSFKVIIRI
PIFAIVLAYFLYI IIGGTGYMTFGENIVGNILTLYPNSISTTIGRLAMLLVMLAFPLQC
HPCRSSVKNII IFIENFRKGKLYDNRASFIPLDNFNSEDPQEAPTQONNEEPNLRSESLR
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HINIITLCILLFsYLLAISITSLAKVLAI VGATGSTS ISFTLPGLFGYKLIGSEFTGTNE  
RVPTSIKIFKYLSSLFIWGIAMVASLSAIVFLGTSSH

SEQ ID NO: 16

YBL099W

>sp|P07251|ATPA\_YEAST ATP synthase subunit alpha, mitochondrial  
OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=ATP1  
PE=1 SV=5

MLARTAAIRSLSRTLINSTKAARPAAAALASTRRLASTKAQPTEVSSILEERIKGVSDEA  
NLNETGRVLAVGDGIARVFLNNIQAEELVE FSSGVKGMALNLE PGQVGIVLFGS DRLVK  
EGELVKRTGNIVDVPVGPGLLGRVVDALGNPIDGKGPIDAAGRSRAQVKAPGILPRRSVH  
EPVQTGLKAVDALVP IGRGQRELI IGDRQTGKTAVALDT ILNQRWNNGS DESKKLYCVY  
VAVGQKRSTVAQLVQTLQHDAMKYSIIVAATASEAAPLQYLAPFTAASIGEWFRDNGKH  
ALIVYDDLKQAVAYRQLSLLRRPPGREAYPGDVFYLSRLLERAAKLSEKEGSGSLTA  
LPVIETQGGDVSAIYPTNVISITDGQIFLEAELFYKIRPAINVGLSVSRVGSAAQVKAL  
KQVAGSLKLFQAQYREVAFAQFGSDLDASTKQTLVLRGERLTQLLKQNYSPATEEQVP  
LIYAGVNGHLDGIELSRIGEFESSFLSYLKSNNHLLTEIREKGELSKELLASLKSATES  
FVATF

SEQ ID NO: 17

YBR241C

>sp|P38142|YB91\_YEAST Probable metabolite transport protein YBR241C  
OS=Saccharomyces cerevisiae (strain ATCC 204508 / 5288c) GN=YBR241C  
PE=1 SV=1

MAETERLMPNGGSRETKPLITGHLLIGTIVACLGSIQYGYHIAELNAPQEFLLSCSRFEAP  
DENISYDDTWVQGHGLKQCIALTDSQYGAITSIFSIGGLFGSYYAGNWANRYGRKYVSMG  
ASAMCMVSSLLLLFFSNSYLQLLFGFLVGMSCGTAIVITPLFINEIAPVEWRGAMGSMNQ  
VSINLGILLTQTLALKYADSYNWRWLLFSGSVIAVANILAWLKVDESPRWLVSHGFVSEA  
ETALFKLRPGTYQQAQKEIQDWQRSHGHNRDPESSEETHSGPTLWQYVTDPSYKKPRTVI  
LAILSCQFCGINSI IFYGVKVIKILPDYSIQVNFASILNVVVTLAASAI IDHVGRRP  
LLLASTTVMTAMSLLSVGLTLSVSVFLVLTATFVYIAAFAIGLPIPLI IGELSYPQDA  
ATAQSFGTVCNWLATFIVGYLFPIGHGLMGGYVFAIFAAIAAMFATYVYKRVPETKGKTT  
YSEVWAGY

SEQ ID NO: 18

YBR294W

>sp|P38359|SUL1\_YEAST Sulfate permease 1 OS=Saccharomyces cerevisiae  
(strain ATCC 204508 / S288c) GN=SUL1 PE=1 SV=2

MSRKSSTEYVHNQEDADIEVFSEYRTYRESEAAENRDGLHNGDEENWKNSSKQKFGVT  
KNELSDVLYDS IPAYEESTVTLKEYYDHS IKNNLTAKSAGS YLVSLFPI IKWFPHYNFTW  
GYADLVAGITVGCVLVPQSMSYAQIASLSPEYGLYSSFIGAFIYSLFATSKDVCIGPVAV  
MSLQTAKVIAEVLKKYPEDQTEVTAPIIATTLCLLCGIVATGLGILRLGFLVELISLNAV  
AGFMTGSAFNI IWGQIPALMGYNLSLVNTREATYKVVINTLKHLNPKLDAVFGLI PLVIL  
YVWKWCGTFGITLADRYRNPQKVANRLKSFYFYAQAMRNAVVIVVFTAISWSITRNKS  
SKDRPISILGTVPSGLNEVGMKIPDGLLSNMSSEIPASI IVLVLEHIAISKSFGRINDY  
KVVPDQELIAIGVTNLIGTFFHSYPATGSFERSALKKCNVRTPFSGVFTGGCVLLALYC  
LTDAFFFPKATLSAVIIHAVSDLLTSYKTTWTFWKTNPLDCISFIVTVFITVFSSIENG  
IYFAMCWSCAMLLKQAFPAGKFLGRVEVAEVLNPTVQEDIDAVISSNELPNELNKQVKS  
TVEVLPAPYKFS VKWVPFDHGYRELNINTTVRPPPGVIVYRLGDSFTYVNCSTRHYDI  
IFDRIKEETRRGQLITLRKSDRPWNDPGEWKMPDSLKSLFKFKRHSATTNSDLPISNGS  
SNGETYEKPLLVCLDFSQVAQVDSTAVQSLVDLRKAVNRYADRQVEFHFAGIIISPWIK

RSL L5VKFGTTNEEYSDDSI IAGH5SFHVAKYLKDDYDYIDED5RISTSYSNYETLCAAT  
 GTNLPFFHIDI PDFSKWDV

SEQ ID NO: 19

YCL069W

>sp|P25594|YBA3\_YEAST Vacuolar basic amino acid transporter 3  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=YBA3  
 PE=1 SV=1

MNMLIVGRVVASVGGSGLQTLCFVIGCTMVGERSRPLVISILSCAFVAAIVGPI IGGAF  
 TTHVTWRWCFYINLPIGGLAI IMFLLTYKAENKGI LQQIKDAIGTISSFTFSKFRHQVNF  
 KRLMNGII FKFDFFGFALCSAGLVFLFLLGLTFGGNKYSWNSGQVIAYLVLVGLLFI FSLV  
 YDFFLFDKFNPEPDNISYRPLLLRRLVAKPAI I IINMVTFLCTGYNGQMIYSVQFFQLI  
 FASSAWKAGLHLIPIVITNVIAAIASGVITKKLGLVKPLLIFGGVLGVIGAGLMTLMTNT  
 STKSTQIGVLLLLPGFSLGFALQASLMSAQLQITKDRPEAAMDFIEVTA FNTFMKS LGTTL  
 GGVLSTTVFSASFHNKVSRAHLEPYEGKTVDMMILYRLQNYDGSHTIGNILSDSIKNVF  
 WMDLGFYALGFLFCSFSSNKLI IPKKDETPEDNLEDK

SEQ ID NO: 20

YCR028C

>sp|P25621|FEN2\_YEAST Pantothenate transporter FEN2 OS=Saccharomyces  
 cerevisiae (strain ATCC 204508 / 5288c) GN=FEN2 PE=1 SV=1

MMKESKSITQHEVERESVSSKRAIKRLLLFKIDLFVLSFVCLQYWINYVDRVGFNTAYI  
 SGMKEDLKMVGNDLTVSNTVFMIGYIVGMVPPNMLLVCVPPRIWLSFCTFAWGLLTLGMY  
 KVTSFKHICAIRFFQALFESCTFSGTHFVLGSWYKEDELPIRSAIFTGSLVGSFMFSGFM  
 QTSIFTHLNGRNLGAWRWLF IIDFCITLPIAI YGFIFFPGLPDQTS AVSKFSMTRYIFN  
 EQELHYARRRLPARDESTRLDWSTIPRVLKRWHWMMFSLVWVLGGENLGFASNSTFALWL  
 QNQKYTLAQRNNYPSGIFAVGIVSTLCSAVYMSKI PRARHWHVSVFISLVMVIVAVLIRA  
 DPLNPKVVFSAQYLGAVAYAGQAVFFSWANI ICHADLQERAIVLASMNMFSGAVNAWWSI  
 LFFASDMVPKFERGCYALLATAISSGIVSVIRSLQIKENLSKKQVPYIDANDMPGEDDD  
 DDNQDNENDGDDESMEVELHNEEMAEISNPF

SEQ ID NO: 21

YCR075C

>sp|P17261|ERS1\_YEAST Cystine transporter OS=Saccharomyces  
 cerevisiae (strain ATCC 204508 / S288c) GN=ERS1 PE=1 SV=1

MVSLDDILGIVYVTSWSISMYPPIITNWRHKSASAIMDFVMLNTAGYSYLVISIFLQLY  
 CWKMTGDES DLGRP KLTQDFWYCLHGCLMNVL LTTQVVAGARIWRFPKGHRKMNPWYL  
 RILLASLAIFSLLTQFMYSNYWDWHNSRTLAYCNNLFL LKISMSLIKYPQVTHNSTR  
 KSMDCFP IQGVFLDVTGGIASLLQLIWQLSNDQGFSLDTFVTNFGKVGLSMVTLIFNFIF  
 IMQWFVYRSRGHDLASEYPL

SEQ ID NO: 22

YDL128W

>sp|Q99385|VCX1\_YEAST Vacuolar calcium ion transporter  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=VCX1  
 PE=1 SV=1

MDATPPLLTVANSHPARNP KHTAWRAAVYDLQYILKASPLN FLLV FVPLGLIWHGFQLSH  
 TLTF LFNFLAI IPLAAILANATEELADKAGNTIGLLNATFGNAVELIVSI IALKKGQVR  
 IVQASMLGSLLSNLLLVLGLCFI FGGYNRVQQTFNQTAQTMSLLAIACASLLI PAAFR

ATLPHGKEDHFIDGKILELSRGTSSIVILIVYFLFLYFQLGSHHALFEQQEEETDEY MSTI
SRNPHHSLSVKSSLVILLGTTVIIISFCADFLGTIDNVESTGLSKTFIGLIYIPIYGNA
AEHYSVLVAMKDKMDLALGYAIGSSLQYAL FVT PFMVLVGWM I DVPMTLNFSTFETATL
FIAVFLSNYLILDGESNWLEGVMSLAMYILIAMAFFYYPDEKTLDSIGNSL

SEQ ID NO:23

YDL185W

>sp| P17255|YAT A\_\_YEAST V-type proton ATPase catalytic subunit A
OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=YMA1
PE=1 S=3

MAGAIENARKEIKRISLEDHAESEYGAIIYSYSGPYIAENMIGCAMYELKVGHDNLVGE
VIRIDGDKATIQVYEETAGLTVGDPVLRGTGKPLSVELGPGMETI YDGIQRPLKAIKEES
QSIYIPRGIDTPALDRTIKWQFTPGKFQYGDHISGGDIYGSYFENSLISSHKILLPPRSR
GTITWIAPAGEYTLDEKILEVEFDGKKSDFTLHTWPVRVPRPVTEKLSADYPLLTGQRV
LDALFPCYQGGTTCIPGAFGCGKTVISQSLSKYSNSDAII YVGCFAKGTNVLMDGSIEC
IENIEYGNKVMGKDGPREYIKLPRGRETMYSYQKSQHRAHKS DSSREVPPELLKFTCNA
THELYYRTPRSYRRLSRTIKGVEYFEVITFEMGQKKAPDGRIVELYKEVSKSYPISEGPE
RANELVESYRKASNKAYFEWTEIARDLSLLGSHVRKATYQTYAPILYENDHFFDYMQSK
FHLTIEGPKVLAYLLGLWIGDGLSDRATFSVDSRDTSLMERYTEYAEKLNLC AEYKDRKE
PQYAKTYNLYSKYRNGIRNNLNTENPLWDAIYGLGFLKDGKNI PSFLSTDNIGTRET
FLAGLIDSDGYVTDEHGKATIKTIHTSVRDGLVSLARSLGLYVSVNAEPAKVDMMNGTKH
KISYAIYMSGGDVLLNLVLSK CAGSKKFRPAPAAAFARECRGFYFELQELKEDDYGITLS
DDSDHQFLLANQYVHNCGERGNEMAEVLMFPELYTEMSGTKEPIMKRTTLYANTSNMP
YAAREASIYTGITLAEYFRDQGNYSMIADSSSRWAEALREISGR LGEMPADQGF PAYLG
AKLAS FYERAGKAVALGS PDRTGSVS IYAAYS PAGGDFS DPYTTATLGI TQVFWGLDKKL
AQRKHFP SINTSYSYKYTNLKNKFYDSNYPEFPVLRDRMKEILSNAEELEQYVQLVGKS
ALSDDSKITLDVATLIKEDFLQONGYSTYDAFCPIWKTFDMMRAFISYHDEAQAANGA
NWSKLADSTGDVKHAYSSSKFFEPSRGEKEVHGFEKLLSTMQERFAESTD

SEQ ID NO: 24

YDL194W

>sp| P10870|SNF3\_\_YEAST High-affinity glucose transporter SNF3
OS=Saccharomyces cerevisiae (strain ATCC 204508 / 5288c) GN=SNF3
PE=1 SV=3

MDPNSNSSSETLRQEKGFLDKALQRVKGIALRRNNSNKDHTTDDTTGSI RPTPSLQRQN
SDRQSNM TSVFTDDISTIDDNSILFSEPPQKQSMMSICYGYFVA YGGFLFGYDTGLINS
ITSMNYVKSHVAPNHDSFTAQQMSILVSFLSLGTFFGALTAPFISDSYGRKPTI IFSTIF
IFSIGNSLQV GAGGITLLIVGRYISGIGIGAISAVVPLYQAEATHKSLRGAI ISTYQWA I
TWGLLVSSAVSQGTHARN DASSYRIPIGLQYWSSFLAIGMFFLPESPRYYLKD KLDEA
AKSLSFLRGVPVHDSGLLEELVEIKATYDYEASFGSSNFIDCFISSKSRPKQTLRMFTGI
ALQAFQQFSGINFIFYGVNFFNKTGVSNSYLVSFITYAVNVVFNVPGLFFVEFFGRRKV
LVVGGVIMTIANFIVAIYGC SLKTYAAAKYMI AFICLFIAAFSATWGGVYWFYISAELYPL
GVRSKCTAICAAANWLVNFCALITPYIYDTGSHTSSLGAKI FFIWGS LNAMGYI VVYLT
YYETKGLTLEEIDELYIKSSTGVYSPKFNKDIRERALKFYDPLQRLEDGKNTFVAKRNN
FDETPRNDFRNTISGEIDHSPNQKEYHSIPERYDIPTSTEILESPNKSSGMTYPPYSPSL
QDVPIPQTTEPAEIRTKYDLGNGLGLNTYNRGPPSLSSDSEDEYDEIIGPSSQGDQS
NRSTMNDINDYMARLIHSTSTASNTTDKFSGNQSTLRYHTASSHSDTTEEDSNLMDLNG
LALNAYNRGPPSILMNSSDEEANGGETSDNLNTAQDLAGMKERMAQFAQSYIDKRGGLEP
ETQSNILSTSL SVMADTNEHNNEILHSSEENATNQPVNENNDLK



SEQ ID NO: 25

YDL210W

>sp|P32837|UGA4\_YEAST GABA-spccific permease OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=UGA4 PE=1 SV=1  
 MSMSKKNENKISVEQRISTDIGQAYQLQGLGSNLSRSKGTGAGEVNYIDAAKSVNDNQL  
 LAEIGYKQELKRQFSTLQVFGIAFSIMGLLPSIASVMGGGLGGGPATLVWGWVFAAFIL  
 LVGITMAEHASSIPTAGGLYYWTTYAPEGYKEI ISFI IGCSNSLALAAGVCSIDYGLAE  
 EIAAAVTLTKDGNFEVTSGLYGFAGAVVVMCICTCVASGAIARLQTLISIFANLFIIVL  
 LFIALPIGTKHRMGGFNDGDFIFGKYENLSDWNNGWQFCLAGFMPAVWTIGSFDSCVHQS  
 EEAKDAKKSVPIGI ISSIAVCWILGWLI IICLMACINPDIDSVLDSKYGFALAQI IYDSL  
 GKKWAIAFMSLIAFCQFLMGASITTAVSRQVWAFSRDNGPLPSKYIKRVDSKYSPFFAI  
 LAACVGLLILGLLCLIDDAATDALFSLAVAGNNLAWSTPTVFRLTSGRDLFRPGPFYLGK  
 IWSPIVAWTGVAFQLFII ILVMFPSQQHGITKSTMNYACVIGPGIWIWLAGI YYKVYKKKY  
 YHGPATNLSDDDYTEAVGADVDTIMSKQEP

SEQ ID NO: 26

YDR061W

>sp|Q12298|YD061\_YEAST Uncharacterized ABC transporter ATP-binding protein YDR061W OS=Saccharomyces cerevisiae (strain ATCC 204508 / 5288c) GN=YDR061W PE=1 SV=1  
 MSTNKFVVRITNALFKSSLASNSPPVYPKRIRHFEILPNEKWWIWGPGKGFLLDVLNNKY  
 ICEPPLSLRFGLKESNILPRIEQVAFKGVMPHTAHLARYEYFKDDYDQTKQFIFDKA  
 SGSNAVSYKVETNNRQINMELYNALVENLNLSSLQDRWVMGLSNGQMRRARLARSILKEP  
 DLLLIDDPFLGLDPAAIATISQFLAKYDSIEVSGGCPVIGLRYQDTIPAWCTHICCVDE  
 KNGILFEGPIEKLQSKMDETRSRALKELEQLKKASNSKEDISINDLICIHMPYGGKEHEI  
 IKMPHLIELDGLSVSYKGEAVLENLHWKVQPGSKWHIRGDNGSGKSTLLSLLTAEHPQSW  
 NSRVIDNGVPRRTGKTNYFDLNSKIGMSSPELHAI FLKNAGGRLNIRESVATGYHEASSN  
 NYLPWIWKRDKNSQEIWNMYLKYFGLDKDADSVLFEQLSVSDQKLVLFVRSLIKMPQIL  
 LDEAFSGMEVEPMMRCHEFLEEWPGTVLVVAHVAEETPKCAHYLRLLISPGEYEIGDMEN

SEQ ID NO: 27

YDR093W

>sp|Q12675|ATC4\_YEAST Phospholipid-transporting ATPase DNF2 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=DNF2 PE=1 SV=1  
 MSSPSKPTSPFVDDIEHESGSASNGLSSMSPFDDSFQFEKPSAHGNIEVAKTGGSVLKR  
 QSKPMKDI STPDLKSVTFDGIIDDYSNDNDINDDDELNGKTEIHEHENEVDDDLHSFQAT  
 PMPNTGGFEDVELDNNEGSNNSQADHKLKRVRFGRTRRNKSGRIDINRSKTLKWAKKNFH  
 NAIDEFSTKEDLENSALQNRSDLRVYVYNNLPLPEDMLDEDGLPLAVYPRNKIRTTKYT  
 PLTFFPKNILFQFHNFIYFLILLILGAFQIFGVTNPGFASVPLIVIVI ITAIKDGIED  
 SRRTVLDLEVNNTTRTHILSGVKNENVAVDNVSLWRRFKKANTRALIKIFEYFSENLTAA  
 REKKLQKKREELRRKRNSRSFGPRGSLDSIGSYRMSADFRPSLDYENLNQTM SQANRYN  
 DGENLVDRTLQPNPECRFAKDYWKNVKVGDIVRVHNNDEIPADMILLSTSDVDGACYVET  
 KNLDGETNLKVRQSLKCSKI IKSSRDITRTRKFWVESEGHANLYSYQGNFKWQDTQNGNI  
 RNEPVNINLLLRGCTLRNTKWAMGMVIFTGDDTKIMINAGVTPTKKSRI SRELNFSVIL  
 NFVLLFILCFTAGIVNGVYKQKPRSRDYFEFGTIGGSASTNGFVSFWAVILYQSLVPI  
 SLYISVEI IKTAQAI FITYDVLVLYNAKLDYPCPKSWNISDDLQIEYIFSDKTGTTLTQN  
 VMEFKCTINGVSYGRAYTEALAGLRKRQGV DVESEGREKEEIAKDRETMIDELRSMDS  
 NTQFCPEDLTFVSKEIVEDLKGSSGDHQKCEHFLALALCHSVLVEPNKDDPKKLDIK  
 AQSPDESALVSTARQLGYSFVGS SKSGLIVEIQGVQKEFQVLNVLEFNSSRKRMSCI IKI  
 PGSTPKDEPKALLICKGADSVIYRDLRDTQNDATLLEKTALHLEEYATEGLRTLCLAQRE  
 LTWSEYERWVKTYDVAASVTNREEELDKVTDVIERELILLGGTAIEDRLQDGVPSIAL

LAEAGIKLWVLTGDKVETAINIGFSCNVLNNDMELLVVKASGEDVEEFGSDPIQVNNLV
TKYLREKFGMSGSEELKEAKREHGLPQGNFAVI IDGDALKVALNGEEMRRKFLLLCKNC
KAVLCCRVSPAQKAADVVKLVKKTLDVMTLAIGDGSNDVAMIQSADVGVGIAGEEGRQAVM
CSDYAIGQFRYVTRLVLVHGKWCYKRLAEMI PQFFYKNVI FTLSLFWYGI YNNFDGSYLF
EYTYLTFYNLAFTSVPVILLAVLDQDVSDTVSMLVPQLYRVGILRKEWNQTKFLWYMLDG
VYQSVICFFFFPYLAYHKNMVVTENGLGLDHRYPVGVFVTAIAVTSCNFYVFMEQYRWDWF
CGLFICLSLAVFYGTWTGIWTSSSSSNEFYKGAARVFAQPAYWAVLFGVLFCLLPRFTID
CIRKIFYPKDIIEIVREMWRGDFDLYPQGYDPTDPSRPRINEIRPLTDFKEPISLDTHFD
GVSHSQETIVTEEIPMSILNGEQSRKGYRVSTTLERRDQLSPVTTTNNLPRRSMASARG
NKLRTSLDRTREEMLANHQLDTRYSVERARASLDLPGINHAETLLSQRSRDR

SEQ ID NO:28

YDR338C

>sp| Q05497 |YD338\_\_YEAST Uncharacterized transporter YDR338C
OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=YDR338C
PE=1 SV=1

MAGILSKTLSEVHPSLRNMGIGNTHRRISLGFLLPKNKNPLVRKFRARTRNIDQRSFR
SLTDDFGSNVHEPNPYLGNIDEEDPDLYHDEEDGELSRTISLPSRVSETPELSPQDQVDWI
LHEHERRYSSVCNSDNEEASQSNTPDRIQEYSGRELEYDEFMNRLQAQKQKLTRSAVTD
KGTSHHRRPSFVSVTSRGSVPYIYQEIENDSEALAEALASHVTFKSEARVLASYSFPLI
FTFLLLEQIFPMVCSLTVGHGKNELAAVSLASMTSNITLAI FEGIATSLDTLCPQAYGSG
RFYSVGVHLQRCIAFSLVI YIPFAVMWWYSEPLLSYI IPEKELINLTSRFLRVLILGAPA
YIFFENLKRFLQAQGI F DAGI YVLTICAPLNVLSYTLVWNKYIGVGF IGAAIAVVLNFW
LMFFLLLFYALYIDGRKCWGGFSRKAFTHWNDLGHAFSGIIMLEAEELSYELLTLFSAY
YGVSYLAAQSAVSTMAALLYMIPFAIGISTSTRIANFIGAKRTDFAHISSQVGLSFSFIA
GFINCCILVFGRNLIANIYSKDPEVIKLIQAQVLPVGVIVQNFDSLNAVAGSCLRQGMQS
LGSIVNLMAYYLFGIPLALILSWFFDMKLYGLWIGIGSAMPLIIGLVEAYYVLPDWDKIM
TYAEILKETEDDEVDSDYEYLTDSDDPDENTALLGA

SEQ ID NO: 29

YDR40 6W

>sp|Q04182 |PDR15\_YEAST ATP-dependent permease PDR15 OS=Saccharomyces
cerevisiae (strain ATCC 204508 / S288c) GN=PDR15 PE=1 SV=1

MSSDIRDVEERNRSDSSSSSSSSNSAAQSIGQHPYRGFDSEAAERVHELARTLTSQSLLYT
ANSNNSSSSNHNAHNADSRVSTDMEGVNPVFTNPDTPGYNPKLDPNSDQFSSTAWVQN
MANICTSDPDFYKPYSLGCVWKNLSASGDSADVSYQSTFANIVPKLLTKGLRLLKPSKEE
DTFQILKPMGDCLNPGELLVVLGRPGSGCTTLLKSISSNSHGFKIAKDSIVSYNGLSSSD
IRKHYRGEVVYNAESDIHLPHLTVYQTLFTVARMKTPQNRKIGVDREAYANHVTEVAMAT
YGLSHTRDTKVGNDLVRGVS GGERKRV SIAEVAICGARFQCWDNATRGLDSATALEFIRA
LKTQADIGKTAATVAIYQCSQDAYDLFDKVCVLDGQYLYFGPAKDAKKYFQDMGYCCPP
RQTTADFLTSITSPTERI ISKEFIEKGRVPTPKDMAEYWLQSESYKNLIKDIDSTLEK
NTDEARNI 1RDAHAKQAKRAPPSPYVVNYGMQVKYLL I RNFWRMKQSA SVTLWQVI GN
SVMFILGSMFYKVMKNDTSTFYFRGAAMFFAILFNAFSCLEI FSLYETRPITEKHRT
YSLYHPSADAFASVLSMPPKLI TAVCFNI IFYFLVDFRRNGGVFFYFLINVIATFTLS
HLFRCVGLSKTKTLQEAMVPASMLLLAISMYTGFAIPKTKILGWSIWIWIINPLAYLFE
SLMINEFHDRRFPCAQYI PAGPAYQNI TGTQRVCSAVGAYPGNDYVLGDDFLKESYDYEHKH
KWRGFGIGMAYVVFVYVYLILCEYNEGAKQKGMVVF LRSKIKQLKKEGKLQEKHRPGD
IENNAGSSPDSATTEKKILDDSSEGSDSSSDNAGLGLSKSEAI FHWDRDLCYDVP IKGQR
RILNNVDGWVKPGTLTALMGASGAGKTTLLDCLAERVMTMGVITGNI FVDGRLRDESFPRS
IGYCQQQDLHLKTATVRESLRFSA YLRQPSSVSI EEKNRYVEEVIKILEMQQYSDAVVGV
AGEGLNVEQRKRLTIGVELAARPKLLVFLDEPTSGLDSQTAWDTCQLMRKLATHGQAILC
TIHQPSAILMQQFDRLLFLQKGGQTVYFGDLGEGCKTMIDYFESKGAHKCPPDANPAEWM

LEVVGAA PGSHATQDYNEVWRNSDEYKAVQEELDWMEKNLPGRSKEPTAEEHKPFAASLY
YQFKMVTIRLFQQYWRSPDYLWSKFI LTI FNQVFIGFTFFKADRSLQGLQNQMLSI FMYT
VIFNPILQQYLPSFVQQRDLYEARERPSRTF SWLAFFLSQI IVEI PWNILAGTIAYCI YY
YAVGFYANASAAGQLHERGALFWLFSIAFYVYIGSMGLLLMISFNEVAETA AHMGTLLFTM
ALSFCGVMATPKVMPRFWIFMYRVSP LTYMI DALLALGVANVDVKCSNYEMVKFTP PPSGT
TCGDYMASYIKLAGTGYLSDPSATDICSFCAVSTTNAFLATFSSHYRRWRNYGIFICYI
AFDYIAATFLYWLSRVPKKNKGKISEKPKK

SEQ ID NO: 30

YDR536W

>sp|P39932|STL1\_YEAST Sugar transporter STL1 OS=Saccharomyces
cerevisiae (strain ATCC 204508 / S288c) GN=STL1 PE=1 SV=2
MKDLKLSNFKGKFI SRTSHWGLTGKRLRYFITIASMTGFSLFGYDQGLMASLITGKQFN Y
EFPATKENGDRHATV VQGATTSCYELGCFAGSLFVMFCGERIGRKP LILMGSVITIIG
AVISTCAFRGYWALGQFI IGRVVTGVGTGLNTSTI PVWQSEMSKAENRGLLVNLEGSTIA
FGTMIAWIDFGLSYTNSSVQWRFPVSMQIVFALFLLAFMIKLPESPRWLISQSRTEEAR
YLVGTLDDADPNDEEVITEVAMLHDAVNRTKHEKHSLSLFSRGRSQNLQRALIAASTQF
FQQFTGCNAAIYYSTVLFNKTIKLDYRLSMIIGGVFATIYALSTIGSFFLIEKLGRRKLF
LLGATGQAVSFTITFACLVKENKENARGAAVGLFLFITFFGLSLLSLPWIYPPEIASMKV
RASTNAFSTCTNWL CNFAVVMFTPIFIFIGSGWGCYLFFAVMNYLYIPVIFFFYPETAGRS
LEEIDI IFAKAYEDGTQPWRVANHLPKLSLQEVEDHANALGSYDDEMEKEDFGEDRVEDT
YNQINGDNSSSSSNIKNEDTVNDKANFEG

SEQ ID NO: 31

YEL031W

>sp|P39986|ATC6\_YEAST Manganese-transporting ATPase 1
OS=Saccharomyces cerevisiae {strain ATCC 204508 / S288c) GN=SPF1
PE=1 SV=1
MTKKS FVSSPIVRDSTLLVPKSLIAKPYVLPFFPLYATFAQLYFQQYDRYIKGPEWTFVY
LGTLVSLNILVMLMPAWN VKIKAKFNYSTTKNVNEATHIL IYTTPNNGSDGIVEIQRVTE
AGSLQTF FQFQK RFLWHENEQVFSSPKFLVDESPKIGDFQKCKGHSGDLTHLKRLYGEN
SFDIPIPTFMELFKEHAVAPL FVFQVFCVALWLLDEFWYYSLFNLFMI ISMEAAAVFQRL
TALKEFR TMGIKPYTIN VFRNKKWVALQTNELLPMDLVSITRTAEESAIPCDLILLDGSA
IVNEAMLSGESTPL LKESIKLRPSEDNLQLDGVDKIAVLHG GTKALQVTPPEHKS DI PPP
PDGGALAI VTKTG FETSQGS LVRVMI YSAERVSVDNKEALMFI LFLIFAVIASWYVWVE
GTKMGRIQSKLILDCILI ITSVPPELPMELTMAVNSSLAALAKFYVYCTEPFRI PFAGR
IDVCCFDK TGTLTGEDLVFEGLAGISADSENIRHLYSAAEPESTILVIGAAHALVKLED
GDIVGDPMEKATLKA VGWAVERKNSNYREGTGKLDI IRRFQFSSALKRSASIASHNDALF
AAVKGAPETIRERLSDIPKNYDEIYKSFTRSGSRVLALASKSLPKMSQSKIDDLNRDDVE
SELT FNGLIFHCPLKDDAIETIKMLNESSHRSIMITGDNPLTAVHVAKEVGIVFGETLI
LDRAGKSDDNQ LFRDVEETVS IPFDPSKDTFDH SKLFDRYDIAVTGYALNALEGHSQ LR
DLLRHTWVYARVSPSQEFLLNTL KDMGYQTL MCGDGTNDVGALKQAHVGIALLNGTEEG
LKKLGEQRRL EGMKMMYIKQTEFMARWNQPQPPVPEPIAHLFP PGPKNPHYLKALESKGT
VITPEIRKAVEEANSK PVEVIKPNGLSEKKPADLASLLLNSAGDAQGDEAPALKLGDASC
AAPFTSKLANVSAVTNI IRQRCALVNTIQMYKILALNCLISAYSLSI IYMAGVKFGDGQ
ATVSGLLLSVCFLSISR GKPLEKLSKQRPQSGIFNVYIMGSILSQFAVHIATLVYITTEI
YKLEPREPQVDLEKEFAPS LLNTGIFIIQLVQQVSTFAVNYQGE PFRENIRSNKGMYYGL
LGVTGLALASATEFLPELNEAMK FVPMTDDFKIKLTLTLLLDFFG SWGVEHFFKFFFMDD
KPSDISVQQVKIASK

SEQ ID NO: 32

YER166W

>spi P32660 !A?C5\_YEAST Phospholipid- transporting ATPase DNF1  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=DNF1  
 PE=1 SV=2

MSGTFHGDGHAPMSPFEDTFQFEDNSSNEDTHIAPTHFDDGATSNKYSRPQVSFNDETPK  
 NKRE DAEETFNDDEYDNHSFQPTPKLNNGSGTFDDVELDNDSGEPHTNYDGMKFRFMG  
 TKRNKKGNPIMGRSKTLKWARKNIPNPFEDFTKDDIDPGAINRAQELRTVYYNMLPKDM  
 IDEEGNPIMQYPRNKIRTTKYTPLTFLPKNILFQFHNFANVYFLVLIILGAFQIFGVNTP  
 GLSAVPLVVIVI ITAIKDAIEDSRRTVLDLEVNNTKTHILEGVENENVSTDNISLWRRFK  
 KANSRLLFKFIQYCKEHLTEEGKKKRMQRKRHEL RVQKT VGTSGPRSSLD SIDSYRVSAD  
 YGRPSLDYDNLEQGAGEANIVDRSLPPRTDCKFAKNYWKGVKVDIVRIHNDEI PADII  
 LLSTSDTDGACYVETKNLDGETNLKVRQSLKCTNTIRT SKDIARTKFWIESEGPHSNLYT  
 YQGNMKWRNLADGEIRNEPITINNVLRLGCTLRNTKWAMGVVMFTGGDTKIMLNSGITPT  
 KKSRI SRELNF SVVINFVLLF ILCFVSGIANGVYYDKKGRSRFSYEFGTIAGSAATNGFV  
 SFWVAVILYQSLVPI SLYISVEI IKTAQAAF IYGDVLLYNAKLDYPCTPKSWNISDDLQ  
 VEYIFSDKTGTLTQNVMEFKKCTINGVSYGRAYTEALAGLRKRQIDVETEGREKAEIA  
 KDRDTMIDELRALSGNSQFYPEEVT FVSKEFVRDLKGASGEVQRCCEHFMLALALCHSV  
 LVEANPDNPKKLDLKAQSPDEAALVATARDVGF SFGVKT KGLI IEMQGIQKEFEILNIL  
 EFNSSRKRMS CIVKI PGLNPGDEPRALLICKGADS IYSRLSRQSGSNSEAILEKTALHL  
 EQYATEGLRTL CIAQRELSWSEYEWNEKYDIAAASLANREDELEV VADSIERELILLGG  
 TAIEDRLQDGVPCIELLAEAGIKLWVLTGDKVETAINIGFSCNLLNEMELLVIKTTGD  
 DVKEFGSEPEIVDALLSKYLKEYFNLTGSEEEIFEAKKDHEFPKGNYAIVIDGDALKLA  
 LYGEDIRRKFLLLCKNCRAVLCCRVSQSQA AVVKLVKDSLDMVMTLAIGDGSNDVAMIQS  
 ADVGIGIAGEEGRQAVMCSDYAIGQFRYLARLVLVHGRWSYKRLAEMIPEFFYKNMIFAL  
 ALFWYGIYNDFDG SYLYEYTYMMFYNLAF TSLPVI FLGILDQDVNDTISLVVPQLYRVGI  
 LRKEWNQRKFLWYMLDGLYQSIICFFFPYLVYHKNMIVTSNGLGLDHRYPVGVVYVTTIAV  
 ISCNTYVLLHQYRWDWFSGLFIALSCLVVF AWTGIWSSAIASREFFKAAARI YGAPSWFA  
 VFFVAVLFCLLPRFTYDSFQKFFYP TDEIVREM WQHGHFDHYPPGYDPTDPNRPKVTKA  
 GQHGEKI IEGIALSDNLGGSNYSRDSVVT EEPMTFMHGEGD GSPSGYQKQETWMTSPKET  
 QDLLQSPQFQQAQTFGRGPSTNVRSSLDRTREQMIATNQLDNRY SVERARTSLDLPGVTN  
 AASLIGTQQNN

SEQ ID NO: 33

YFL011W

>sp| P43581 IHXT10JYEAST Kexcse transporter HXT10 OS=Saccharomyces  
 cerevisiae (strain ATCC 204508 / S288c) GN=HXT10 PE=1 SV=1  
 MVSSSVSILGTSAKASTSLSRKDEIKLTPETREASLDIPYKPIIAYWTVMGLCLMIAFGG  
 FIFGWDGTISGFINQTD FKRFRFGELQRDGSFQLSDVRTGLIVGI FNIGCALGGLTLGRL  
**GDIYGRKIGLMCVILVYWGIVIQIASSDKWYQYFIGRIVSGMGVGGVAVLSPTLISEIS**  
 PKHLRGTCVSFYQLMITLGI FLGYCTNYGTTKYSNSIQWRVPLGLCFAWAIFMVMIGMVMV  
 PESPRYLVEKGKYEARRSLAKSNKVTVDPGVVF EFDTIVANMELERAVGNASWHELFS  
 NKGAILPRVIMGIVIQSLQQLTGCNYFFYYGTTIFNAVGMQDSFETSIVLGAVNFASTFV  
 ALYIVDKFGRRKCLLWGSASMAICFVIFATVGVTRLWPQKDKQPSQSAGNVMIVFTCF  
 IFSFAITWAPIAYVIVAETYPLRVKNRAMAIAVGANWMMWGFLIGFFTPFITRSIGFSYGY  
 VFMGCLI FSYFYVFFFVCE TKGLTLEEVNEMYEERIKPWKSGGWI PSSRRTPOPTSSTPL  
 VIVDSK

SEQ ID NO: 34

YGL006W

>sp| P38929 |ATC2\_\_YEAST Calcium-transporting ATPase 2 OS=Saccharomyces  
 cerevisiae (strain ATCC 204508 / S288c) GN=PMC1 PE=1 SV=1  
 MSRQDENSALLANNENKPSYTG NENGVYDNFKLSKSQLSDLHNP KSIRSFVRLFGYESN

SLFKYLKT DKNAGISLPEISNYRKTNRYKNYGDNS LPERI PKSFLQLVWAA FNDKTMQLL
TVAAVVSFVLGLYELWMQPPQYDPEGNKIKQVDWIEGVAIMIAVFFVVLVSAANDYQKEL
QFAKLNKKKENRKII VIRNDQEILISIHVVLVGDVLSLQTDVVPADCVMI SGKCEADES
SITGESNTIQKFPVDNSLRDFKKFNSIDSHNHKPLDIGDVNEDGNKIADCMLISGSRIL
SGLGRGVITSVGINSVYGQTM TSLNAEPESTPLQLHLSQLADNISVYGCVSAI I LFLVLF
TRYLFYII PEDGRFHDLDPAQKGSKFMNIFITSITVI VVAVPEGLPLAVTLALAFATTRM
TKDGNLVRVLRSCETMGSATAVCSDKTGTLTENVM TVVRGFPGNSKFDDSKSLPVSEQRK
LNSKKVFEENCSSSLRNDLLANIVLNSTAFENRDYKKNKNTNGSKNMSKNLSFLDKCKS
RLSFFKKGNREDDQLFKNVNKGREQEPFIGSKTETALLSLARLSLGLQPGELOYLQDQP
MEKFNIEKVVQTI PFESSRKWAGLVV KYKEGKNKPFYRFFIKGAAEIVSKNCSYKRNSD
DTLEEINEDNKKETDDEIKN LASDALRAISVAHKDFCECD SWPPEQLRDKDSPNIAALDL
LFNSQKGLILDGLLGIQDPLRAGVRESVQQCQRAGVTVRMVTGDNILTAKAIARNCAILS
TDISSEAYSAMEGTEFRKLT KNERIRILPNLRVLARSSPEDKRLLVETLKGMDVVAVTG
DGTNDAPALKLADVGFSGMISGTEVAREASDIILMTDDFS AIVNAIKWGRCSVSVSIK KFI
QFQLIVNITAVILTFVSSVASSDETSVLTAVQLLWINLIMDTLAALALATDKPDPNIMDR
KPRGRSTSLISVSTWKMILSQATLQLIVTFILHFYGP ELPFFKKHEDEITSHQQQLNAMT
FNTFVWLQFFTMLVSRKLD EGDGISNWRGRISAANLNFFQDLGRNYFLTIMAIIGSCQV
LIMFFGGAPFSIARQTKSMWITAVLCGMLSLIMGVLVRI CPDEVAVKVPAAAFVQRFKYV
FGLEFLRKNHTGKHDDEEALLEESDSPESTAFY

SEQ ID NO: 35

YGL013C

>sp|P12383|PDR1\_\_YEAST Transcription factor PDR1 OS=Saccharomyces
cerevisiae (strain ATCC 204508 / S288c) GN=PDR1 PE=1 SV=2
MRGLTPKNGVHIETGPDTESSADSSNFSTGFSGKIRKPRSKVSKACDNCRKRKIKCNGKF
PCASCEIYSCECTFSTRQGGARIKLNHKTSL EGTTVQVKEETDSSSTSFSNPQRCTDGPC
AVEQPTKFFENFKLGGRSSGDNSGSDGKNDDDVNRNGFYEDDSESQATLTSLOTTLKNLK
EMAHLGTHVTS AIESIELQISDLLKRWEKVRTELATTKFYPNKSIETQLMKNKYCDVV
HLTRYAAWSNNKKDQDTSSQPLIDEIFGLYSPFQFLSLQIGIGKCFQNYRSKSKCEIFPRT
AKETIYIMLRFFDVCFHHINQGCVSIANPLENYLQKMNLLPSTPSSISSAGSPNTAHTKS
HVALVINHLPQPFVRNITGISNSELLSEMNDI SMFGILLKMLDMHKNSYQNFLMEITSN
PSVAKNTQSIDVLQEFIH YCQAGEALIALCYSYYNSTLYNYVDFTCDITHLEQLLYFLDL
LFWLSEIYGFEKVLNVAVHFVSRVGLSRWEFYVGLDENFAERRRNLWWKAFYFEKTLASK
LGYP SNIDDSKINCLLPKNFRDVGFLDNRDFIENVHLVRRSEAFDNMCISDLKYYGELAV
LQIVSHFSSSVLFNEKFTSIRNTSKPSVVR EKLLEFEVLEIFNETEMKYDAIKEQTGKLF
IAFSKSTELKVSREDKIMASKFVLFYEH HFCRMVNESDNIVARLCVHRRPSIL IENLKI
YLHKIYKSWTDMNKILLDFDNDYSVYRSFAHYSISCIILVSQAFSVAEFIKVNDVVMIR
VFKRFLDIKIFSENETNEHVNSQSFKDYTRAFSFLTIVTRIMLLAYGESSTNLDVISK
YIDENAPDLKGI IELVLDTNSCAYRFLLEPVQKSGFHLTVSQMLKNRKFQEP LMSNEDNK
QMKHNSGKNLNPDLPSLKTGTSCLLNGIESPQLPFNGRSAPSPVRNNSLPEFAQLPSFRS
LSVSDMINPDYAQPTNGQNNTQVQSNKPINAQQQIPTSVQVPFMNTNEINNNNNNNNNNK
NNINNINNNNSNNFSATSFNLGTLDEFVNNGDLEDLYSILWSDVYPDS

SEQ ID NO: 36

YGL255W

>sp|P32804|ZRT1\_\_YEAST Zinc-regulated transporter 1 OS=Saccharomyces
cerevisiae (strain ATCC 204508 / S288c) GN=ZRT1 PE=1 SV=1
MSNVTTTPWVKQWDPSEVTLADKTPDDVWKTCV LQGVYFGGNEYNGNLGARISSVFVILFV
STFFTMFPLISTKVKRLRIPLYVYLF AKYFGSGVIVATAFIHLM DPAYGAIGGTT CVGQT
GNWGLYSWCPAIMLTSLTF TFLTDLFSSVWVERKYGLSHDHTHDEIKDTVVRNTAAVSSE
NDNENGTANGSHDTKNGVEYYEDSDATSM DVVQS FQAQFYAFLILEFGVIFH SVMIGLNL
GSVGDEFSSLYPVLVHFHQSFEGLGIGARLSAIEFPRSKRWWPWALCVAYGLTTPICVAIG

LG¥RTRY¥5G5YTAL¥I5G¥LDAI5AGILLYTGL¥ELLARDFIFNPQRIKDLRELSFN¥I  
CTLFGAG IMALIGKWA

SEQ ID NO: 37

YGR125W

>sp|P53273|YG35\_\_YEAST Uncharacterized vacuolar membrane protein  
YGR125W OS=Saccharomyces cerevisiae (strain ATCC 204508 / 5288c)  
GN=YGR125W PE=1 S¥=1  
MGRITIRRRRSNSSLSEAISS¥SLGINQDSS¥NKMHRAS¥SAMPPLCRSYMSGFFFTGGNSP  
MINNLSDSKLPISNKQHPK¥IHGSENLRQTAQLSNEFCSSS¥EENSPTIKDYMDI IGNG  
DRKDDQSMRTIEENIDEEYSDEYSRLLLSPASSN¥DDDRNRGLQNSSLPELEDGYAGGYQ  
SLRPSHNLRFPRNLWHMCTSFPSKFAHYLPAA¥LGLLLNILDALSYGMIIFPITEPVFS  
HLGPTGISMFYISTI ISQA¥YSGGWSSFPSPGIGSEMIEITPFYHTMALAIKEALAGNDDE  
IITTTIFCY¥ISSMLTG¥¥FYALGKLRGKI¥GFFPRHILIGCIGG¥GYFLI ITGIEVTT  
R¥AKFEYSWPFSSGLFTDYDTLAKWLLPVLLT¥¥LIGTQRYFKNSL¥LPSFYILTLVLFH  
FIVAIIPTLSLDALRQAGWI FPIANS DSKWYDHYRLFN¥HK¥HWSL¥LQOI PTMMALTFF  
GILH¥PIN¥PALAMSLQMDKYDVDRELIAHGYSNFFSGLLGS¥QNYL¥YTNS¥LFIIRAGA  
DSPFAGFLLIALTICIMI IGP¥I ISFIPICI¥GSLIFLLGYELL¥EAL¥DTWNKLNRFY  
LTV¥II¥FTMGIFDF¥LGII¥GILIACFSSFL¥DSTKLQTINGEYNGN¥ARST¥YRDYVQT  
KFLDGIGEQI Y¥LKLQNLFFGTIISIEEKIERLLQISNKDATKRRIKYLILDFKNINAD  
NIDYSAEAGFNRIKRFTETKRIKLI ISSIKERDRIYNAFN¥GLLND¥ELFADLNSALEW  
CENEFLFQYKQLRKKAKERLEEGKQNN¥¥SA¥IAATKNKKIDTIGNGLNRGSNGDTARNL  
MSLPTNTPRNYQILS¥AQNVF¥NDEQA¥KNFKKEYKDDEP¥LPILLFALKQYRPDIISE¥  
QKVREKEIKFWAQLCPYFTRRRRLASQSHLLHADNIFFL¥ETGMLKATYELPQGTLYEIFS  
NGTCFGKIIAPGNAMPREQKLTIEETETDS¥LW¥IDSSSLNKLKEDNLALYVE¥ALM¥MCI  
KDRFKELLGYTL¥SA

SEQ ID NO: 38

YGR181W

>sp|P53299|TIM13\_\_YEAST Mitochondrial import inner membrane  
translocase subunit TIM13 OS=Saccharomyces cerevisiae (strain ATCC  
204508 / 5288c) GN=TIM13 PE=1 S¥=1  
MGLSSI FGGGAPSQQKEAATTAKTTPNPIAKELKNQIAQELAYANATEL¥NKISENCFEK  
CLTSPYATRNDACIDQCLAKYMRSWN¥ISKAYISRIQNASASGEI

SEQ ID NO: 39

YGR217W

>sp|P50077|CCH1\_\_YEAST Calcium-channel protein CCH1 OS=Saccharomyces  
cerevisiae (strain ATCC 204508 / S288c) GN=CCH1 PE=1 S¥=1  
MQGRKRTLTEPFEPNTNPFQDAAVMTEN¥EDNSETDGNRLESKPQAL¥PPALNIVPPES  
SIHSTEEKKGDEYNGNDKSSSLISNIFRTRVGRSSHENLSRPKLSLKTASFGAAESSRN  
¥SPSTKSASSSQYIDLNDERLRRRSFSSYSRSSRRVSNSPSSTDRPPRSK¥LSLIAA  
DDMDDFEDLQKGFKSAIDEEGLTWLPQLKSEKSRP¥SD¥GEDRGEQESIPD¥HTPN¥G  
ASATPGSIHLTPEPAQNGSVSEGLEGSINNSRKKPSPKFFHLLSPQKEDKDQTE¥IEYAE  
DILDFTLQKLESRPVLYGHSLGVFSPTNPLRIKIARFLLHRRYSLLYNTLLTFYAIL  
LAIRTYNPHN¥¥FLYRFNSWTDYFIFILSACFTGNDIAKIIAFGFWDSEMFKAYGREYK  
SILQRSGIMKLYIYLREKYGRKLIDFII PFRIISPGEETKYQRSSLSTSLTKPYGAKENQ  
RPFQTPRAFARSSWNRIDL¥SS¥SFWLGMFLSIKSYDTKTGIRIFKPLAILRILRLVN¥D  
TGMPSILRGLKYGI PQL¥NVSSML¥YFWIFFGILG¥QIFQGSFRRQCVWFNPEDPTDITYQ

YDMQFCGGYLDPYTKRKQNYIYEDGSEGSY5KGFLCPQY5KC¥5NANPYNGRI5FDNI¥N  
 SMELVVFVIMSANTFTDLMYTMDSEMAACLFVFCIFVLTIWLLNLLIAVLVSSFEIAN  
 EEYKKKKFIYGSRKTGYVARIVTGYWKYFKLKANQTKFPNWSQKGLAIYSHVEFIFVILI  
 ICDIGMRASVKVSTSANCNILLKTDGRGISIVLFIESLARLVLYLPNMWKFLTKPSYVYD  
 FIISII TLVI SCLAVEGVLGHMYAWLS IFHISRFRVI ISFNLTKKLWKQI LSNGVMI WN  
 LSSFYFFFTFLVAIIMAVYFEGVIPPEEMADQPFMGYSLPNSFLSLFIIIGSTENWTDILY  
 ALQKHSPNISSTFFCSVFFI IWFLLSNSVILNIFIALISESMEVKEEEKRPQIKHYLKF  
 VYPQKIQEYTHASLVARIRKKFFGGHRNEDTRDFKQFLMRGTAIMNIAQNMGELADEFK  
 PPSENLFKKGLSKLTIGVPSLKRRLRMFANNPFYKNSDVVFTETNDINGRTYILELNEYED  
 EKLDYLKKYPLFNYSYFFSPQHRFRRFCQRLVPPSTGKRTDGSRFFEDSTDLYNKRSYF  
 HHIERDVVFVIFALATILLIVCSCYVTPLYRMHHKMGTNWSSALDCAFIGAFSIEFIVK  
 TVADGFYSPNAYLRNPWNFIDFCVLI SMWINLIAYLKNNGNLSRIFKGLTALRALRCLT  
 ISNTARQTFNLVMDGLNKIFEAGLISLSLLFPFTVWGLSIFKGRGLGTCNDGSLGRADCY  
 NEYSNSVQWDIMSPRVYQQPYLHLD SFASAFSSLYQIISLEGWVDLLENMMNSSGIGTP  
 ATVMGSAGNALFLVLFNFLSMVFI LNLFVSVFVNNQARTTGSAYFTIEEKAWLESQKLLS  
 QAKPKAI PNLIELSRVRQFFYQLAVEKKNFYAS FLQVVL YLHI IMLL3RSYN PGNLIGY  
 QGVYFMFSTSVFLIQEALHMCGEPRLYFRQKWN SIRLSIII IAFIMNAVAFHVPASHYW  
 FHNKIGFFLLVIFLFI IPQNDTLELLETAMASLPPILSLTYTWGVFLVYAIALNQIFG  
 LTRLGSNTTDNINFRVIVIKSMIVLFRCSFGEGWNYIMADLTVSEPYCSSDDNSTYTDGCS  
 ETYAYLLLMSWNIISMYIFVNMVSLI IGNSYVYRSGGSRSGINRSEIKKYIEAWSKFD  
 TDGTGELELSYLPRIMHSFDGPLSFKIWEGRLTIKSLVENYMEVNPDDPYDKIDLIGLN  
 KELNTIDKAKIIQRKLQYRRFVQSIHYTNAYNGCIRFSDLLQIPLYTAYSARECLGIDQ  
 YVHHLYLKGVKDYLENQRNFDVLEMVVTRWKFHCRMKRTIEPEWDVKDPTVSSHISNIN  
 VNLEPAPGILEREPIATPRMDYGVNNFMWSPRMNQDSTMEPPEEPIDNNDSDANDLIDR

SEQ ID NO: 40

YGR224W

>sp|P50080|AZR1\_YEAST Azole resistance protein 1 OS=Saccharomyces  
 cerevisiae (strain ATCC 204508 / S288c) GN=AZR1 PE=1 SV=1  
 MKGEPKTYSMSDLSYYGEKAQQQNEKQQKQYVVRNSTQSTSKQNVSVLEDNASESNEL  
 PKGFILYASLIALALSLFLAALDIMIVSTIIIEVAKQFGSYSEIGWLFTGYSLPNALLAL  
 IWGRIATPIGFKETMLFAIVIFEIGSLISALANSMSMLIGGRVIAGVGGCGIQSLSFVIG  
 STLVEESQRGILIAVLSCSFAIASVVGPFLLGGVFTSSVTWRWCFYVNLPIGGLAFFLFLF  
 FYNPGLSTFQETMDNIRKFPSQFIEIVRVNAVYHLLKIKGFSKLNWGRKPFMELI FMYDII  
 EFVFCAGFTCILLAFTFGNRYAWNSASI IILFIIGIVLWLAGIYDFLVFPKFNIKVA  
 TPHYQPLMSWTNIKKPGIFTVNIALFLTACAGYISQFTYIVQYFQLIYNDSAWRAAVHLVA  
 CIISTVVTAILCGAITDKTRQIKPIIVISSIFGVVAGILTLLNANNNSAHIGLLILPG  
 VAFGGLAQSSMLASQIQLDKKSPTFRSDFVSITTFNTFCKNLGQALGGVISNTVFSAAAI  
 KKLTKANIQLPDGTTVDNLVIYRQTNFDGSHSKLGNIISES LTDVFYMALGFYALSLIFA  
 VFASNKKVTASLR

SEQ ID NO: 41

YGR281W

>sp|P53049\_YOR1\_YEAST Oligomycin resistance ATP-dependent permease  
 YOR1 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c)  
 GN=YOR1 PE=1 SV=1  
 MTITVGDAVSETELNKSQNVVLSPKASASSDISTDVKDTSSSWDDKSLLPPTGEYIVDR  
 NKPQTYLNSDDIEKVTESDIFPQKRLFSFLHSKKIPVQTDDERKIYPLFHTNI ISNMF  
 FWWVLPILRVGYKRTIQPNDLFKMDPRMSIETLYDDFEKNMIYYFEKTRKKYRKRHPEAT  
 EEEVMENAKLPKHTVLRALLFTFKKQYFMSIVFAILANCTSGFNPMITKRLIEFVEEKAI  
 FSHMHVNKGIGYAIACLMFMVNGLTFNHFFHTSQLTGTVQAKSILTKAAMKMFNASNYA

RHCFPNGKVISFVTTDLARIEFALSFPFLAGFPAILAICIVLLIVNLGPALYGIGIFF  
GGFFISLSLFAFKLILGFRIAANIFTDARVTMMREVLNNIKMIKYTTWEDAYEKNIQDIRTK  
EISKVRKMQLSRNFILIAMAMSLPSIASLVTFILAMYKVNKGGRQPGNIFASLSLQVLSLQ  
MFFLPPIAIGTGIDMIIGLGRQLQSLEAPEDDPNQMIEMKPSPGFDPKLALKMTHCSFEWE  
DYELNDAIIEAKGEAKDEGKKNKKRDKDTWGKPSASTNKAKRLDNMLKDRDGPEDLEKTS  
FRGFKDLNFDIKKGEFIMITGPIGTGKSSLLNAMAGSMRKTGKVEVNGDLLMCGYPWIQ  
NASVRDNI IFGSPFNKEKYDEVVRVCSLKADLDILPAGDMTEIGERGITLSGGQKARINL  
ARSVYKKKDI YLFDDVLSAVDSRVGKHIMDECLTGMLANKTRILATHQLSLIERASRVIV  
LGTGQVDIGTVDELKARNQTLINLLQFSSQNSEKEDEEQEAVVAGELGQLKYESEVKEL  
TELKKKATEMSQTANSKIVADGHTSSKEERAVNSISLKIYREYIKA AVGKWFIALPLY  
AILVVGTTFCSLFSSVWLSYWTENKFKNRPPSFYMGLYSFFVFAAFIFMNGQFTILCANG  
IMASKWLNLRVAVKRI LHTPMSYIDTTP LGRILNRFTKDTDSLDELTESLRLMTSQFANI  
VGVCVMCIVYLPWFIAIAI PFLVI FVLIADHYQS SGRE IKRLEAVQRS FVYNNLNEVLGG  
MDTIKAYRSQERFLAKSDFLINKMNEAGYLVVVLQRWVGIFLDMVAIAFALIIITLLCVTR  
AFPISAASVGVLLTYVLQPLGLLNTILRAJMTQTENDMNSAERLVTYATELPLEASYRKPPE  
MTPPESWPSMGEI IFENVDFA YRPGLPI VLKLNLNLIKSGEKIGICGRTGAGKSTIMSAL  
YRLNELTAGKILIDNVDISQLGLFDLRRKLAII PQDPVLFRTIRKNLDPFNERTDDELW  
DALVRGGAIKDDLPEVKLQKPDENGTHGKMHK FHLQAVEEEGNSFSLGERQLLALTRA  
LVRQSKILILDEATSSVDYETDGKIQTTRIVEEFGDCTILCIAHRLKTIVNYDRILVLEKG  
EVAEFDTPWTLFSQEDSIFRSMCSRSGIVENDFENRS

SEQ ID NO: 42  
YHL016C

>sp|P33413|DUR3\_YEAST Urea active transporter OS=Saccharomyces  
cerevisiae (strain ATCC 204508 / 5288c) GN=DUR3 PE=1 S#=2  
MGEFKPPLPQAGYAIVLGLGAVFAGMMVLTYYLLKRYQKEIITAEFFTAGRSVKTLV  
AAAVSSWIWCSTLLTSSTKEYADGIFGGYAYAAGACFQIIAFAILAIKTKQMAPNAHTY  
LELVTRYGKIGHGCYLFYAIATNILVTSMLLTSGSAVFSDLTGMNTIASCFLLPVGVVV  
YTLFGGIKATFLTDYMHTCVIII IVLVFAFKVYATSDVLGSPGKVYDLVREAAKRHPVDG  
NYQGEYMTMTSKSAGILLI INLIGNFGTVFLDNGYWNKAISASPAASLKAYAIGGLAWFA  
VPSLISLTMGLACLAVETSPNFPTYDPDPLTSFQANSGLVLPAAAIAIMGKGGAVASLLMI  
FMAVTSAMSAELI AVSSVFTYDIYREYIDPRASGKLI YTSHVACI FFGLAMSGFSVGLY  
YGGISMGYIYEMMGI ISSAVLPVVLTLCSKDMNLVA AVVSPILGTGLAIMSWLVCTKSL  
YKELTVDTTFMDYPMLTGNLVALLSPAIFIPILTYVFKPQNF DWEKMKDITRVDETAELV  
QADPDIQLYDAEANDKEQEEETNSLVS DSEKNDVRVNNEKLI EPNLGVVISNAIFQEDDT  
QLQNELDEEQRELARGLK IAYFLCVFFALAFVWVWMPMPYGSKYI FSKKFFTGWVVVMI I  
WLFFSAFAVCIYPLWEGRHGIYTTLRGLYWDLSGQTYKLREWQNSNPQDLHVVTSSQISAR  
AHRQSSHFQVDEII

SEQ ID NO: 43  
YIL088C

>sp|P40501|AVT7\_YEAST Vacuolar amino acid transporter 7  
OS=Saccharomyces cerevisiae (strain ATCC 204508 / 5288c) GN=AVT7  
PE=1 SV=1  
MEATSSALSSTANLVKTI V GAGTLAIPYSFKSDGVLVGVILTLAAVTSGLGLFVLSKCS  
KTLINPRNSSFFTL CMLTYPTLAPIFDLAMIVQC FGVGLSYLVLIGDLFPGLFGGERNYW  
IIASAVIIIPLCLVKKLDQLKYSSILGLFALAYISILVFSHFVFELGKGELTNILRNDIC  
WVKIHDFKGLLSTFSI IFAFTGSMNLFPMINELKDNSMENITFVINNSISLSTALFLIV  
GLSGYLTFGNETLGNLMLNYDPNSIWIVIGKFC LGSMLILSFPLLFHPLRIAVNNV IWI  
EITYGGANPEEDPQVSEYTRASNLRPISMTVEDPAQPSDALDATSYNEQECLLPNGNFDN  
GSIESQENNNDERGTM AVAGDNEHHAPFVKSRFYWITALLLISMYTLALS VQSFALVLSF



VGATGSTSISFTLPGLLGYKLIGLDSLAIKMI PPKDRFYKRCSLLL VFYGLSVMFLSLY  
 VTVFNRSDEA

SEQ ID NO: 44

YJL093C

>sp| P40310 IT0K1\_\_YEAST Outward-rectifier potassium channel TOK1  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / 5288c) GN=TOK1  
 PE=1 SV=1

MTRFMNSFAKQTLGYGNMATVEQESSAQAVDSHSNNTPKQAKGVLAEEELKDALRFRDERV  
 SIINAEPSSTLFVFWFVSCYFPVITACLGPVANTISIAWKEKWRSLKNNNSVVTNPRSN  
 DTDVLMNQVKT VFDPPGI FAVNIISLVLGFTSNI ILMLHFSKKLTYLKSQNLINITGWITIA  
 GGMLLDVIVCSLNDMPSSIYSKTIGFWFACISSGLYLVTI ILTIHFIFIGYKLGKYPPTFN  
 LLPNERSIMAYTVLLSLWLIWAGMFGSLLHITYGNALYFCTVSLTLTVGLGDILPKSVGA  
 KIMVLI FSLSGVVMGLIVFMTRS IIQKSSGPI FFFHRVEKGRSKSWKHYMDSKKNLSE  
 EAFDLMKCIRQTASRKQHWFSLSVTIAIFMAFWLLGALVFKFAENWSYFNCI YFCFLCLL  
 TIGYGDYAPRTGAGRAFFVIWALGAVPLMGAILSTVGDLLFDISTSLDIKIGESFNKVK  
 SIVFNQRQALSFMVNTGEIFEESDTADGDLEENTTSSQSSQISEFNNDNNEENDSGVTS  
 PPASLQESFSSLSKASSPEGI LPLEYVSSAEYALQDSGTCNLRNLQELLKAVKKLHRICL  
 ADKDYTLFSFSDWSYIHKLHLRNITDIEEYTRGPEFWISPDTPKFPLNEPHFAFMMLFKN  
 IEELVGNLVEDEELYKVISKRKFLGHRKTL

SEQ ID NO: 45

YJL094C

>sp IP40309 IKHA1\_YEAST K (+)/H (+) an.tiporter 1 OS=Saccharomyces  
 cerevisiae (strain ATCC 204508 / S288c) GN=KHA1 PE=1 SV=1

MANTVGGILSGVNPFFHYNSSSPLTLFLFQACLILLVCNLIHIPFSMMRQPKVISEVISGV  
 ILGPTIFGQIPNYTNTIFPTSSI PGLNLVANLGI ILFMFFLGLEVDIAFIKKHLKALVI  
 GIVTLAVPFGFGLLAIPLFHTYANKTEGERHIKFSVFMVFI AVSISVTAFPVLCRILNE  
 LRLIKDRAGIVVLAAGIINDIMGWILLALSIISSAEGSPVNTVYILLITFAWFLIYFFP  
 LKYLLRWVLRIRTHELDRSKPS PLATMC ILFIMFI SAY FTDIIGVHPIFGAFIAGLVVPRD  
 DHYVVKLTERMEDI PNIVFIPI YFAVAGLNVDLTLNNEGRDWGYVFATIGIAIFTKI ISG  
 TLTAKLTGLFWREATAAGVLMSCKGIVEI VVLTVGLNAGI ISRKIFGMFVLMALVSTFVT  
 TPLTQLVYPDSYRDGVRKSLSTPAEDDGAADGLDSEGVDKTEINTQLNSLADVSKYRIGE  
 LTTVINTEAISPSLKLNLNLSLGVSPKPKNNKHNETSLSRMTTATDSTLKSNTFKIKK  
 MVHIWKSVDVDTNLSVIDEKLTPFEGVGALRAIHLRLLTERTDQLSSSLYNDDPHF  
 TANTDSLQIFDIFSNLSKIPFSSEVIFSTMREKAANIATMKMDSTDLILLPLKGASYEY  
 RGSVPFIDEKYANFDHIYSHLLGLNELSSTFFKSIFQSLKANFAVQISNTYGRNLNADRFK  
 RKRFNLLPKPYLTQSDYLGLYLLLLICYRDGYNNNDNASCIFINSKNIDFAKDLSTAF  
 EHDWLNRESTIKIVDIPFETKVPEEAIEKPSFIETVLDVGLSDTALADIEETTFI IGEDLP  
 DESEPFSEEVRTVIFEGSNRRFDTLIVHHFSE

SEQ ID NO: 46

YJL108C

>sp| P42946 |PRM10\_YEAST Pheromone-regulated membrane protein 10  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=PRM10  
 PE=1 SV=1

MIVSFGDATTRTSEVQLVRC TQGLNLWKLHQVHAVYKRVVHDTLGADEGNALLDQILADT  
 NLYPPWMCVLLYAFCSAMVTPYAFGGDWNLAISFFMGLCVGSLQFILSQSYMYSNVFE  
 ISASI WSFCGRAFGSIPRSHICFGAVTQGSLALILPGYIILCGALELQSRSLVAGAVRM  
 FYAIIYSLFLGFGITLGSALFGWMYHNATNEISCPQLI SPWFRFLFVPAFTIISILLNQA

HISQLPVMVFTSCTGYVVITYWAGKHFANSTEFATAALAAAFVIGVLGNLYSRIWKGLAVSAM  
LPAIFVQVPSGSIASQNSLLSGLQSANTI VNaNETITTSTSDPSSSMSFGMTMIQVCVGIS  
VGLFASSLFFVYPFGKKTGLFSL

SEQ ID NO: 47

YJL212C

>sp|P40897|OPT1\_\_YEAST Oligopeptide transporter 1 OS=Saccharoirc.yces  
cerevisiae (strain ATCC 204508 / S288c) GN=OPT1 PE=1 SV=1  
MSTIYRESDSLESESPPTPTTPIQINMEEKKDAFVKNIDEDVNNLTATTDEEDRDPES  
QKFDRHSIQEEGLVWKGDPYLPNSPYPEVRSVAVSIEDDPTIRLNHWRTWFLTTVFVVVF  
AGVNQFFSLRYPSEINFLVAQVVCYPIGRILALLPDWKCSKVPFFDLNPGPFTKKEHAV  
VTIA¥ALTSSSTAYAMYILNAQGSFYNMKLN¥GYQFLL¥WTSQMIGYGAAGLTRRW¥¥NPA  
SSIWPQTLIS¥SLFDSLHSRK¥EKT¥ANGWMPRYRFFLI¥LIGSFIWY¥PGFLFTGLS  
YFN¥ILWGSKTRHNFIAANTIFGTQSGLGALPITFDYTQ¥SQAMSGSVFATPFY¥SANTYA  
S¥LIFFFVI¥LPLCLYFTNTWYAKYMPVISGSTDYDNTQNKYNVTKILNEDYSINLEKYKEYS  
P¥FVFPFSYLLSYALNFAA¥IA¥F¥HCILYHGKDI¥AKFKDRKNGGTDIHMRI YSKNYKDC  
PDWWYLLLQI¥MIGLGF¥A¥CCFDTKFPAWAF¥IAILISLVNFIPQGILEAMTNQH¥GLN  
IITELICGYMLPLRPMANLLFKLYGFI¥MRQGLNLSRDLKCLAMYMK¥SPRLIFAY¥QIYAT  
IISGM¥N¥GVQEWMMHNIDGLCTTDQPNGFTCANGRTVFNASI IWSLPKYLFSGRI YNP  
LMWFFLI GLLFPLAVYAVQWKFPKFKFAKH IHTPVFFTGPNI P PST PYNYSLFFFAMSFC  
LNLIRKRWRWFNKNYFN¥MGAGVEAG¥AISV¥IIFLC¥QYPGGKLSWWGNNVWKR TYDND  
YKKFYTLKKGETFGYDKWW

SEQ ID NO: 48

YJR106W

>sp|P47144|ECM27\_YEAST Protein ECM27 OS=Saccharomyces cerevisiae  
(strain ATCC 204508 / 5288c) GN=ECM27 PE=1 SV=2  
MDWAINVAHPRLLYKDPKLS¥TFI¥PSLFHII IAF¥LLGICASDFLCPNVAHISDPNSLR  
SNGSL¥SKTASHASHTGALMA¥LLSWCNSPDLFSNLMSWATSTRETRSTS¥SLSIGE¥L  
GACGIILCI¥EGSI Fi IMSRTHIEI S QIQKLS IMRDLFSLAAMC¥MSY¥SLMNQVTVLN  
CLLMAFLYAFYL¥VKLTFKLNHSAETPDETAADTSLRENS¥SPFLDDSLMASGLLPPIQP  
GFDISNSITHGIKPSLLSAMDNSFLSMLNENSSLEEDSRNEMAELNLTLSMTPGQHWSA  
SATVAGEATSAGRPFSEPTNAFTEYRDSERAINSSPA¥FAPYRDNPDDEESQEQLLETT  
THGHFQAQEMRRFSKRS LGWI IKI Fi PHLNS¥FSQKSISDAI FSI IT¥PFFI IFKLSCPQP  
PSDILSYDPTLNRYSLTTLPI ILLFIQS ITAPFL LCS ILSVLLTYHLGYLVYLFPLILAM  
ALILLTAFITK¥NLHNKFTLSLDSSNILQEKLQKRKLLERLNTSIQIIIFLAIGIINII I  
WISLLANSLIEMMEIYQKILGLSKAILGLTIFAWGNSVGD LISNISMCRLYKTQTHYQDR  
¥RLATKFFMISCASCLGG¥MLNSMGGIGFSGL¥SMLFIGAFNDNEWWFLRKY¥KLQETSQ L  
DNTLNYKFI¥SC¥FI ILQI ILLLLFFGGPNNIKRRRLTKEMKLY¥GISMCGLWALATLINIL  
LELFS

SEQ ID NO :49

YJR160C

>sp|POCEO0|MPH3\_YEAST Alpha-giucosides permease MPH3  
OS=Saccharorayces cerevisiae (strain ATCC 204508 / S288c) GN=MPH3  
PE=1 SV=1  
MKNLSFLINRRKENTS DSN¥YPGKAKSHEPSWIEMDDQTKKDGLDI¥H¥EFSPDTRAPSD  
SNKVITEI FDATEDAKEADESERGMPLATALNTYPKAAAWSLL¥STTLIMEGYDTAILGA  
FYALPIFQRKFGSQNDKTGEWEISASWQIGLTL CYMAGEI¥GLQLTGPS¥DLVGNRYTLI  
IALFF' LAAFTFILIYFCNSL GMI AV' GQALCGMPWGCFQCLT¥S YASEICPLALRYYLTTYS

NLCWLFQGFLFAAGIMKNSQKKYADSELGYKLPFALQWILPVPLALGIFFAPESPWWLVKK
GRFDEARRSLRRTLSGKGPEKEILVTLEVDKIKVTIDKEKRLTSKEGSYSDCFEDKINRR
RTRITCLCWAGQATCGSILIGYSTYFYEKAGVSTEMSFTFSI IQYCLGICATFLSWWASK
YFGRYDLYAFGLAFQTIIVFFIIIGLGCSSSTHGSKMGSGLLMAVAFFYNLGIAPVVFCLV
SEMPSSRLRKTIIILARNTYNVVIS ICSVLILYQLNSKKWNWGAKSGFFWGLCFCTLIW
AVVDLPETAGKTFVEINELFKLGVSAKRFKSTKVDPFVVKTPPKDVSNDPKGDIEASIA
EE

SEQ ID NO: 50

YKL064W

>sp|P35724|MNR2\_YEAST Manganese resistance protein MNR2
OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=MNR2
PE=1 SV=1

MSTDNSQKDEGVPLLSPYSSSPQLRKKRNQKRRKDKFVGHKSDSRRPTQLLHDNLQHN
HGQITDFDQIDSWGMLHESDSTSNDI IKSEDP SLKGAFIDHRPSMSQPREGPQSVSSTVQ
PQPIMKFSTPSYKKPAGLRPSDQNRSLVSDLSPSELESWLKRRKSVHKS FVDENSPTDRR
QSNANNDVVIDVDALMNHVNNNASTGVNDNSKRRKKKRGSDSSSNKNSKSTSSDSNDEED
EYNSRPSSSLSSNSSLDDVCLVLDDEGSEVPKAWPDCTVLEEF SKEETERLRSQAIQDA
EAFHFQYDEDEEDGTSNEDGILFSKPIVTNIDVPELGNRRVNETENLNKNGRLRPKRIAPW
HLIQRPMVLGSNSTKDSKSRISQGLQDNLLVGRNIQYPPHI ISNNPEHFRFTYFRVDLDS
TVHSPTISGLLQPGQKFDL FVASIYSQDNSAGHIKTHPNSPTPGIKAETVSQ LQGLTAK
NPSTLSSMSVANIEDVPPFWLDVSNPTEEMKILSKAFGIHPLTTEDI FLGEVREKVELF
RDYYLICFRSFDIVA EKHVRRRRRKEKQESATLDHESISRRKSQAYGATMSNESNANNNNS
TSNASRSKWLPSILRARRRSSANRTTNTSSSYKRRVKSEKKKMEENEKFKRKSGDRHKP
REGELEPLNVYI IVFRTGVLTFHFAPT PHPINVRRRARLLKDYLNVTSDWIAYALIDDI
DAFAPMIELIEDEVYEIEDAILKMHQSDSSSDSDSDSDSGASDEDAFPFDVYSK KTS
YSSAKSSVSSRSMSTSEAS FNANLIGWKRKGDMLRRIGECRKR VMS ILRLLGSKADVIK
FAKRYNEQWEASPQSEIAMYLGD IQDHIVTMVSSLNHYEKL LSRSHSNYLAQINIDMTKV
NNDMNDVLGKITILGTI VLP MNVITGLWGMNVI VPGQYRDSL TWFIGIVLFMCM LACSAY
MYTKRRFGF

SEQ ID NO: 51

YKR050W

>sp|P28584|TRK2\_YEAST Low-affinity potassium transport protein
OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=TRK2
PE=1 SV=1

MPTAKRTSSRASLALPFQLRLVHKKSWGHR LRDFISGFLKSCRPIAKYVFPNFIVVHYIY
LITLSIIGSILLYPCKNTAFIDVLF LAAGASTQGGLATKSTNDFNLYQQIVVYVITLLST
PILIHGFLAFVRLYWFERYFDNIRDISKQNFKLRRTMTLQORELSGSSGNAARS SFKDN
LFRGKFVSREDPRQSASDVPMDSPTDTSALSSISPLNVSSSKEESSDTQSSPPNFSSKRQP
SDVDPRDI YKSIMMLQKQEKSNANSTDSFSSETNGPAFIVQERHERRAPHCSLKRHSVL
PSSQELNKLAQTKSFQKLLGLRRDEGDHDYFDGAPHKYMVTKKKKISRTQSCNIPTYTAS
PSPKTSQVVENHRNLAKSAPSSFVDEEMSFSPQESLNLQFQAHPKPKRREGDIGHPFT
RTMSTNYLSWQPTFGRNSVFIGLTKQKEELGGVEYRALRLLCCILMVYIYIGFNILAFVT
IVPWACTRHHYSEI IRRNGVSPTWWGFF TAMSAFNSNLGSLTADSMVSFD TAPYPLIFMM
FFI IIGNTGFPIMLRFI IWIMFKTSRDL SQFKESLGLLDHPRRCFTLLFP SGPWWLFT
TLVVLNATDWILFI ILDFNSAVVRQVAKGYRALMGLFQSVCTR TAGFNVDLSKLHPSIQ
VSYMLMMYVSVLPLAISIRRTNVEEQSLGLYDSGQDDENITHEDDIKETDHDGESEERD
TVSTKSKPKKQSPKSFVGAHLRRQLSFDLWYFLGLFI ICICEGRKIEDVNKPDFNVFAI
LFEVVSAYGTVGLSLGYPNTNTSLSAQFTVLSK LVI IAMLIRGNRGLPYTL DRAIMLPS
DKLEQIDRLQDMKAKGKLLAKVGEDPMTTYVKKRSHK LKKIATKFWGKH

SEQ ID NO: 52

YKR105C

>sp IP36172 VBA5\_YEAST Vacuolar basic amino acid transporter 5  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=VBA5  
 PE=3 SV=1

MEETKYSSQOEIEGACGSDASLNARGSNDSMPMGLSPLYLCLASLTLVLFITALDILIVGTI  
 IDVVAEQFGNYSKTGWLVTGYSLPNAILSLIWGRFASI IGFQHSLILAILIFEAGSLIAA  
 LASSMNLIFGRVAVGGSGLQTLCFVIGCTMVGERSRPLVISILSCAFAVAAIVGPII  
 GGAFTHVTRWCFYINLPIGGLAI IMFLLTYKAENKGILQQIKDAIGTISSFTFSKFRH  
 QVNFKRLMNGI IFKFDFFGFALCSAGLVFLGLTFGGNKYSWNSGQVITYLVLGVLLEFI  
 FSLVYDFFLFDKFNPEPDNISYRPLLLRRLVAKPAI IIVNMVTFLLCTGYNGQMI YSVQF  
 FQLI FASSAWKAGLHLIPIVITNVIAAASGVITKKLGLVKPLLIFGGVLGVIGAGLMTL  
 MTNTSTKSTQIGVLLLPFGFSLGFALQASLMSAQLQITKDRPEAAMDFIEVTAFTNFMKSL  
 GTTLGGVLSTTVFSASFHNKVSRAHLEPYEGKTVDDMILYRLQNYDGSHTIGNILSDSI  
 KNVFWMDLGFYALGFLLFCFSFNKKLI I PKKDDTPEDNLEDK

SEQ ID NO: 53

YKR106W

>sp IP36173 GEX2\_YEAST Glutathione exchanger 2 OS=Saccharomyces  
 cerevisiae (strain ATCC 204508 / S288c) GN=GEX2 PE=1 SV=1  
 MSSSVVGASSNKKSGIRQSCIEIERERHSNDDTYSMTSTFFKLENEIMSAQFDSLKYKI  
 LLISTAFVCGFGISLDYTLRSTYTGATNSYSEHSLSTVQVINAVSVGSQVVYSRLSD  
 HFGRLRLFLVATIFYIMGTIIQSQATRLTMYAAGSVFYNCYVGTNLLLTLILSDFSSLK  
 WRMFYQYASYWPYII IPWISGNI ITAANPQKNWSWNIAMWAFI YPLSTLPIIFLILYMKY  
 KSSKTAEWRSLEQARKERTGGFLFENLVFLFWKLDIVGILLITVSLGCILVPLTLANETS  
 QKWHNSKI IATLVSGGCLFFI FLYWEAKFAKSPLLPFKLLSDRGIWAPLGVTFNFNFTFF  
 ISCDYLYPVLVSMKESSTSAARIVNLPDFVAATASPFYSLLVAKTRKLLSVIGGCAAW  
 MVCMLGFYKYRGGSGSHEGVIAASVIMGLSGLLCSNSVIVILQAMTTHSRMAVITGIQYT  
 FSKLGAAIGASVSGAIWTQTMPNQLYKNLGNDTLAEIAYASPYTFISDYPWGSPERDAVV  
 ESYRYVQRIIMTVGLACTVPPFTFTMFMRNPELIDKATHEEFTEDGLVVLVLPDEENIFSQI  
 KALFRHNRSNKKSGC

SEQ ID NO: 54

YLR447C

>sp IP32366 VA0DJYEAST V-type proton ATPase subunit d  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=VMA6  
 PE=1 SV=2

MEGVYFNIDNGFIEGVVRYRNGLLSNNQYINLTQCDTLEDLKLQLSSTDYGNFLSSVSS  
 ESLTTSLIQEYASSKLYHEFNIRYDQSSGSTRKFMFYITYGYMIDNVALMITGTIHDRDK  
 GEILQRCHPLGWFDLPTLS VATDLESLEYETVLVDTPPLAPYFKNCFDTEELDDMNIEII  
 RNKLYKAYLEDFYNFVTEEIPPEPAKCEMQLLGFADRRSINIALNSLQSSDIDPDLKSD  
 LLPNIGKLYPLATFHQAQDFEGVRAALANVYEURGFLETGNLEDHFYQLEMELCRDAF  
 TQQFAISTVWAWMKSKEQEVNITWIAECIAQNQRERINNYISVY

SEQ ID NO: 55

YML116W

>sp IP13090 ATR1\_YEAST Aminotriazole resistance protein  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=ATR1  
 PE=1 SV=2

MGNQSLVVLTESKGEYENETELPVKKSSRDNNIGESLTATAFTQSEDEMVDNQNQKWNPN  
 YFKYAWQEYLFIFTCMISQLLNQAGTTQTLSIMNILSDSFGSEGNKSWLMASFPLVSGS  
 FILISGR LGDI YGLKKMLLVGYVLVLIWSLICGITKYSGSDTFFI ISRAFQGLGIAFVLP  
 NVLGI IGNIYVGGTFRKNIVISFVGAMAPIGATLGCLFAGLIGTEDPKQWPWAFYAYSIA  
 AFINFLVLSIYAIPSTIPTNIHHFSMDWIGSVLGVIGLILLNFVWNQAPISGWNQAYIIVI  
 LIISVIFLVVFI IYEIRFAKTPLLRAVIKDRHMIQIMLALFFGWGSFGIFTFYFQFQL  
 NIRQYTALWAGGTYFMFLIWGI IAALLVGFTIKNVSPSVFLFFSMVAFNVGSIMASVTPV  
 HETYFRTQLGTMIILSFGMDLSFPASSI IFSDNLPMEYQGMAGSLVNTVVNYSMSLCLGM  
 GATVETQVNS DGKHLKGYRGAQYLG I GLASLACMISGLYMVES FIKRRARAAAE YDCT  
 VA

SEQ ID NO: 56

YMR034C

>sp rQ05131 rYMS4\_\_YEAST Uncharacterized membrane protein YMR034C  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=YMR034C  
 PE=1 SV=1

MKTQYSLIRKIWAHSVTEFLKSQWFFICLAILIVIARFAPNFARDGGLIKQYSIGYGCV  
 AWIFLQSGLGKSRSLMANMLNWRHATILVLSFLITSSIVYGFCAVKAANDPKIDDWV  
 LIGLILTATCPTTVASNVIMTTNAGGNSLLCVCEVFIGNLLGAFITPALVQMFTNRAPFA  
 YGNPATGNIGALYGRVMKQVGLSVFVPLFVGVQVIQNCFPKGTAYYLGFLLKYYHIKIGSY  
 MLLLIMFSSFSTAFYQDAFTSVSHVCIIFLCFFNLGI YIFFTGLSYLCARPWFILKLFPH  
 EPIEGKSTRLYRYSYNIFRPFYYSKEDAICIMFCGPAKTAALGVSLITSQYGDKKEHLGK  
 LLVPLVLYQVEQVMTANFFVSLFKRWIQKDAQADGSESSCANENEEVDLEKI ISIGTGEN  
 QSVLSNNVPYTQPR

SEQ ID NO: 57

YMR056C

>sp lP04710 |ADT1\_\_YEAST ADP, ATP carrier protein 1 OS=Saccharomyces  
 cerevisiae (strain ATCC 204508 / S288c) GN=AAC1 PE=1 SV=1

MSHTETQTQSSHFGVDFLMGGVSAIAKTGAAPIERVKLLMQNQEEMLKQGLDTRYKGI  
 LDCFKRTATHEGIVSFWRGNTANVLRYPFTQALNFAFKDKIKSLLSYDRERDGYAKWFAG  
 NLFSGGAAGGLSLLFVYSLDYARTRLAADARGSKSTSQRQFNGLLDVYKKTLDGLLGL  
 YRGFVPSVLGI IVYRGLYFGLYDSFKPVLLTGALGFSFVASFLLGWVITMGASTASYPLD  
 TVRRRMMMTSGQTIKYDGDALDCLRKIVQKEGAYSFLKGCGANI FRGVAAAGVISLYDQLQ  
 LIMFGKFKF

SEQ ID NO: 58

YMR253C

>sp | Q04835 |YM87\_YEAST Oncharacterized membrane protein YMR253C  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=YMR253C  
 PE=1 SV=1

MNPSVPKVMKRENNTHLLVSKEMNDTSLQLPSTTRSLSPKESNSNEDFNVDGNETTLQRI  
 SKDYLPNIGLVLLTVSYFFNSAMVVSTKVLNENDPDDIANDRQIKPLQILLVRMVITYIG  
 TL IYMYINKST ISDVP FGKPE VRKWLVLVRGCTGFFGVFGMYYS LMYLT ISDAVL ITFLAP  
 SLTIFLSWVILRERFTKVEALGSLISLLGVVLIIVRPSFLFGTPELTDSSSQIVESSDPKS  
 RLIATLVGLWGVLMSCVYII IRYIGKRAHAIMSVSYFSLITAIVSFIGINTIPSMKFQI  
 PHSKKQWILFGNLGVSGFI FQLLLTMGIQRERAGRGLMITYTQLLYAVFWDVALYKHWP  
 IWSWIGMI I I ISATLWVIRIRAANNETTAKDLTPIIDDEENSIPLTEFDLSDSK

SEQ ID NO: 59

YNL065W

>sp|P53943|AQR1\_YEAST Probable transporter AQR1 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=AQR1 PE=1 SV=1  
 MSRSNSIYTEDIEMYPHNEQHLTREYTKPDGQTKSEKLNFEFAYINSHGTLTKTTTREI  
 EGDLDSETSSHSSDDKVDPTQQITAETKAPYTLSSYGQKWGMVAILTMCGFWSSSLGSPYIY  
 YPALRQLEKQFNVDENMVNVTVVVYLLFQGISPTVSGGLADCFGRRP IILAGMLIYVIAS  
 IGLACAPSYGVIIIFLRCIQSIGISPTIAISSGVVGDFTLKHHERGTFVGFVLLGQCF  
 GSLIGAVLTARWDWRAIFWFLTIGCGSCFLIAFLILPETKRTIAGNLSIKPKRFINRAPI  
 FLLGPPVRRRFKYDNPDYETLDPTIPKLDLSSAGKILVLP E I ILSLFP SGLLFAMWTLMLS  
 SISSGLSVAPYNYHLV I IGV CYLP GGIGGLMGSFFFTGRIIDMYFKRKIKKFEQDKANGLI  
 PQDAEINMFKVRLVCLLPQNFLAVVAYLLFGWSIDKGWRIESILITSFVCSYCAMSTLST  
 STTLVLVDL YPTKS STAS SCFN FVRCS LST I FMGC FAKMKAAMT VGGTFT FLCALVFFFN F  
 LMF I PMKYGMKWREDRLKQQRQSWLNTLAVKAKKGTKRQNDNHN

SEQ ID NO: 60

YNL070W

>sp|P53507|TOM7\_YEAST Mitochondrial import receptor subunit TOM7 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=TOM7 PE=1 SV=2  
 MSFLPSFILSDESKERISKILTLTHNVAHYGWIPFVLYLGWAHTSNRPNFLNLLSPLPSV

SEQ ID NO: 61

YNL083W

>sp|D6W196|CMC1\_\_YEAST Truncated non-functional calcium-binding mitochondrial carrier SAL1-1 OS=Saccharomyces cerevisiae (strain ATCC 204508 / 5288c) GN=SAL1 PE=1 SV=2  
 MLLKNCETDKQRDIRYACLFKELDVKGNGQVTLDNLISAFEKNDHPLKGNDEAIKMLFTA  
 MDVNKDSVVDLSDFKKYASNAESQIWNQFQRIDLDHDGKIGINEINRYLSDLDNQSICNN  
 ELNHELNSNEKVNKFSRFFEWAFPKRKANIALRGQASHKKNNTDNDRSKKT TDS DLYV TYDQ  
 WRDFLLLVPRKQGSRLHTAYSFYLFNEDVLSSEGDVTLINDFIRGFGFFIAGGISGVI  
 SRTCTAPFDRLLKVF LIARTDLSSILLNSKTDLLAKNPADINKISSPLAKAVKSLYRQGG  
 IKAFYVGNGLNVIKVFP ESSI KFGSFEVTKKIMTKLEGCRD TKDLSKFS TYIAGGLAGMA  
 AQFSVPIDTLKFRVQCAPLDTKLGNNLLFQTAKDMFREGGGQI ILQRCHSRYSGHISL  
 CCIRFGDFFLCKMVCYQGTGKDEPTTRSGHSGKQPGCTSNQCIQWNCRSFCCLSNQSFKN  
 KTTSPRNIC TSLCV

SEQ ID NO: 62

YNL095C

>sp|P53932|YNJ5\_YEAST Uncharacterized transporter YNL095C OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=YNL095C PE=1 SV=1  
 MVHITLQAIWVSVKPI IkiYLIIGVGFMAKMGILTVEATRI ISDIVLTVLLP SLSFNK  
 IVANIEDKDIKSVGIICLSALLIFGSGFFFAYVVRLLF LPVPKQWYGGILAGGMFPNISDL  
 PIAYLQSMDOGLVFSEEEGNKGVANV I IFLTMFLICIFNLGGFRLIESDFEYNDES AVR  
 VSETTKTQPAVSANTTNTDTSERFFSNEQQLFNNKYTARDSLTEAIGTKGENADVPPISR  
 RSTNSIAPLSLPDTSNSKITKPVQVKARNTI ACTQSEESQATRGSNPLDSQSSASTIHS  
 YNTSESYESSIDTMRARRTASQPRAYNTTTLL EENCLDEKCPKNMSMAALEP I R SIDMRA

LPSONIHHLIREYSNVDQYGHQRRNSSLRGADMNDVHSISSNSTLQTIKTANLTRILTSD
ATVSKKDIETSGESLPQWMRKFSLTPLLVFLLKNCLRPCSMAVIALTVAFI PWVKALFV
TTANTPHISQAPDNAPPLSFFMDFTGYVGAACVPFGLILLGATLGRKIGNLYPGFWKAA
VTLVILRQCVMPI FGVLWCDRLVKAGWVNWQDDRMLLFVIAISWNLPMTTTLIYFTASFT
PPETTAPIQMECVSFFLMLQYPLMVVSLPFLVSYFLKVQMNL

SEQ ID NO: 63

YNL121C

>sp|P07213|TOM70\_\_YEAST Mitochondrial import receptor subunit TOM70
OS=Saccharomyces cerevisiae (strain ATCC 204508 / 5288c) GN=TOM70
PE=1 SV=2

MKSFITRNKTAILATVAATGTAIGAYYYNQLQQQQQRGKNTINKDEKKDTKDSQKETE
GAKKSTAPSNPPIYPVSSNGEPDFSNKANFTAEKDKYALALKDKGNQFFRNKKYDDAIK
YYNWALELKEDPVFYSNLSACYVSVGDLKVVEMSTKALELKPDISKVLRRASANEGLG
KFADAMFDLSVLSLNGDFN DASIEPMLERNLNKQAMSKLKEKFGDIDTATATPTLSTQP
AKERKDKQENLPSVTSMAFFGI FKPELTFANYDESNEADKELMNGLSNLYKRSPESYDK
ADESFTKAARLFEEQLDKNNEDEKLKEKLAISLEHTGIFKFLKNDPLGAHEDIKKAIELF
PRVNSYI YMALIMADRNDSTEYNYFDKALKLDSNNSVYYHRGQMNFIQNYDQAGKDF
DKAKELDPENI FPYIQLACLAYRENKFDDCETLFSEAKRKFPEAPEVFNFFAEILTDKND
FDKALKQYDLAIELENKLDGI YVGIAPLVGKATLLTRNPTVENFIEATNLLEKASKLDPR
SEQAKIGLAQMKLQQEDIDEAITLFEESADLARTMEEKLQAITFAEAAKVQQRIRSDPVL
AKKIQUETLAKLREQGLM

SEQ ID NO: 64

YNL142W

>sp|P41948|MEP2\_YEAST Ammonium transporter MEP2 OS=Saccharomyces
cerevisiae (strain ATCC 204508 / S288c) GN=MEP2 PE=1 SV=1

MSYNFTGTPTGEGTGGNSLTTLNLTQFDLANMGWIGVASAGVWIMVPGIGLLYSGLSRKK
HALSLLWASMMASAVCIFQWFFWGYSLAFSHNTRGNFGITLEFFGFRNVLGAPSSVSSL
PDILFAVYQGM FAAVT GALML GGACE RARL FPMMVFL FLWMT IVYC P IACWVWNAEGWLV
KLGSLDYAGGLCVHLTSGHGGLVYALILGKRNDPVTRKGM PKYKPHSVTSVVLGTVFLWF
GWMFFNGGSAGNATIRAWYSIMSTNLAAACGGLTWMVIDYFRCGRKWTTVGLCSGIIAGL
VGITPAAGFVPIWSAVVIGVVTGAGCNLAVDLKSLLRIDDGLDCYSIHGVGCGIGS VLTG
IFAADYVNATAGSYISPIDGGWINHHYKQVGYQLAGICAALAWTVTVTSILLTMAIIPF
LKLRLSADEEELGTDAQIGEFITYEESTAYIPEPIRSKTS AQMPPPHENIDDKIVGNTDA
EKNSTPSDASSTKNTDHIV

SEQ ID NO: 65

YOL020W

>sp|P38967|TAT2\_\_YEAST Tryptophan permease OS=Saccharomyces
cerevisiae (strain ATCC 204508 / S288c) GN=TAT2 PE=1 SV=1

MTEDFISSVKRSNEELKERKSNFGFVEYKSKQLTSSSSHNSNSSHHDDDNQHGKRNIQR
CVDSFKSPLDGSFDTSNLKR TLKPRHLIMIAIGGSIGTGLFVSGKAIAEGGPLGVVIGW
AIAGSQI IGTIHGLGEITVRFPVVGAFANYGTRFLDPSISFVVSTIYVLQWFFVLPLEI I
AAAMTVQYWNSSIDPVIWVAIFYAVIVSINLFGVRGFGAEFAFSTIKAITVCGFIILCV
VLICGGPDHEFIGAKYWHPGCLANGFPVGLSVLVVASYSLGGIEMTCLASGETDPKGL
PSAIKQVFWRILFFFLISLTLVGFVLPYTNQNLGGSSVDNSPFVIAIKLHHIKALPSIV
NAVILISVLSVGNCSIFASSRTLCSMAHQGLIPWWFGYIDRAGRPLVGIMANSLFGLLAF
LVKSGSMSEVFNWLMIAIAGLATCIVWLSINLSHIRFRLAMKAQGKSLDELEFVSAVGIWG
SAYSALINCLILIAQFYCSLWPIGGWTS GKERAKIFFQNYLCALIMLFIFIVHKIYYKQ

TGKWWGVKALKDI DLETDRKDI D I E I V k Q E I A E k k M Y L D s R p W Y V R Q F H F W C

SEQ ID NO: 66

YOL075C

>sp|Q08234|YO075\_\_YEAST Uncharacterized ABC transporter **ATP-binding** protein/permease YOL075C OS=*Saccharomyces cerevisiae* (strain ATCC 204508 / S288c) GN=YOL075C PE=1 SV=3  
 MSQQENGDVATELIENRSLFSRIPRISLHVRDLSI VASKTNTTLVNTFSDMLPSGSMVAV  
 MGGSGSGKTTLLNLVASKISGGLTHNGSIRYVLEDTGSEPNETEPKRAHLDGQDHPIQKH  
 VIMAYLPQQDVLSPRLTCRETLKFAADLKLNSSERTKMLMVEQLIEELGLKDCADTLVGD  
 NSHRGLSGGEKRRLSIGTQMISNPSIMFLDEPTTGLDAYS AFLVIKTLKKLAKEDGRTFI  
 MSIHQPRSDILFLLDQVCILSKGNVYCDKMDNTIPYFESIGYHVPQLVNPADYFIDLSS  
 VDS RSDKE EAATQS RLN SLIDHWHDYERTHLQLQAE SYISNAT EI Q I Q N M T T R L P F W X Q V  
 TVLTRRNFKLNFSDYVTLISTFAEPLI IGTVCGWI YKPKDKSS IGGLRTTACLYASTIL  
 QCYLYLLFDYTRLCEQDIALYDRERAEGSVTPLAFIVARKISLFLSDDFAMTMIFVSITY  
 FMFGLEADARKFFYQFAVFLCQLSGLSMLSVAVSRDFSKASLVGNMFTVLSMGCGF  
 FVNAKVMPVYVRWIKYIAFTWYSFGTLMSSFTNSYCTTDNLDECLGNQILEVYGFPRNW  
 ITVPAVLLCWSVGYFVVGAI ILYLHKIDITLQNEVKSKQKKIKKKSPTGMKPEIQLLDD  
 VYHQKLEAEKGKNIHITIKLEDIDL RVI FSAPFSNWKEGNFHETKEILQSVNAI FKPG  
 MINA IMGPSGSGKSSLNLISGR LKSSVFAKFDTS G S I M F N D I Q V S E L M F K N V C S Y V S Q D  
 DDHLLAALT VKETLKYAAALRLHHLTEAERMERTDNLIRSLGLKHCENNI IGNEFVKGIS  
 GGEKRRVTMGVQLLNDPILLLEPTSGLD SFTSATILEILEKLCREQGKTI IITIHQPR  
 SELFKRFGNVLLAKSGRTAFNGSPDEMIAYFTELGYNCPSFTNVADFFLDLISVNTQNE  
 QNEISSRARVEKILSAWKANMDNESLSPTPISEKQQYSQESFFTEYSEFVRK PANLVLAY  
 IVNVKRQFTTTRRSFDSLMARIAQIPGLGVIFALFFAPVKHNYTSSINRLGLAQESTALY  
 FVGMLGNLACYPTERDYFEEYNDNVYGIAPFFLAYMTLELPLSALASVLYAVFTVLACG  
 LPRTAGNFFATVYCSFIVTCCGEALGIMTNTFFERPGFVVNCISI ILSIGTQMSGLMSLG  
 MSRVLKGFNLYLNPVGYTSMI IINFAPGNLKLTCEDGGKNSDGTCEFANGHDVLSYGLV  
 RNTQKYLGI IVCVAIIYRLIAFFILKAKLEWIKW

SEQ ID NO: 67

YOL077W-A

>sp|P81451|ATP19\_YEAST ATP synthase subunit k, mitochondrial OS=*Saccharomyces cerevisiae* (strain ATCC 204508 / S288c) GN=ATP19 PE=1 SV=1  
 MGAAYHFMGKAIPPHQLAIGTLGLLGLLVVPPNPFKSAKPKTVDIKTDNKDEEKFIENYLK  
 KHSEKQDA

SEQ ID NO: 68

YOL122C

>sp|P38925|SMF1\_\_YEAST Manganese transporter SMF1 OS=*Saccharomyces cerevisiae* (strain ATCC 204508 / S288c) GN=SMF1 PE=1 SV=2  
 MVNVGPSHA AVDA SEARKRNISEEV FELRDKKDSTV VIEGEAPVRTFTSSSSNHERED  
 TYVSKRQVMRDIFAKYLKFIGPLMVSVAI DPGNYSTAVDAGASNQFSLLCI ILLSNFI  
 AIFLQCLCIKLGSVTGLDL SRACREYLPRWLNWTLYFFAECAVIATDIAEVI G T A I A L N I  
 LIKVPLPAGVAITVVDVFLIMFTYKPGASSIRFIRIFECFVAVLVVGVICFAIELAYIP  
 KSTSVKQVFRGFVPSAQMFDHNGIYTAISILGATVMPHSLFLGSALVQPRLLDYDVKHGN  
 YTVSEEQDKVKKSKSTEEIMEEKYFNYPNTNAAIKYCMKYSMVELSITLFTLALFVNCAI  
 LVVAGSTLYNSPEADGADLFTIHELLSRNLAPAAGTI FMLALLLSGQSAGVVCTMSGQIV  
 SEGHI NWKLQ PWQRRLATRC IS I P C L V I S I C I G R E A L S K A L N A S Q V V L S I V L P F L V A P L  
 IFFTCKKSIMKTEITVDHTEEDSHNHQNNDRSAGSVIEQDGSSGMEIENGKDVKIVYMA  
 NNW I I T V I A I I V W L F L S L L N V Y A I V Q L G M S H G D I S



SEQ ID NO: 69

YOR079C

>sp|Q12067|ATX2\_YEAST Metal homeostasis factor ATX2 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=ATX2 PE=1 SV=1  
 MKFLGVILLASFLLIATFLIGLIPLYIDKQKSSIVTNQEGADSISDFTTNADTQTINDD  
 VSSYRVKIAVLSQFGIGMLLGTSFMLVI PEGIKACVEHDGNVGVNLLIGFLGVYVLDRLV  
 TLWVSRKQTVYTHDAVKFQSWKDIINHPRQIWMNLIQNNVVFALFIHGLSDGIALGTTTN  
 NDSLLI VVLI A I V I H K I PAVLSLTSLMVSQRNLMKWEVICNVFLFASSTPIGYIVLSLLN  
 LSHSPTMDWISGNLLMSGGSLLYASFTA FVGGDSHDHDL SVEQEVVLP HDESVYVLIGV  
 CIPLVISYCISEE

SEQ ID NO: 70

YOR087W

>sp|Q12324|YVC1\_YEAST Calcium channel YVC1 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=YVC1 PE=1 SV=2  
 MVSANGDLHLPISNEQCMPEENNGSLGFEAPTPRQILRVTLNLKYLIDKVVPIVYDPNDIV  
 CDHSEILSPKVVKLAYEACGGNPKDKANKRKYQSVII FSLKVCWEYSILATMEVHNAKL  
 YETRNLASQQCKLLIEREETRDQLFLMQLLRRYVINENDEDEQEPNLELAT DMHCT  
 TVIGSSGFQRCLKWIWRGWIVQNGLDPTTFIKDDSLAEVSLISHFNPVRLKAPVYQNYLQ  
 MIFSFLFLGLYTLVVNGKDSERVQSFDDLLESIFYVFNTGFILDELTKLYYIGYAHLSFWN  
 LFNDTTYLIITFAMGFRAMSVTPLNAKYSSSEDWDKISYRVLSCAAPFVWSRLLLYLESQR  
 FIGIMLVILKHMMEKESIVFFLLFLIMIGFTQGFLGLDSADGKRITGPILGNLTITVLG  
 LGSFDVFEFAPPYAAILYGYFFIVSVILLNILIALYSTAYQKVIDNADDEYMALMSQK  
 TLRIRAPDEDVYVSPNLNIEVFMTPI FRILPPKRAKDLSYTVMTIVYSPFLLLI SVKET  
 REARRIKYNRMKRLNDDANEYDTPWDLTDGYLDDDDGLFSDNRNSGMRATQLKNSRSLKL  
 QRTAEQEDVHFVKVPPKWKYKNVKKCSPSFEQYDNDDEDDAGEDKDEVKELTKKVENLTAV  
 ITDLLEKLDIKDKKE

SEQ ID NO: 71

YOR092W

>sp|Q99252|ECM3\_YEAST Protein ECM3 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=ECM3 PE=1 SV=1  
 MTHITLGQA IWA SVRPIIKIYLIIGVGFGLCKMNILTVQATRSISDIVLTILLPCLSFNK  
 IVANIEDNDIKDVGI ICLTSVILFATGLGFIVRSVLPVPKRWRGGILAGGMFPNISDL  
 PIAYLQSMQDQGFIFTEAEGEKGVANVI IFLAMFLICVFNLGGFRLIENDFHYKGDDEEN  
 TLTNDDSAQQPTQPIEGNSSSSSNQDILKEPNESTVPSNSQASYISEKNKKEKTELSVPK  
 PTHTAPPAIDDRSSNSAVVSIDSITHSLRTNHVDAQSVSELNDPTYRTRSQPIAYTTES  
 RTSHVHNNRRNSITGSLRSIDMRELPAEGMSDLIREYSNVDQYGRRRKSSISSQGAPSVL  
 QADGTISPNTLRTSTLQRVKTSNLTRI ITSDATVSKKDIETSGSSLPKWLQKFPLTKFFV  
 FFLKNCLRPCSMAVILALI IAFI P W V K A L F V T T S N T P K I K Q A P D N A P A L T F I M D F T S Y V G  
 AASVPPFGLILLGATLGRKIGKLYPGFWKSAVVLVFLRQCIMPFGVLWCRLVKAGWLN  
 WENDKMLLFVTAITWNLPTMTTLIYFTASYTPEDETEPVQMECTSFFLMLQYPLMVVSLP  
 FLVSYFIKVMQKL

SEQ ID NO: 72

YOR130C

>sp|Q12375|ORT1\_YEAST Mitochondrial ornithine transporter 1 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=ORT1

PE=1 S#2

MEDSKKKGLIEGAILDI INGSIAGACGKVIIEFPDFTVKVRLQTQASNVFPTTWSCKIFTY  
 QNEGIARGFFQGIASPLVGACLENATLFVSYNQCSKFLEKHTNVSPGQILISGGVAGSC  
 ASLVLTPVELVKCKLQVANLQVASAKTKHTKVLPTIKAI ITERGLAGLWQGQSGTFIRES  
 FGGVAWFATYIEIVKSLKDRHSLDDPKRDESKIWELLISGGSAGLAFNASIFPADTVKSV  
 MQTEHISLTNAVKKIFGKFGKGFYRGLGITLFRVAVPANA AVFYIFETLSAL

SEQ ID NO: 73

YOR222W

>sp|Q99297|ODC2\_YEAST Mitochondrial 2-oxodicarboxylate carrier 2  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=ODC2  
 PE=1 SV=1

MSSDSNAKPLPFYQFISGAVAGISELTVMYPLDVVKTRFQLEVTTPATAAVGKQVERYN  
 GVIDCLKKIVKKEGFSRLYRGISSPMLMEAPKRATKFCNDQYQKIFKNLFNTNETTQKI  
 SIAAGASAGMTEAAVIVPFELIKIRMQDVKSSYLGPMDCCLKKTIKNEGIMGLYKGIESTM  
 WRNALWNGGYFGVIYQVRNSMPVAKTKGQKTRNDLIAGAIGGTVGTMLNTPFDVVKSRIO  
 SVDAVSSAVKKNWCLPSLLVI YREEGFRALYKGFVVKVCR LAPGGSMLLWFTGMMNFF  
 RDLKYGH

SEQ ID NO: 74

YOR291W

>sp|Q12697|YPK9\_YEAST Vacuolar cation-transporting ATPase YPK9  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=YPK9  
 PE=1 SV=1

MDIPSSNQIQHGQRSENRMRPRASFSSSTATTTSTAATLTSAMVLDQNNSEPYAGATFEAV  
 PSSIVSFHHPHSFQSSNLPSPHSSGNLEQRGRRLTESEPLVLSAEQSRSSSRNPSHFRF  
 FTQEQISNAEGASTLENTDYDMAWDATPAYEQDRI YGTGLSSRRSS IRSFSRASSLSNAK  
 SYGSFSKRGRSGSRAPQRLGENSDTGFVYHSATHSSSSLSRYTTREIRPIELESQTDEIL  
 EDESSTHSLESSDSRRSASENNRGSFSGHDDVHNQHSEYLKPDYHEKFYPQYAPNLHYQR  
 FYIAEEDLVIGIAAYQTSKFWYI IYNLCCFLTFLGLVYLLTRWLPKLVKLYGVKVLAKA  
 EWVVIENEFGEFVIQPI DRQWYNRPLSTVLPFENYPNPSYEPNDINLSHHHANEINPNVP  
 ILITFEYRYIKFI YSPLDDLFKTNNNWDIPDWVDLSTVSNGLTKGVQEDRELAFGKNQIN  
 LRMKTTSEILFNEVLHPFYVFQVFSI ILWGIDEYYYAAACI FLISVLSIFDSLNEQKKVS  
 RNLAEMSHFHCDVRVLRDKFWTTISSSELVPGDI YEVSDPNITILPCDSILLSSDCIVNE  
 SMLTGESVPVSKFPATEETMYQLCDDFQSTQISSFVSKSFLYNGTNI IRARIAPGQTAAL  
 AMVVRTGFSTTKGSLVRSVMVFPKPTGFKFYRDSFKYIGFMSLIAIFGFCVSCVQFIKLGL  
 DKKTMLRALDIITI VVPPALPATLTIGTNFALSRLKEKGIFCISPTRLNISGKIDVMCF  
 DKTGTLTEDGLDVLGVQISEPNGVRGQKFGELLSDIRQVFPKFSLNDCSSPLDFKSRNFF  
 MSLLTCHSLRSVDGNLLGDPLDFKMFQFTGWSFEEDFQKRAFHSLYEGRHEDDVFPENSE  
 IIPAVVHPDSNNRENTFTDNDPHNFLGVVRSFEFLSELRRMSVIVKTNNDVYWSFTKGA  
 PEVISEICNKSTLPADFEVLRRCYTHNGYRVACAGKTLPKRTWLYSQKVSREEVESNLE  
 FLGFI IFQNKLKKETSETLKSLODANIRTIMCTGDNILTAI SVGREAGLIQCSRYYVPS I  
 NDTPLHGEPVIVWRDVNEPDKILDTKTLKPVKLGNNSSVESLRECNITLAVSGDVFRLLFR  
 DENEIPEEYLNEILLNSSIYARMSPEKHELMIQKLDYTVGFCGDCGANDCGALKAADV  
 GISLSEAEASVAAPFTSKI FNISCVLDVIREGRAALVTSFACFQYMSLYSAIQFITITIL  
 YSRGNSLNGDFQFLYIDLILLIVPIAICMSWSKSYEKIDKKRPSANLVSPKILVPLLISVFL  
 VFLFQFIPWI IVQKMSWYIKPIVGGDDAVQSSDNTVLFVSNFYILTAVLSVGGPPYRE  
 PMSKNFEFIVDITVSIGASLLLMTLDTESYLGKMLQLTPISNSFTMFI IVWVILNYAQL  
 YIPPSIKGWLKKKKSSKKYKLLIQEEMKLKEV

SEQ ID NO: 75

YOR306C

>sp|Q08777|MCH5\_YEAST Riboflavin transporter MCH5 OS=Saccharomyces cerevisiae {strain ATCC 204508 / S288c) GN=MCH5 PE=1 SV=2  
 MSSDSLTPKDTIVPEEQTNQLRQPDLEDESIHYDPEADDLESLETTASYASTSVSAKVYT  
 KKEVNKGTDIESQPHWGENTSSSTHSDSKEEDSNEEIEISFPEGGFKAWVVTFCFLGLIAC  
 FGLLNSTGVIESHLQDNQLSSESVSTIGWLFSLFLFVCSASCI ISGTYFDRNGFRTIMIV  
 GTVHFVAGLFATANSTKYWHFILSFAIVCGFGNGIVLSPLVSVPAHYFFKRRGTALAMAT  
 IGGSVGGVVFPIMLRSFFSMKSDTDPTYGFWGIRTLGFLDLALLTLS IILVKERLPHVI  
 ENSKDGESRWRYILRVYILQCFDAKAFKLDKYLFCVLGTVFSELSINSALTYYGSYATSH  
 GISANDAYTLIMI INVCGIPGRWVPGYLSDKFGRFNVAIATLLTLFIVMFVWGLPFGTNL  
 TNMYVISALYGFCSGSVFSLLPVCCGQISKTEEFKGRYSTMYFVVGFGTLVGIPITGAI I  
 SIKTTADYQHYI IFCGLATFVSAVCYI ISRAYCVGFKWVRF

SEQ ID NO: 76

YOR316C

>sp|P32798|COT1\_YEAST Cobalt **uptake** protein COT1 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=COT1 PE=1 SV=2  
 MKLGSKQVKI ISLLLLDTVFFGIEITTYGLSHSLALIADSFHMLNDIISLVVALWAVNVA  
 KNRNPDSTYTYGWKRAEILGALINAVFLIALCVSILIEALQRIIAPPVIENPKFVLYVGV  
 AGLISNTVGLFLFHDNDQEHGHGHGHSHGGIFADHEMHMPSSHTHTHAHVDGIENITPMD  
 STDNISEIMPNAIVDSFMNENTRLLTPENASKTPSYSTSSHTIASGGNYTEHNKRKRLN  
 MHGVFLHVLGDALGNIGVMLSAFFIWKTDYSWKYYTDPLVSLI ITGI IFSSALPLSCKAS  
 KILLQATPSTLSGDQVEGDLLKI PGI IAIHDFHIWNLTESIFIASLHIQLDISPEQFTDL  
 AKIVRSKLRHYGIHSATLQPEFITREVTSTERAGDSQGDHLQNDPLSLRPKTYGTGISGS  
 TCLIDDAANCNTADCLEDH

SEQ ID NO: 77

YOR334W

>sp|Q01926|MRS2\_YEAST Magnesium **transporter** MRS2, mitochondrial OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=MRS2 PE=1 SV=2  
 MNRRLLVRSISCFQPLSRITFGRPNTPFLRKYADTSTAANTNSTILRKQLLSLKPISASD  
 SLFISCTVFNSKGN I SMSEKFPKWSFLTEHSLFPRDLRKIDNSSIDI IPTIMCKPNCI V  
 INLLHIKALIERDKVYVFDTTNPSAAAKLSVLMYDLESKLSSTKNNSQFYEHRALESIFI  
 NVMSALETDFKLHSQICIQILNDLENEVNRLKLRHLLIKSKDLTLFYQKTLIRDLLDEL  
 LENDDDLANMYLTVKKS PKDNFSDLEMLIETYYTQCDEYVQQSESLIQDIKSTEEIVNI I  
 LDANRNSLMLELKVTIYTLGFTVASVLPAFYGMNLKNFIEESEWGFTSVAVFSIVSALY  
 ITKKNFNSLRSVTKMTMYPNSPANSSVYPKTSASIALTNKLRRRKWWKSTKQRLGVLLY  
 GSSYTNKANLSNNKINKGFSKVKKFNMENDIKNKQNRDMIWKWLIEDKKN

SEQ ID NO: 78

YPL078C

>sp|P05626|ATPF\_YEAST ATP synthase subunit 4, mitochondrial OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=ATP4 PE=1 SV=2  
 MSMSGVRGLALRSVSKTLFSQGVRCPSMVIGARYMSSTPEKQTDPKAKANSIINAIPGN  
 NILTKTGVLTGTSAAAVIYAI SNELYVINDESILLTLFLGFTGLVAKYLAPAYKDFADARM  
 KKVSDVLNASRNKHVEAVKDRIDSVSQQLQNVAAETTKVLFDVSKETVELESEAFELKQKVE  
 LAHEAKAVLDSWVRYEASLRQLEQRQLAKSVISRVQSELGNPKFQEKVLQQSISEIEQLL  
 SKLK

SEQ ID NO: 79

YPL270W

>sp|P33311 **MDL2\_YEAST** ATP-dependent permease MDL2 , mitochondrial  
OS=*Saccharomyces cerevisiae* (strain ATCC 204508 / 5288c) GN=MDL2  
PE=1 SV=3

MLNGRLPLLRGICRNMLSRPRLAKLPS IRFRSLVTPSSSqli PLSRLCLRSPAVGKSLI  
LQSFRCNSSKTVPETSLPSASPISKGSARSAHAKEQSKTDDYKDIIRLFMLAKRDWKL  
TAILLLTISCSIGMSIPKVIGIVLDTLKTSSGSDFFDLKIPIFSLPLYEFLSFFTVAL  
GCAANFGRFILLRILSERVVARLRANVIKKTTLHQDAEFFDNHKVGDLSRLGSDAYV  
SMTQKVSDGVKALICGVVGVMMCSLSPQLSILLFFTPPVLFASVFGKQIRNTSKDLQ  
EATGQLTRVAEEQLSGIKTVQSFVAEGNELSRYNVAIRDI FQVGKTAFTNAKFFTTTSL  
LGDLSFLTIVLAYGSYLVLSQSLSIGDLTAFMLYTEYTGNAVFGLSTFYSEIMQGAGAASR  
LFELTDRKPSISPTVGHKYPDRGVIEFKDVSFSYPTPRPSVQIFKNLNFKIAPGSSVCIV  
GPSGRGKSTIALLLLRYNPTTGTITIDNQDISKLNCKSLRRHIGIVQQEPVLMSTGIRD  
NITYGLTYTPTKKEIRSAKQCFCHNFITKFPNTYDVTIGPHGTLLSGGQKORIAIARAL  
IKKPTILILDEATSAL DVESEGA INYTFGQLMKS KSMT IVSIAHRLST IRRSENVILGH  
DGSVVEMGKFKELYANPTSALSQLLNEKAAPGPSDQQLQIEKVIKEDLNESKEHDDQKK  
DDNDDNDNNDNDSNNQSPETKDNNSDDIEKSVEHLLKDAAKEANPIKITPQP

SEQ ID NO: 80

YPL274W

>sp|Q08986 **SAM3\_YEAST** S-adenosylmethionine permease SAM3  
OS=*Saccharomyces cerevisiae* (strain ATCC 204508 / S288c) GN=SAM3  
PE=1 SV=1

MDILKRGNESDKFTKIETESTTIPNDSDRSGSLIRRMKDSFKQSNLHVIPEDLENSEQTE  
QEKIQWKLASQPYQKVLRSQRHLTMI AIGGTLGTGLFIGLGYSLASGPAALLIGFLLVGTS  
MFCVVQSAELSCQFPVSGSYATHVSRFIDESVGFVATNYALAWLISFPSELIGCALTI  
SYWNQTVNPAVWVAIFYVIMVNLNLFVGRGFAETEFALSI IKVIAI FIFIIIGIVLIAGG  
GPNSTGYIGAKYWHDPGAFAPVFKNLNCFVSAAFSFGGSELVLLTSTESKNISAI  
AKGTFWRIAIFYITTVV IIGCLVPYNDPRLLSGSNSDVSASPFVIALSNTGSMGAKVSN  
FMNVVILVAVVSVCSNCSVYASSRLIQALGASGQLPSVCSYMDRKGRLPLVGIGISGAFGLL  
GFLVASKKEDEVFTWLFALCSISSFFTWFICMSQIRFRMALKAQGRSNDEIAYKSILGV  
YGGILGCVLNALLIAGEIYVSAAPVGSPPSAAEAFFEYCLSIPIIMIVVYFAHRFYRRDWKH  
FYIKRSEIDLDTGCSVENLELFKAQKEAEQLIASKPFYKIRYRWC

SEQ ID NO: 81

YPR003C

>sp|P53394 **SULX\_YEAST** Putative sulfate transporter YPR003C  
OS=*Saccharomyces cerevisiae* (strain ATCC 204508 / 5288c) GN=YPR003C  
PE=1 SV=1

MTSNNSLLGRGRMSYSSTAPPRFKRSVDQRDTFSDNFDYDKDSSNRGRITYIAASNSTTG  
PPPNNSRSGCTNNTNNTNNTSNTSNTNNDSDVENTVFETLPYYLPCFSWLPEYTFNKLW  
GDVIAGISVASFQIPLALSYYTTSIAHVPLCGLYSLAISPFVYGILGSVPQMIVGPESAI  
SLVVGQAVESITLHKENVSLIDISTVITFVSGTILLFSGISRFGFLGNVLSKALLRGFIS  
SVGLVMI INSLISELKLKDLVSLPQHYHTPFKILFLIDYAPAQYHIPTAIFSGCCLIV  
LFLTRLLKRKLMKYHKSIAIFFPDILLVVIVTILISMKFNKHYGISI IGDFSMDNDFDEL  
KNPLTRPRRKLIPDLFSASLIVAMLGFFESTTASKSLGTTYNLTVSSNRELVALGFMNIV  
ISLFGALPAFGGYGRSKINALSGAQSVMGVMGVITLITMNLQFVHYIPNCVLSVIT  
TIIGISLLEEVPGDIKFHLRCGGFSELFVFAVTFCTTIFYSIEAGICIGCVYSI INI IKH  
SAKSRIQILARVAGTSNFTNLDYMMNMKRNSLDVEGTEEIEGCMIVRIPEPLTFTNSD  
LKQRDLRIERYGSSKIHPGRKSLRSKDSIKYVIFDLGGMTSIDSSAAQVLEEI ITS  
YKRRNVFI YLVNVSINDKVRRLFKAGVAASVERAQAANNENNTSNTFS DAGETYS  
PYFDSIDAALYEIEKMKIKGNNVNPNDSEFSMSNTLFNSSLV

SEQ ID NO: 82

YPR011C

>sp|Q12251|YP011\_\_YEAST Uncharacterized mitochondrial carrier YPR011C  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=YPR011C  
 PE=1 SV=1  
 MAEVLTVLEQPNSIKDFLKQDSNIAFLAGGVAGAVSRTVVSPFERVKILLQVQSSTTSYN  
 RGIFSSIRQVYHEEGTKGLFRGNGLNCIRIFPYSAVQFVVVEACKKKLFHVNGNNGQEQL  
 TNTQRLFSGALCGGCSVVATYPLDLIKTRLSIQTANLSSLNRSKAKSISKPPGIWQLLSE  
 TYRLEGGLRGLYRGVWPTSLGVVPYVALNFAVYEQREFGVNSSDAQPSWKSPLYKLTIG  
 AISGGVAQTITYPFLLRRRFQVLAMGGNELGFRYTSVWDALVTIGRAEGVSGYYKGLAA  
 NLFKVVVPSTAVSWLVYEVVCDVSRNW

SEQ ID NO: 83

YPR058W

>sp|P32331|YMC1\_YEAST Carrier protein YMC1, mitochondrial  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / 5288c) GN=YMC1  
 PE=1 SV=2  
 MSEEFPSQLIDDLEEHPQHDNARVVKDLLAGTAGGIAQVLVGQPFDTTKVRLQTSSTPT  
 TAMEVVRKLLANEGPRGFYKGTLTPLIGVGACVSLQFGVNEAMKRFFHHRNADMSSTLSL  
 PQYYACGVTGGIVNSFLASPIEHVIRLQTQTGSGTNAEFKGPLECIKKLRHNKALLRGL  
 TPTILREGHCGTYFLVYEALIANQMNRRLERKRDIPAWKLCI FGALSGTALWLMVYPL  
 DVIKSVMTDNLQPKPFGNSISSVAKTLYANGGIGAFFKGFPTMLRAAPANGATFATFE  
 LAMRLLG

SEQ ID NO: 84

YPR128C

>sp|Q06497|ANT1\_YEAST Peroxisomal adenine nucleotide transporter 1  
 OS=5accharomyces cerevisiae (strain ATCC 204508 / S288c) GN=ANT1  
 PE=1 SV=1  
 MLTLESALTGAVASAMANI AVYPLDLSKTI IQSQVSPSSSEDSNEGKVLPNRRYKNVVD  
 M INIFKEKGILGLYQGMTVTT VATFVQNFVYFFWYTFIRKSYMKHKLLGLQSLKNRDGPI  
 TPSTIEELVLGVAASISQLFTSPMAVVATRQQTVHSAESAKFTNVIKDIYRENGDITA  
 FWKGLRTGLALTINPSITYASFQRLKEVFFHDHSNDAGSLSAVQNFILGVLSKMISTLVT  
 QPLIVAKAMLQSAQSKFTTFQEALLYLYKNEGLKSLWKGVLPQLTKGVIVQGLLFAFRGE  
 LTKSLKRLI FLYSSFFLKHNGQRKLAST

SEQ ID NO: 85

YPR201W

>sp|Q06598|ARR3\_YEAST Arsenical-resistance protein 3  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=ARR3  
 PE=1 SV=1  
 MSEDQKSENSVPSKVMVNRTDILTTIKSLSWLDLMLPFTI ILSII IAVI ISVYVPSR  
 TFDAEGHPNLMGVSIPLTVGMIVMMIPPICKVSWESIHKYFYRSYIRKQLALSFLNWVI  
 GPLLMTALAWMALFDYKEYRQGIIMIGVARIAMVLIWNQIAGGDNDLCVVLVITNSLLQ  
 MVLYAPLQIFCYVISHDHLNTRVLFEEVAKSVGVFLGIPLGIGIIIRLGSITIAGKS  
 NYEKYILRFISPWAMIGFHYTLFVI FISRGYQFIHEIGSAILCFVPLVLYFFIAWFLTFA  
 LMRYLSI SRSDTQRECSCDQELLLKRVWGRKSCEASFSITMTQCFTMASNNFELSLAIAI  
 SLYGNNSKQAI AATFGPLLEVPILLILAIVARILKPYIWNRRN

SEQ ID NO: 86

YBR008C

>sp|P38124|FLR1\_YEAST Fluconazole resistance protein 1  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=FLR1  
 PE=1 SV=1

MVYTSTYRHTIVVDLLEYLGIVSNLETLQSAREDETRKPENTDKKECKPDYDIECGPNRS  
 CSESSTDSDDSSGSQIEKNDPFRVDWNGPSDPENPQNWPLLKSLVVFQIMLLTCVTYMG  
 SIYTPGQEIYQEEFHVGHVATLNLVSLYVGLGPI IFSPLSETARYGRLNLYMVTLLFF  
 FMIFQVGCATVHNIGGLIVMRFIGILCSPSLATGGGTVADI ISPEMVPLVLMWSAGAV  
 AAPVLAPLLGAAMVDAKNWRFIFWLLMWLSAATFILLAFFFPETQHHNILYRRALKRKE  
 TGDDRYYTEQDKLDREVDARTFLINTLYRPLKMI IKEPAILAFDLYIAVAYGCFYLFEEA  
 FPIVFGVIYHFSLVEVGLAYMGFCVGCVLAYGLFGILNMRIIVPRFRNGTFTPEAFLIVA  
 MCVCWCLPLSLFLFGWTARVHWILPVISEVFFVLAVFNIFQATFAYLATCYPKYVASVFA  
 GNGFCRASFACAFPLFGRAMYDNLATKNYPVAWGSSLVGFLTLGLAIPFILYKYGPSLR  
 TRSSYTEE

SEQ ID NO: 87

YBR021W

>sp|P05316|FUR4\_YEAST Uracil permease OS=Saccharomyces cerevisiae  
 (strain ATCC 204508 / S288c) GN=FUR4 PE=3 SV=2

MPDNLHLHLSGSSKRLNSRQLMESSNETFAPNNVDLEKEYKSSQSNITTEVYEASSFE  
 VSSEKPOYSSFWKKI YYEYVWDKSIILGVSILDSFMYNQDLKPVEKERRVWSWYNYCYFW  
 LAECFNINTWQIAATGLQLGLNWWQCWITIIWIGYGFVGAFFVLASRVGSAYHLSFP  
 ASFGIFFSLWPVINRVVMAIVWYSVQAYIAATPVSLMLKSIFGKDLQDKI PDHFGSPNAT  
 TYEFMCFIFWAASLPFLVPPHKIRHLFTVKAVLVPFASFGFLIWAIRRAHGRIALGSL  
 TDVQPHGSAFSAWFLRSLMGCMANFSTMVINAPDFSRFSKNPNSALWSQLVCIPFLFSIT  
 CLIGILVTAAGYEIYGINYWSPLDVLEKFLQTTYNKGTRAGVFLISFVFAVAQLGTNISA  
 NSLSCGTDMSAIFPKFINIKRGSFLCAAMALCICPWNLMATSSKFTMALSAYAIFLSSIA  
 GVVCSDFVVRGRIKLTTHIYSHQKGSFYMYGNRFGINWRALAAAYLCGVAPCLPGFIAEV  
 GAPAIKVSDBGAMKLYLSYVWVGYGLSFSSYALCYFFPVPVPGCPVNNI IKDKGWFQRWANV  
 DDFEEEWKDTIERDDLVDNISVYEHEHEKTFI

SEQ ID NO: 88

YBR043C

>sp|P38227|QDR3\_YEAST Quinidine resistance protein 3  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=QDR3  
 PE=1 SV=2

MQAQSQSNVGLSRNSCDNSLPNNHVMHCDSESSGSPHSEHNDYSYEKTNLESTASNSR  
 EHRDNQLSRLKSEEVVVPKNQRRGLLPQLAIIPEFKDARDYPPMMKKMIVFLIAFSSMMG  
 PMGTSI IFPA INSITTEFKTSVIMVNVS IGVYLLSLGVFPLWSSSLSELEGRRTTYITSF  
 ALLFAFNIGSALAPDINSFIALRMLCGAASASVQSVGAGTVADLYISEDGKLNLSYYLGL  
 PLLAPLLSPIFGSLLVNRWPWRSTQWFMVILSGCNVILLTVLLPETLRKQDSKGAIAQIL  
 AERRIQVDNNERGEIQEDYQRGEDETDRIVENQVATLSTEKHNYVGEVRDQDSDLLESHSS  
 PNTYDGRAGETQLQRI YTEASRSLYEYQLDDSGIDATTAQVTRIRSTDPKLARSIRENSL  
 RKLQTNLEEQVKKVLSNNGGEIAPKQVSAVRKVDWDTFFVYFIKPLKSLHFLEYPPVALAI  
 TFSAISFSTVYFVNMTVEYKYSRPPYNFKPLYIGLLYIPNSVTYFFASI YGGRVWDMLLK  
 RYKEYGILAPEARISWNVVTSVISFPIALLIIFGWCLDKKCHWVTPLIGTALFGYAAMMT  
 IGATLSYLVDLSPGKATGVALNLLIRQILAATAVAVVTTPLMLNGMGTGWAFTMLAFIVLG  
 ASSVLIILKKHGDYWRENYDLQKLYDKID

SEQ ID NO: 89

YBR287W

>sp|P38355|YB8B\_YEAST Uncharacteri<sup>zed</sup> transporter YBR287W  
 OS=Saccharomyces cerevisiae {strain ATCC 204508 / S288c) GN=YBR287W  
 PE=1 SV=1  
 MVETFSFAHLAYLVFESVLQVVIIALAGFWSASSGLLPKQSQKI ISLLNVDLFTPCLIFS  
 KLAKSLSMKIFEIAI IPIFFGLTTGISFISGKIMSRILDLDKDETNFVVANSVFGNSNS  
 LPVSLTSLAYTLPNLTWDQIPNDNRDNVARGILYLLI FQQIGQMLRWSWGYNKLKMKWS  
 GENTQHMPPSQVQSLLETPNIDNEELVNEEQEEQELLEEEENRMNSSFSSSSIGDKIW  
 QKCTVFERIRANLNPPLYSMIFAVVVAAGPLQRELFMEDGFINNTFAEAVTQLGSVSI  
 PLILVVLGSNLPSAEVFPKTVHHSKLLIGSI IGRMILPSCFLLPIIAIAVKYINVSILD  
 DPIFLVVGFLTLVSPPAIQLTQITQLNEFFEAEMADILFWGYAVLSLPVSI IVVSGA IYV  
 LQWANPT

SEQ ID NO: 90

YBR295W

>sp|P38360|ATU1\_YEAST P-type cation-transporting ATPase  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=PCA1  
 PE=1 SV=2  
 MKPEKLFSGLGTSDGEYGVVNSENISIDAMQDNRGECHRRSIEMHANDNLGLVSQRDCTN  
 RPKITPQECLSETEQICHHGENRTKAGLDVDDAETGGDHTNESRVDECCA EKVNDTETGL  
 DVDSCCGDAQTGGDHTNESCVDGCCVRDSSVMVEEVTGSCEAVSSKEQLLTSFEVVPKSKS  
 EGLQSIHDIRETTRCNTNSNQHTGKGRLCIESSDSTLKKRCKVSRQKIEVSSKPECCNI  
 SCVERIASRSCEKRTFKGSTNVGISGSSSTDSLSEKFFSEQYSRMYNRYSSILKNLGCIC  
 NYLRTLKGESCCLPKVRFCGEGASKKTKYSYRNSSGCLTKKKTHGDKERLSNDNGHADF  
 VCSKSCCTKMKDCAVTSTISGHSSSEISRIVSMEPIENHLNLEAGSTGTEHIVLSVSGMS  
 CTGCESKLLKKSFGALKCVHGLKTSLILSQAEFNLDLAQGSVKDVIKHLKTTTEFKYEQIS  
 NHGSTIDVVVYPYAAKDFINEEWPQGVTELKIVERNIIRI YFDPKVI GARDLVNEGWSVPV  
 SIAPFSCHPTIEVGRKHLVRVGCTTALSIIILTIPILVMAWAPQLREKISTISASMLATI  
 IQFVIAGPFYLNALKSLI FSRLIEMDLLIVLSTSAAYIFSIVSFGYFVVG RPLSTEQFFE  
 TSSLLVTLIMVGRFVSELARHRAVKSISVRSLSQASSAILVDKTGKETEINIRLLQYGDIF  
 KVL PDSRIPTDGTVISGSSEVDEALITGESMPVPKKCQSI VVAGSVNGTGTFLVFKLSKLP  
 GNNTISTIATMVDEAKLTKPKIQNIADKIASYFVPTIIGITVVTFVWIAV GIRVEKQSR  
 SDAVIQAIYAITVLIVSCPCVIGLAVPIV FVIASGVAAKRGVIFKSAESIEVAHNTSHV  
 VFDKTGTLTEGKLTVVHETVRGDRHNSQSLLLGLTEGIKHPVSMIAIASYLKEKGVSAQNV  
 SNTKAVTGRKVEGTSYGLKLGQGNCRWLGHNNDPDVRKALEQGYSVFVCFVNGSVTAVY  
 ALED SLRADAVSTINLLRQGISLHILSGDDGAVRSMAARLGISSNIRSHATPAEKSE  
 YIKDIVEGRNCDSSSQSRPVVFCGDTNDAIGLTQATIGVHINEGSEVAKLAADVML  
 KPCLNNILTMITVSQKAMFRVKNLFLWSFTYNLFAILLAAGAFVDFHI PPEYAGLGELVS  
 ILPVIFVAILLRYAKI

SEQ ID NO: 91

YBR296C

>sp|P38361|PH089\_YEAST Phosphate permease PH089 OS=Saccharomyces  
 cerevisiae (strain ATCC 204508 / S288c) GN=PH089 PE=1 SV=1  
 MALHQFDYIFAIAMLFAFLDAFNIGANDVANSFASSISSRSLKYWQAMVLAGLCEFLGAV  
 LAGARVSGTIKNNI IDSSIFTNDPAVLMLTMTSALIGSSCWLT FATAIGMPVSTTHSIVG  
 GTIGAGIAAGGANGVWGWGVSQI IASWFIAPILAGAIAAIVFSISRFSVLEVKSLERS  
 IKNALLLVGVLV FAFS ILTMLIVWKGPNLHLDDLSETETAVSIVLTGAIASIVYFIF  
 YPFYRRKVLQDWTCLKLIDIFRGPSFYFKSTDDIPPMP EGHQLTIDYYEGRRNLGTTVSV  
 EDEENKAASNSNDSVKNKEDIQEVDLVRTETETEPETKLS TKQYWWSSLLKQGPKKWPLLFWL

VISHGWTQDVIHAQVNDRDMLSGDLKGMYSKFDNRVEYIYSVLQAITAATMSFAHGA  
 NDVANATGPLSAVYVIWKTNTI GAKSEVPVVVLA YGGVALVIGCWTYGYN I IKNLGNKMI  
 LQSPSRGFSIELAVAITTVMATQLGIPTSTTQIAVGGIVAVGLCNKDLKSVNWRMVAWCY  
 SGWFLTLPIAGLIAGI INGI ILNAPRFGVEYQMT

SEQ ID NO: 92

YCL038C

>sp|P25568|ATG22\_YEAST Autophagy-related protein 22 OS=Saccharomyces  
 cerevisiae (strain ATCC 204508 / 5288c) GN=ATG22 PE=1 SV=1  
 MSYGTINDMNESVTNYRIKKAQNNIKGWYAYSFSSEPFVVS AVSTYIPLLLQQFASINGV  
 KVHDHSIPCLSETGSDSKCVLGLFNNRI FVDTSSFALYVFSLSVLFQTI IVISVSGIVD  
 LWGSVKFKGRILVWFGVIGALSTVAISKLNDDTQI YSLAGLYIVANGCFGVINVVGNLLP  
 I FVKDSLKCQSQGAYEPDKVDSLTTVISGRGASLGYSSALIVQIVSMFLVASKKGSKQDV  
 QVAVLFGIWWFVWQLPMIWLIDDVTIPIRVDDSTLASARSPYPGEQDALGQLNWKNYLS  
 YGWSLFEFSFKHARLLKDVMIFLIAWFIISDSITTTINSTAVLFSKAELHMSTLNLIMISV  
 LTVVNAMLGAFMI PQFLATKFRWTSSQTLMYI I IWASFIPFYGILGFFFNAFGLKHKFEM  
 FLLAIWYGLSLGGLSAVRSVFSLVPPGKESTFFSMFSITDKGSSILGPFVLVGLLTDKT  
 HNI RYSFYFFFLLMLSLPVLNCLDVKRGREAEELSQVLPESERRLD

SEQ ID NO: 93

YCR011C

>sp|P25371|ADP1\_YEAST Probable ATP-dependent permease  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=ADP1  
 PE=1 SV=2  
 MGSHRRYLYYSILSFLLLSCSVVLAKQDKTPFFEGTSSKNSRLTAQDKGNDTCPFCFCM  
 LPIFECKQFSECNSYTGRCECIEGFAGDDCSLPLCGGLSPDESGNKDRPIRAQNDTCHCD  
 NGWGGINCDVCQEDFVCDAFMPDPSIKGTCTYKNGMIVDKVFSGCNVTNEKILQILNGKIP  
 QITFACDKPNQECNFQFVIDQLESFYCGLSDCAFEDLEQNTSHYKCNDVQCKCVPDPTVL  
 CGAKGSIDISDFLTETIKGPGDFSCDLETRQCKFSEPSMNDLILTVFGDPYITLKCESGE  
 CVHYSEIPGYKSPSKDPTVSWQGLVLAALTAVMVLALFTFATFYISKSPFRNGLGSSKS  
 PIRLPDEDAVNNFLQNEDDLATLSFENITYSVPSINSDGVEETVLNEISGIVKPGQILA  
 IMGGSGAGKTTLLDILAMKRKTGHVSGSIKVNIGISMDRKSFSKIIGFVDQDDFLLPTLTV  
 FETVLNSALLRPLKALSFEAKKARVYKVLEELRIIDIKDRIIGNEFDRGISGGEKRRVSI  
 ACELVTSPLVLFLEPTSGLDASNANNVIECLVRLSSDYNRTLVLVLSIHQPRSNIFYLFDK  
 LVLLSKGEMVYSGNAKKVSEFLRNEGYICPDNYNIADYLIDITFEAGPQGKRRRIRNISR  
 LEAGTDTNDIDNTIHQTTFTSSDGTQREWAHLAAHRDEIRSLRDEEDVEGTDGRRGAT  
 EIDLNTKLLHDKYKDSVYYAELSQEIEEVLSEGDEESNVLNGDLPTGQQSAGFLQQLSIL  
 NSRSFKNMYRNPKLLGLNYLLTILLSLFLGTLTYNVSNDISGFQNRMGFLFFILTYFGFV  
 TFGTGLSSFALERIIFIKERSNNYYSPLAYYISKIMSEVVPLRVVPPILLSLIVYPMTGLN  
 MKDNAFFKCGIGILILFNLGISLEILTIGIIFEDLNNSI ILSVLVLLGSLFLSGLFINTKN  
 ITNVAFKYLKNFSVFYAYESLLINEVKTLMLKERKYGLNIEVPGATILSTFGFVVQNLV  
 FDIKILALFNVVFLIMGYLALKWIVVEQK

SEQ ID NO: 94

YDL054C

>sp|Q07376|MCH1\_YEAST Probable transporter MCH1 OS=Saccharomyces  
 cerevisiae (strain ATCC 204508 / S288c) GN=MCH1 PE=1 SV=1  
 MPLSKVEHYLSYHTRLLLLPHVLSLQSSHRVAYIFSLLSAVSTGFITLISLYSQPWQKHLN  
 YSSWQINTIASMTNLGMYLTPPILGMIADSHGPITLSLLAIIGFI PSYSYLAYVFNHPEL  
 SLGGNGDSSFNLSIICFVFIGISTSAFYFALLTCTKLYPHTKLLSISLPTTCYGISSVV  
 GSQLLRKWFSSNASSSSNSDLNLRVVFQTFALVYVVIGLLAWIATSVVSLLFHNEEQ



DNQKRLDDQTDVEQS PLLERSNHVQEKFTQTMLRI FSDPVTYI LAVS ILLSLGPLEMFI A  
 NMGSLTNLLVQLDAPTLSTKLLSTYALSSTFTRLLTGIVADFFAKKKISIKWILLTFLSL  
 GVCAQLFLKMTSSASPWGLVPTGSLVGIIVYGGLFTVYPTLVLLVWGERSFRTVYGSLLI  
 APAIGSMI FCMLYAKFYDSRCMSGGDLRNPSCISAVYKYSSIAFVVS AVLSAVVFWKLLK  
 SRKLR I

SEQ ID NO: 95

YDL100C

>sp|Q12154|**GET3\_YEAST** ATPase GET3 OS=Saccharomyces cerevisiae  
 (strain ATCC 204508 / S288c) GN=GET3 PE=1 SV=1  
 MDLTVEPNLHSLITSTTHKWIFVGGKGGVGGKTTSSCSIAIQMALSQPNKQFLLISTDPAH  
 NLSDAFGEKFGKDARKVTGMNNLSCMEIDPSAALKDMNDMAVSRANNNGSDGQGDLLGSL  
 LQGGALADLTGS IPGI DEALS FMEVMKHIKRQEQGEGETFDTVI FDTAPTGH TLRFLQLP  
 NTL SKLLEKFG EITNKLGPMLNSFMGAGNVDISGKLNELKANVETIRQQFTDPDLTTFVC  
 VCISEFLSLYETERLIQELISYDMDVNSIIVNQLLFAENDQEHNCRCQARWKMQKKYLD  
 QIDELYEDFHVVKMPLCAGEIRGLNNLTKFSQFLNKEYNPITDGKVIYELEDKE

SEQ ID NO: 96

YDL245C

>sp|P54854|**HXT15\_YEAST** Hexose transporter HXT15 OS=Saccharomyces  
 cerevisiae (strain ATCC 204508 / S288c) GN=HXT15 PE=1 SV=1  
 MASEQSSPEINADNLNSSAADVHVQPPGEKEWSDGFYDKEVINGNTPDAPKRGFLGYLI I  
 YLLCYPVSFGGFLPGWDSGITAGFINMDNFKMNFSGYKHSTGEYYLSNVRMGLLVAMFSV  
 GCSIGGVAFARLADTLGRRLAIVIVVLVYMGVGAIIQISSNHKWYQYFVGKI IYGLGAGGC  
 SVLCPMLLSEIAPTDLRGLVSLYQLNMTFGIFLGYCSVYGT RKYSNTAQWRIPVGLCFL  
 WALIIIVGMLLVPESPRYLIECERHEEACVSI AKINKVSPEDPWVLKQADEINAGVLAQR  
 ELGEASWKELFSVKT KVLQRLITGILVQTFQLTGENYFFFYGT TI FKS VGLTDGFETSI  
 VLGTVNFSTIIAVMVVDKIGRRKCLLFGAASMMACMVIFASIGVKCLYPHGQDGPSSKG  
 AGNAMIVFTCFYIFCFATWAPVAYIVVAESFP SKVSKAMSISTAFNWLWQFLIGFFTP  
 FITGSIHFYGYVFGCLVAMFLYVFFFLPETIGLSLEEIQLLYE EGIKPKWSASWVPPS  
 RRGASSRETEAKKKSWEV LKFPKSFN

SEQ ID NO: 97

YDL247W

>sp|P0CD99|**MPH2\_YEAST** Alpha-glucosides permease MPH2  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / 5288c) GN=MPH2  
 PE=2 SV=1  
 MKNLSFLINRRKENTSDSNVYPGKAKSHEPSWIE MDDQTKKDGLDIVHVEFSPDTRAPSD  
 SNKVITEIFDATEDAKEADESERGMPLATALNTYPKAAAWSLLVSTTLIMEGYDTAILGA  
 FYALPIFQRKFGSQNDKTGEWEISASWQIGLTL CYMAGEIVGLQLTGPSVDLVGNRYTLI  
 IALFFLAAFTFILYFCNSLGMIAVGQALCGMPWGC FQCLTVSYASEICPLALRYYLTTYS  
 NLCWLFQQLFAAGIMKNSQKKYADSELGYKLPFALQWILPVPLALGIFPAPESPWWLVKK  
 GRFDEARRSLRRTL SGKGPEKEILVTLEVDKIKVTIDKEKRLTSKEGSYSDCFEDKINRR  
 RTRITCLCWAGQATCGSILIGYSTYFYEKAGVSTEMSFTFSI IQYCLGICATFLSWWASK  
 YFGRYDLYAFGLAFQTIVFFI IGGLGCSSTHSGSKMGSGLLMAVAFFYNLGIAPVVFCLV  
 SEMPSRLRKT TI ILARNTYNNVSI ICSVLILYQLNSKKWNWGA KSGFFWGLCFCTLIW  
 AVVDLPETAGKTFVEINELFKLGVSARKFKSTKVD PFFVVKTP LKTSLITTPREISKLPLQ  
 RNSNVSHHL

SEQ ID NO: 98

YDR011W

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>sp IP32568 |SNQ2_YEAST Protein SNQ2 OS=Saccharomyces cerevisiae
(strain ATCC 204508 / S288c) GN=SNQ2 PE=1 SV=2
MSNIKSTQDSSHNAVARSASSFAASEESFTGITHDKDEQSDTPADKLTKMLTGPARDTA
SQISATVSEMAPDVVSKVESFADALSRHTTRSGAFNMDSDDGDFDAHAIFESFVRDADE
QG IHIRKAGVT IEDV'SAKG VDAS ALEGAT FGN ILC LPLT I FKG IKA KRHQKMRQ1I SNVN
ALAEAGEMILVLRPGAGCSSFLKVTAGEIDQFAGGVS GEVAYDGI PQEEMMKRYKADVI
YNGELDVHFPYLTVKQTLDFAIACKTPALRVNNVSKKEYIASRRDLYATIFGLRHTYNTK
VGND FVRGV SGG ERKRVS IA EA LAA KGS I YCWDNATRGLDASTALEYAKAIRIMTNLLKS
TAFVTIYQASENI YETFDKVTVLVYSGKQI YFGLIHEAKPYFAKMGYLCPPRQATAEFLTA
LTDPNGFHLIKPGYENKVPRTAEFEFETYWLN SFEFAQMKKDIAAYKEKVNTEKTKEVYDE
SMAQEKSKYTRKKSYYTVSYW EQVKLCTQRGFQRI YGNKSYTVINVC SAIIQSFITGSLF
YNTPSSTSGAFSRGGVLYFALLYSLMGLANISFEHRPILQKHGYSLYHPSAE AIGSTL
ASFPPRMIGLTCFFIILFFLSGLHRTAGSFFTIYLF LTMCEAINGLFEMVSSVCDTLSQ
ANSISGILMMSISMYSTYMIQLPSMHPWFKWISYVLP IRYAFESMLNAEFHGRHMDCANT
LVPSGGDYDNLSDDYKVC AFVGSKPGQSYVLGDDYLKNQFYVYKHTWRNFGILWCFLLG
YVVLKVI FTEYKRPVKGGDALIFKKGSKRFIAHADEESPDNVNDIDAKEQFSSESSGAN
DEVFDDLEAKGVFIWKDVCTTI PYEGGKRMLLDN VSGYCI PGTMTALMGESGAGKTTLLN
TLAQRNVGI ITGMDLVNGRP IDASFERRTGYVQQDIHIAELTVRESLQFSARMRRPQHL
PDSEKMDYVEKI IRVLGMEEYAEALVGEVGCGLNVEQRKKLS IGV'ELVAKPDLLLFLDEP
TSGLDSQSSWAI IQLLRKLKSKAGQSILCTIHQPSATLFEEDRLLLLLRKGGQTVYFGDIG
KN SATILN YFERNGARKC D SSENPAEYI LEA IGAGATASVKEDWKEKWLNSVEFEQTK EK
VQDLINDLSKQETKSEVGDKPSKYATSYAYQFRYVLRITSTSFWRSLNYIMSKMMLMLVG
GLYIGFTFFNVGKSYVGLQNAMFAAFISIILSAPAMNQIQGRAIASRELFEVRESQSNMF
HWSLVLITQYLSELPYHLFFSTIFFVSSYFPLRIFFEASRS AVYFLNYCIMFQLYYVGLG
LMILYMSPNLP SANVILGLCLSFMLSFCGVTQPVSLMPGFWTFMWKASPYTYFVQNLVGI
MLHKKPVVCKKKELNYFNPPNGSTCGEYMKPFLEKATGYIENPDATSDCAYCIYEVGDNY
LTH ISSKYSYLWRNFGI FWIYIFFN1IAMVCVYYLFHVRQ3 SFLSPVS ILNKIKNIRKKK
Q
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SEQ ID NO: 99

YDR2 92C

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>sp|P32916 |SRPR_YEAST Signal recognition particle receptor subunit
alpha homolog OS=Saccharomyces cerevisiae (strain ATCC 204508 /
S288c) GN=SRP101 PE=1 SV=2
MFDQLAVFTPQGQVLYQYNCLGKGFSEIQINSFISQLITSPVTRKESVANANTDGFDFNL
LTINSEHKNSPSFNALFYLNKQPELYFVVTFAEQTLELNQETQQTLALVLKLNLSLHLS E
SILKNRQGGNEKNKHN YVDILQGIEDDLK KFEQYFRIKYEESIKQDHINPDNFTKNGSVP
QSHNKNTKKKLRDTKGGKQSTGNVGSGRKWGRDGGMLDEMNHEDA AKLDFSSSN SHNSSQ
VALDSTINKDSFGDRTEGGDFLIKEIDDLSSHKDEITSGNEAKNSGYVSTAFGFLQKHV
LGNKTINESDLKSVLEKLTQQ LITKNVAPEAADYLTQQVSHDLVGSKTANWTSVENTARE
SLTKALTQILTPGVSDLLREIQSKRSKKDEEGKCDPVVFSIVGVNGVGKSTNLSKLAFW
LLQNNFKVLIVACDTFRSGAVEQLRVHVENLAQLMDDSHVRGSKNKRKGTGNDYVELFEA
GYGGS DLVTKIAKQAIKYSRDQNFDIVLMDTAGRRHNDPTLMSPLKSFADQAKPDKIIMV
GEALVGTDSVQQAKN FNDAFGKGRNL DFFIISKCDTVGEMLGTMVNMVYATGI P I L F V G V
GQTYTDLRRTL SVKWAVNTLMS
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SEQ ID NO: 100

YDR497C

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>sp IP30605 |ITR1_YEAST Myo-inositol transporter 1 OS=Saccharomyces
cerevisiae (strain ATCC 204508 / S288c) GN=ITR1 PE=1 SV=2
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MG IHI PYLTSKTSQSNVGDVGNADSVVEFNSEHDSPSKRGKITLESHEIQRAPASDDEDR  
 IQIKPVNDEDDTSVMITFNQSLSPFI ITLTFVASISGFMFGYDTGYISSALISIGTDLDH  
 KVLTYGEKEIVTAATSLGALITSIFAGTAADI FGRKRCLMGSNLMFVIGAI LQVSAHTFW  
 QMAVGRLLIMFGVGVIGSLIAPLFISEIAPKMIRGRLTVINSLWLTGGQLVAYGCGAGLNY  
 VNNGWRILVGLSLIPTAVQFTCLCFLPDTPRYVVMKGDRLARATEVLKRSYTDTSSEEIER  
 KVEELVTLNQSIPGKNVPEKVWNTIKELHTVPSNLRALIIGCGLQAIQQFTGWNSLMYFS  
 GTIFETVGFKNSSAVSIIIVSGTNFIFTLVAFFSIDKIGRRTILLIGLPGMTMALVVCIA  
 FHFLGIKFDGAVAVVSSGFSSWGIVIIIFI IVFAAFYALGIGTVPWQQSELFPQNVIRGI  
 GTSYATATNWAGSLVIASFTLMLQNITPAGTFAFFAGLSCLSTIFCYFCYPELSGLELE  
 EVQTIKDGFINI KASKALAKKRKQQVARVHELKYE PTQEIIEDI

SEQ ID NO: 101

YEL006W

>sp|P39953|YEA6\_\_YEAST Mitochondrial nicotinamide adenine  
 dinucleotide transporter 2 OS=Saccharomyces cerevisiae (strain ATCC  
 204508 / S288c) GN=YEA6 PE=1 SV=1  
 MNNGDNKTTLENSKNASLANGNYAIPTKLNRLKKNADPRVAAIS GALS GALS AMLVCPFD  
 VAKTRLQAQGLQNMTHQSQHYKGGFFGTATI FKDEGAAGLYKGLQPTVLGYI PTLMIYFS  
 VYDFCRKYSVDIFPHSPFLSNASSAITAGAISTVATNPIWVVKTRLMLQTGIGKYSTHYK  
 GTIDTFRKI IQQEGAKALYAGLVPALLGMLNVAIQFPLYENLKIRFGYSESTDVSTDVTS  
 SNFQKLILASMLSKMVASTVTYPHEILRTRMQLKSDLPNTVQRHLLPLIKITYRQEGFAG  
 FYSGFATNLVVRTVPAAVVTLVVSFEYSKYLTTFFQ

SEQ ID NO: 102

YEL027W

>sp|P25515|VATL1\_YEAST V-type proton ATPase subunit c  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=VMA3  
 PE=1 SV=1  
 MTELCPVYAPFFGAIGCASAIIFTSLGAAAGTAKSGVGICATCVLRPDLLFKNIVPVIMA  
 GI IAIYGLVVSVLVCYSLGQKQALYTGFIQLGAGLSVGLSGLAAGFAIGIVGDAGVRGSS  
 QPRLVFGMILILIFAELVGLYGLIVALLNSRATQDVVC

SEQ ID NO: 103

YEL065W

>sp|P39980|SIT1\_YEAST Siderophore iron transporter 1  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=SIT1  
 PE=3 SV=1  
 MDPGIANHTLPEEFEEVVPEMLEKEVGAQVDVKPTLTSSPAPSYIELIDPGVHNEIY  
 AEMYNRPI YRVALFFSLFLIAYAYGLDGNIRYTFQAYATSSYSQHSLSTVNCIKTVIAA  
 VGQIFFARLSDI FGRFSIMIVSI IFYSMGTI IESQAVNITRFAVGGCFYQLGLTGIIILIL  
 EVIASDFSNLNWRLALFIPALPFIINTWISGNVTS AIDANWKGIGMWAFILPLACIPL  
 GICMLHMYRLARKHAKDRLKPEFEALNKLKWKSFCDIAFWKLDIIGMLLITVFFGCVLV  
 PFTLAGGLKEEWKTAH IIVPEVIGWVVVLPPLYMLWE IKYSRHPLTPDWL IQDRGI FFALL  
 IAFFINFNWYMQGDYMYTVLVAVHESIKSATRITSLYSFVSVIVGTILGFILIKVRRTK  
 PFIIIFGISCWIVSFGLLVHYRGDSGAHSGI IGSLCLLGFAGSFTYVTQASIQASAKTHA  
 RMAVVTSLYLATYNIGSAFGSSVSGAVWTNLPKEISKRISDPTLAAQAYGSPFTFITTY  
 TWGTPERIALVMSYRYVQKILCIIGLVFCFPLLGCFAFMLRNHKLTD SIALEGNDHLESKN  
 TFEIEEKEESFLKNKFFTHFTSSKDRKD

SEQ ID NO: 104  
 YER019C-A  
 >sp|P52871|SC6B2\_YEAST Protein transport protein SBH2  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=SBH2  
 PE=1 SV=1  
 MAASVPPGGQRILQKRRQAQSIKEKQAKQTPPTSTRQAGYGGSSSSILKLYTDEANGFRVD  
 SLVVLFSLVGFIFSIALHLLTKFTHII

SEQ ID NO: 105  
 YER053C  
 >sp|P40035|PIC2\_YEAST Mitochondrial phosphate carrier protein 2  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=PIC2  
 PE=1 SV=1  
 MESNKQPRKIQLYTKEFYATCTLGGI IACGPTHSSITPLDLVKCRLQVNPPLYTSNLQGF  
 RKIIANEGWKKVYTGFGATFVGYSLQGAGKYGGYEFKHLVSSWLSPGVTVYLMASATAE  
 FLADIMLCPFEAIKVKQQTTPPFCCNNVVDGWKKMYAESGGMKAFYKGIPLWCRQIPYT  
 MCKFTSFEEKIVQKIYSVLPKKKEEMNALQQISVSFVGGYLAGILCAAVSHPADVMMVSKIN  
 SERKANESMSVASKRIYQKIGFTGLWNGLMVRIVMIGTLTSFQWLIYDSFKAYVGLPTTG

SEQ ID NO: 106  
 YER119C  
 >sp|P40074|AVT6\_YEAST Vacuolar amino acid **transporter** 6  
 OS=Saccharomyces **cerevisiae** (strain ATCC 204508 / S288c) GN=AVT6  
 PE=1 SV=1  
 MVASIRSGVLTLLHTACGAGILAMPYAFKPFGLIPGVIMIVLCGACAMQSLFIQARVAKY  
 VPQGRASFALTRLINPNLGIVFDLAIKCFVGVSYMIVVGDLMQPIMSVWTRNAWLL  
 NRVQISLIMLFFVAPLSFLKLNLSRYASMVAI SSVAYLCVLVLLHYVAPS DEILRLKG  
 RISYLLPPQSHDLNVLNLTLPFVVFAYTCHHNMFS IINEQRSSRFEHVMKIPLIAISLALI  
 LYIAIGCAGYLTFGDNI IGNI IMLYPQAVSSTIGRIAIVLLVMLAFPLQCHPARASIHQI  
 LQHFAEENVSISATSADPTVATESSPLIRDSSLDLNEVIEEESI YQPKETPLRGKSFIV  
 ITCSILVASYLVAISVSSLARVLAIVGATGSTSISFILPGLFGYKLGTEHKTAVPLTTK  
 IFKYTGLLLFIWGLIIMITCLTAALKLN

SEQ ID NO: 107  
 YFL028C  
 >sp|P43569|CAF16\_YEAST CCR4-associated factor 16 OS=Saccharomyces  
 cerevisiae (strain ATCC 204508 / S288c) GN=CAF16 PE=1 SV=1  
 MVSQFAIEVRNLYTKFKESSDPSVVDINLQIPWNTSLVVGANGAGKSTLLKLLSGKHLK  
 LDGKILVNGLDPFSPLSMNQVDDDESVEDSTNYQTTTYLGTWCHMSI INRDIGVLELLK  
 SIGFDHFRERGERLVRILDIDVRWRMHRLSDGQKRRVQLAMGLLKPWRVLLLDEVTVDLK  
 VIARARLLEFLKWEETTRCS VVYATHIFDGLAKWPNQVYHMKSGKI VDNLDYQKDVEFS  
 EVVNAKVNQVAFENDNNKVVISKVNSLHPLALEWLKRDNPDPKEIGI

SEQ ID NO: 108  
 YFR04 5W  
 >sp|P43617|YFL5\_YEAST Uncharacterized mitochondrial **carrier** YFR04 5W  
 OS=Saccharomyces **cerevisiae** (strain ATCC 204508 / S288c) GN=YFR045W  
 PE=1 SV=3  
 MANQNSDLYKQITAGSVAAVFQTTMTYPFEYLKTGLQLQPKGTAFEIILPQIKSYFVGCS  
 ALNVAAFVKITILRFVTFDKLCHSLNNDNNDNFQRLTGYNLLIAGTLTGIVESLFI IPF  
 ENIKTTLIQSAMIDHKKLEKNQPVVNAKATFHKVATKSTPVARIEKLLPAVKHMYQTRGP

AA FVQG7TATTFRQ IANTS IQFTAYTAFKRL LQARN DKAS SVITGLATSFTLVAMTQPID  
 VVKTRMMSQNAKTEYKNTLNCMYRI FVQEGMATFWKGSIFRFMKVGI SGGLTFTVYEQVS  
 LLLGFSSRS

SEQ ID NO: 109

YGL084C

>sp|P53154|GUP1\_YEAST Glycerol uptake protein 1 OS=Saccharomyces  
 cerevisiae {strain ATCC 204508 / 5288c) GN=GUP1 PE=1 SV=1  
 MSLISILSPLITSEGLDSRIKPSPKKDASTTTKPSLWKTTEFKFYIAFLVVVPLMFYAG  
 LQASSPENPNYARYERLLSQGWLFGRKVDNSDSQYRFFRDNFALLSVLMLVHTSIKRIVL  
 YSTNITKLRFDLIFGLIFLVAAHGVNSIRILAHMLLILYAIAHVLKNFRRIATISIWYGI  
 STLFINDNFRAYPFGNICSFLSPLDHWYRGI IPRWDVFFNFLLRVLSYNLDFLERWENL  
 QKKKSPSYESKEAKSAILLNERARLTAAHPIQDYSLMNYIAYVTYTPLFIA GPI ITFNDY  
 VYQSKHTLPSINFKFIYYAVRFVIALLSMEFILHFLHVVAISKTKAWENDTPFQISMIG  
 LFNLNI IWLKLLI PWRLFRLWALLDGI DTPENMIRCVDNNYSSLAFWRAWHRS YNKWVVR  
 YIYIPLGGSKNRVLTSLAVFSFVAIWHDIELKLLWGWLI VLFLLPEIFATQIFSHYTDA  
 VWYRHVCAVGAVFNIWMMIANLFGFCLGSDGTKLLSDMFCTVSGFKFVILASVSLFIA  
 VQIMFEIREEEKRHGI YLKC

SEQ ID NO: 110

YGL104C

>sp|P53142|VPS73\_YEAST Vacuolar protein sorting-associated protein  
 73 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=VPS73  
 PE=1 SV=1  
 MNRILSSASLLSNVSMRPNQKHKITKALCYAI IVASIGSIQFGYHLSELNAPQQVLSCE  
 FDIPMEGYPYDRTWLGKRGYKQCI PLNDEQIGIVTSVFCIGGILGSYFATSLANI YGRKF  
 SSLINCTLNIVGSLII FNSNSYRGLI IGRILVGISCGSLIVI IPLFIKEVAPSGWEGLL  
 SMTQICIRLGVLLTQGIAPLTDYSYRWRWILFGSFLI AVLNFMMWFIVDESPKWLLAHGR  
 VTDAKLSLCKLRGVTFDEAAQEIQDWQLQIESGDPLIEPTTTNSISGSNSLWKYLDRDTN  
 VKSRHVITVLLFGQQFCGINSIVLYGTKI ISQLYPQHAIRINFFISMVNVLVITILVSLLI  
 HSLPRKPLLMSTVLSVTA FIMGIAMNHKNMNLIVFSFI YMGVFTMGLNPLPFIIMRE  
 VSKPQDMVLAQRYGTICNVWGTFI IAYTFPI IHDVLSGYVFI IFAI IACSISAFIWKVVP  
 ETKRSG

SEQ ID NO: 111

YGL114W

>sp|P53134|YGL4\_YEAST Putative oligopeptide transporter YGL114W  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=YGL114W  
 PE=1 SV=1  
 MPQSTPSQEVQRPWDNKPALKQITLRATIAGIAIGSLVLTSNFQFGLQGTGWVSMMSLPS  
 ALLACAFFKNIWPLIFPNDRPFSDVENVYVQSMVAVGTGPLAFGFVGVIPAIKFLTND  
 ESGGLREQQSFTEFRELLIWSTALAFFGIFFAVPLRKQVIVREKLPFPSPGSATATLISVL  
 NGTEILQEVSKESELLEMQRRLNECPEVLQPNRDPEEADYLMNSSHSELGDYTATSQDGS  
 SILSTGSENYRANI IILLKTFVVSPLYTMVSYFVPIRSIPVFGKYLSNNYLWNFQPSPA  
 YIGQGI IMGLPTVSYMLIGCFLGWGVLAPLARYKRWVPPDADVHDWEEGVQGWILWSSLS  
 IMVADSVVAFI VVTVKSIVKFI LI DDKAALLNNIIDDTFQSMLEEEERAINSSRRNTYVD  
 GRQDTVRLVSRDNEIEVDSKHLVRYTTVISGCLVSSIICIVSII YLFGIQVIPLYAI ITA  
 LILALFLSILGIRALGETDLNPNVSGIGKISQLIFAFI IPRDRPGSVLMNVVSGGIAEASA  
 QQAGDLMQDLKTGHLLGAS PRAQFCAQLI GACWS 1LSSFMYLC YNKVYS IPSEQFRI PT  
 AVVWIDCARLVTKGLPDKALECSMILGVIFAVLSLIRNTYRDYGYGWILYIPSGVAVGV  
 GIFNSPSTIARFIGGWASHFWLKNHRGDLNAKTKMIVFSSGLVLGEGIFSVINMLFICL  
 NVPHY

SEQ ID NO: 112

YGL167C

>sp|P13586|ATC1\_YEAST Calcium-transporting ATPase 1 OS=Saccharomyces cerevisiae (strain ATCC 204508 / 5288c) GN=PMR1 PE=1 SV=1  
MSDNPFNASLLDEDSNREREIILDATAEALSKPSPSLEYCTLSVDEALEKLDTDKNGGLRS  
SNEANNRRSLYGPNIEITVEDDESFLFKKFLSNFIEDRMILLIGSAVVSLFMGNI DDAVSI  
TLAIFI VVTVGFVQEYRSEKSLEALNKLVPAECHLMRCGQESHVLASTLVPGDLVHFRIG  
DRIPADIRI IEAIDLSDIDESNLGTGENEPVHKTSQTIEKSSFNDQPNSIVPISERSCIAYM  
GTLVKEGHGKGI VVGTTNTSFGAVFEMMNIEKPKTPLQLTMDKLGKDSLVSFIVIGM  
ICLVGI IQGRSWLEMFQISVSLAVAAI PEGLP IIVTVTLALGVLRMAKRKAI VRRLP SVE  
TLGSVNVICSDKTGTLTSNHMTVSKLWCLDSMSNKLNVLSLDKNKTKNSNGNLKNYLTE  
DVRETLTIGNLNCNNASFQEHAI FLGNPTDVALLEQLANFEMPDIRNTVQKVQELPFNSK  
RKLMTAKILNVPDNKCTVYVKGAFERILEYSTSYLKS KKKTEKLTEAQKATINECANSM  
ASEGLRVFGFAKLTLSDSSTPLTEDLIKDLTFTGLIGMNDPPRPNVKFAIEQLLQGGVHI  
IMITGSENTAVNIAKQIGIPVI DPKLSVLSGDKLDEM SDDQLANVI DHVN IFARATPEH  
KLNIVRALRKRGDVAMTGDGVN DAPALKLSDIGVSMGRIGTDVAKEASDMVLTDDDFST  
ILTAIEEGKGI FNNIQNFLT FQLST SVAALSLVALSTAFKLPNPLNAMQILWINILMDGP  
PAQSLGVEPVDHEVMKKPPRKR TDKILTHDVMKRLLTAAACI IVGTVYIFVKEMAEDGKV  
TARDTTMTFTCFVFFDMFNALACRHNTKSIFEIGFFTNKMFNYAVGLSLLGQMCAIYIPF  
FQSIFKTEKLGISDILLLLLISSSVFIVDEL RKLWTRKKNEEDSTYFSNV

SEQ ID NO: 113

YGR257C

>sp|P53320|MTM1\_YEAST Mitochondrial carrier protein MTM1 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=MTM1 PE=1 SV=1  
MSDRNTSNSLTLKERMLSAGAGSVLTSILITPMDVVRI RLQQQQMI PDCSCDGAAEVPNA  
VSSGSKMKTFTNVGGQNLNNAKIFWESACFQELHCKNSSLKFNGTLEAFTKIASVEGITS  
LWRGISLTLMLAIPANMVYFSGYEYIRDVSP IASTYPTLNPLFCGAIARVFAATS IAPLE  
LVKTKLQSI PRSSKSTKTWMMVKDLLNETRQEMKMVGPSRALFKGLEITLWRDVPFSAI Y  
WSSYELCKERLWLDSTRFASKDANWVHFINSFASGCISGMIAAICTHPFDVGKTRWQISM  
MNNSDPKGGNRSRNMFKFLETIWRTEGLAALYTGLAARVIKIRPSCAIMISSYEISKKV F  
GNKLHQ

SEQ ID NO: 114

YHL035C

>sp|P38735|VMR1\_YEAST ABC transporter ATP-binding protein/permease VMR1 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=VMR1 PE=2 SV=1  
MGTDPLI IRNNGSFWEVDDFTRLGR TQLLSYYLPLAIIASIGI FALCRSGLSRYVRS AEC  
DLVNEYLFGAQEERKEDNSIERLLRNSNTQANYVNVKKQGRILKLRHFDIT TIDVKQIDA  
KNHGGLTFSRPSTSDHLRKSSEIVLMSLQI IGLSFLRVTKINIELTNRDVTTLL LFWLIL  
LSLSILRVYKRSTNLWAICFTAHTTIWISTWI PIRSVYIGNIDDVPSQI FYI FEFVITST  
LQPIKLTSPIKDNSSII YVRDHTSPSREHIS SILSCITWSWITNFIWEAQNTIKL KDI  
WGLSMEDYSIFILKGFTRRNKHINNLTLALFESFKTYLLIGMLWVLVNSIVNLLPTILMK  
RFLEIVDNPNRSSSCMNLAWLYI IGMFICRLTLAICNSQGFVSDKICLRIRAILIGE I Y  
AKGLRRRLFTSPKTS SDDSISANLGTI INLISIDSFKVSELANYLYVTVQAVIMI I VVV  
GLLNFNLGVSAFAGISII LVMFPLNFL LANLLGKFQKQTLKCTDQRISKLNECLQNIRI V  
KYFAWERNIINEIKSIRQKELRSL LKSLVWSVTSFLWFVTPTLVTGVTF AICTFVQHED  
LNAPLAFTTSLSLFTLLKTPLDQLSNM LSFINQSKVSLKRISDFLRMD DTEKYNQLTISPD  
KNKIEFKNATLTWNENDSDMNAFKLCGLNIK FQIGKLNILGSTGSGKSALLLGLLGELN

LISGSIIVPSLEPKHDLIPDCEGLTNSFAYCSQSAWLLNDTVKNNI I FDNFYNEDRYNKV  
 IDACGLKRDLEILPAGDLTEIGEKGITLSGGQKQRISLARAVYSSAKHVLLDDCLSAVDS  
 HTAVWIYENCITGPLMKNRTCILVTHNVSLTLRNAHFIVLENGKVKNQGTITELQSKGL  
 FKEKYVQLSSRDSINEKNANRLKAPRKNDKQIEPV TENINFDANFVNDGQLIEEEEKSN  
 GAISPDVYKWLKFFGGFKALTALFALYITAQILFISQSWWIRHWVNDTNVRINAPGFAM  
 DTLPLKGMTDSSKNKHNAFYLLTVYFLIGI IQAMLGGFKTMFTLSGMRASRKI FNNLLD  
 LVLHAQIRFFDVTVPVGRIMNRFKSDIEGVDQELI PYLEVTIFCLIQCASI IFLITVITPR  
 FLTVAVIVFVLYFFVVGKWLTA SRELKRLDSITKSPIFQHFSETLVGVCTIRAFGDERRF  
 ILENMNKIDQNNRAFFYLSVTVKWFSFRVDMIGAFIVLASGSFILLNIANIDSGLAGISL  
 TYAILFTDGALWLVRLYSTFEMNMNSVERLKEYSSIEQENYLGHDEGRILLNNEPSWPKD  
 GEIEIENLSLRYAPNLPVIRNVSFKVDPQSKIGIVGRTGAGKSTI ITALFRLLEPITGC  
 IKIDGQDISKIDLVTLLRSITI IPQDPILFAGTIKSNVDPYDEYDEKKI FKALSQVNLIS  
 SHEFEEVLNSEERFNSTHNKFLNLHTEIAEGGLNLSQGERQLLFIARSLREPKI ILLDE  
 ATSSIDYDSHLIQGI IRSEFNKSTILTIAHRLRSVIDYDRIIVMDAGEVKEYDRPSELL  
 KDERGIFYSMCRDSSGGLLELLKQIAKQSSKMMK

SEQ ID NO: 115  
 YHL036W

>sp|P38734|MUP3\_\_YEAST Low-affinity methionine permease  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=MUP3  
 PE=1 SV=1  
 MEPLLFNSGKANPSQDVFIDVEVGDITTKYGSTNTGSFSSMDTVEAQAIKAETARFMEVP  
 QGRHLGVFSTVVLVFSRIMGSGIFAVPSVILLNTGGNKLIYFAIWVFSAAIAFAGLYLFL  
 EFGSWIPKSGGRKNFLERSFERPRLIS VVFSCYSVLTGYALTGSIVFGKYVLSAFGVTD  
 DSWSKYVSISFIIFAVLIHGVSVRHGVFIQNALGGLKLIMIVLMCFAGLYTLFFYKSTGQ  
 VAWDLPVTQVEKDSLLSVSSIATAFISSFFCFSGWDTVHTVTSEIKNPVTKLVSGPLSL  
 IICFVCYTMNVAYLKVLTYYEIVSAGPLVGSVLF'TKLFGPRVGGKFIASFIAISAASNI  
 LVVI YSISRNVQEIFKEGYLPFSIHMSKNWPFDAPLPSISLCGFITIAWILILPKEGESF  
 NYLVSMGDYGNQFFLLVAIGLFIWRFKHKNEVPEIRASTFGVLAI ITLSLYMLMAPFFA  
 DPSLNRVGFPPYQIMSLLVIVACFFFWLVKVVLLPKFFHYKLLPKITYLHDGLI VTEWV  
 KKPCLC

SEQ ID NO: 116  
 YHR002W

>sp|P38702|LEU5\_\_YEAST Mitochondrial carrier protein LEU5  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=LEU5  
 PE=3 SV=1  
 MTRDSPDSNDSYKHINKNTTQKTSFDRNSFDYIVRSLAGGISGCAKTLIAPLDRIKIL  
 FQTSNPHYTKYTGSLIGLVEAAKHIWINDGVRGFFQGH SATLLRIFPYAAVKFVAYEQIR  
 NTLIPSKEFESHWRRLVSGSLAGLCSVFITYPLDLVRVRLAYETEHKRVKLGRI IKKI YK  
 EPASATLIKNDYIPNWFCHWCNFRYRGYVPTVLGMIPYAGVSFFAHDLLHDVLKSPFFAPY  
 SVLELSEDELERVQKKQRRPLRTWAELISGGLAGMASQTAAYPFEI IRRRLQVSALSPK  
 TMYDHKFQS ISEIAH 11FKERGVGFFVGLSI GYIKVT'PMVACS FFVYERMKWNFGI

SEQ ID NO: 117  
 YHR096C

>sp|IP38695|HXT5\_\_YEAST Probable glucose transporter HXT5  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=HXT5  
 PE=1 SV=1  
 MSELNAHQGPLEGSATVSTNSNSYNEKSGNSTAPGTAGYNDNLAQAKPVSSYISHEGPP

KDELEELQKEVDKQLEKKSXSDLLFVSVCCLMVAFGGFVFGWDTGTISGFVRQTDFFIRRF  
 GSTRANGTTYLSDVVRTGLMVSIFNIGCAIGGIVLSKLGDMYGRKIGLMTVVVIYSIGIII  
 QIASIDKWYQYFIGRIISGLGVGGITVLAPMLISEVSPKQLRGTLVSCYQLMITFGIFLG  
 YCTNFGTKNYSNSVQWRVPLGLCFAWSIFMIVGMTFVPEsprylVEVGKIEEAKRSLARA  
 NKTTEdSPLVTLEMENYQSS IEAERLAGSASWGEIvTGKpQmFRRTLMGMMIQSLQQLTG  
 DNYFFYYGTTIFQAVGLEDSFETAIVLGVVNFVSTFFSLYTVDRFGRRNCLLWGCVGMIC  
 CYVVYAS VGVTRLWPNQDQPSSKAGAGNCMIVFACFYIFCFATTWAPVAYVLI SESYPLR  
 VRGKAMSIAACNWIWGLISFFTPFITSAINFYYGYVFMGCMVFAYFYVFFFVFPETKGL  
 TLEEVENEMYEENVLPWKSTKWIPPSRRRTDYDLDATRNDPRPFYKRMFTKEK

SEQ ID NO: 118  
 YIL006W

>sp|P40556|YIA6\_\_YEAST Mitochondrial nicotinamide adenine  
 dinucleotide transporter 1 OS=Saccharomyces cerevisiae (strain ATCC  
 204508 / S288c) GN=YIA6 PE=1 SV=1  
 MTQTDNPVPCNGLLPEQQYCSADHEEPLLLHHEEQLIIFPDHSSQLSSADI IEPKMNSSSTE  
 SIIGTTLRKKWVPLSSTQITALSGAFAGFLSGVAVCPDLVAKTRLQAQGLQTRFENPYR  
 GIMGTLSSTIVRDEGPRGLYKGLVPIVLGYFPTWMI YFSVYEFsKkFFHGIFPQDFVAQS  
 CAAITAGAASTTLTNPIWVVKTRLMLQSNLGEHPHthyKGTfDAFRKLFYQEGFKALYAGL  
 VPSLLGLFHVAIHFIYEDLKVRFHCYSRENNTNSINLQRLIMASSVSKMIASAVTYPHE  
 ILRTRMQLKS DI PDS IQRRLFPLI KATYAQEGlKGFYSgFTTNLVRT IPASAITLVS FEY  
 FRNRLENISTMVI

SEQ ID NO: 119  
 YIL120W

>sp|P40475|QDR1\_\_YEAST Quinidine resistance protein 1  
 OS=Saccharomyces cerevisiae {strain ATCC 204508 / S288c) GN=QDR1  
 PE=1 SV=1  
 MTKQQTsvMRNASIAKEEREGSDNNNVDRSSSDAISDNDAERSNSHSEIDNESNfDMVPY  
 SRFSHKQKMLLVVQCAFTGFFSTVAGSIYYPVLTIIERKFNITEELANVTI VVYFIFQGV  
 APS IMGGLADTFGRRPI VLWAILAYFCAC IGLACAHNYAQI LALRCLQAAGI SPVIAINS  
 GIMGDVTTKVERGGYVGLVAGFQVVGTAFGALIGAGLSSKKGWRAIFWFLAIGSGICLVF  
 STLLMPETKRTL VGNGSVTPRSFLNRSILHVGSVKKTLHLDDPDPEPTLEPRTSVDFLAP  
 LKILHIREI DILLS IAGLQFSTWTHQTALTIVLSKKYNLSVAKIGLCFLPAGISTLTSI  
 ISAGRYLNWSYRTRKVKYNRWIkeQELQlMEKYKGDKNKVAELIHSNSHYAFNLVEARLH  
 PAFVTLSSIGFTAFGWCISVKTPLAAVLCTSAFASLFSNCILTFSTTLIVDLFPSKAS  
 TATGCLNLFRCLLSAIFIAALTKMVEKMRYGGVFTFLSAITSSSSLLLLFYLLKNGKQLSF  
 DRIRANDKSAGRSVGKNSEKSVT

SEQ ID NO: 120  
 YIL121W

>sp|P40474|QDR2\_\_YEAST Quinidine resistance protein 2  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=QDR2  
 PE=1 SV=1  
 MAGATSSIIRENDFEDELAESMQSYNRETADKLALTRTESVKPEPEITAPPHSRFSRSFK  
 TVLIAQCAFTGFFSTIAGAIYYPVLS VIERKFDIDEELVNVTVVVYFVFQGLAPTfMGGF  
 ADSLGRPVVLAIVI YFGACIGLACAQTYAQI IVLRCLQAAGI SPVIAINSgIMGDVTT  
 RAERGGYVGYVAGFQVLGSAFGALIGAGLSSRWGWRAI FWFLAIGSGICFLASFLILPET  
 KRNISGNGSVTPKSYLNRAPILVLPVTRKSLHLDNPDYETLELPTQLNLLAPFKILKAYE  
 ICILMLVAGLQFAMYTTHLTALSTALSKQYHLTVAKVGLCYLPSGICTLCSI VIAGRYLN  
 WNYRRRLKYYQNWLKGRKSKLLEHDNDLNLVQRIIENDPKYTFNIFKARLQPAFVTLILL



SSSGFCAYGWCITVKAPLA AVL CMSGFASLFSNCILTFSTTLI VDLFP TKTSTATGCLNL  
 FRCILSAVFIAALS KMVEKMKFGGVFTFLGALTSSSSILLFILLRKGKELAFKRKKQELG  
 VN

SEQ ID NO: 121

YIL166C

>sp|P40445|YIQ6\_\_YEAST Uncharacterized transporter YIL166C  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=YIL166C  
 PE=1 SV=1  
 MSVQKEEYDIVEKAQLSVSAESLTSDESISHNPFDDFHKAEWRKRVYESSGYEGLSKFD  
 PEFTWTKDEEKKLVRKMDLKIFLWVFI MFAFLDLIRKNIARAVSDNFIVDLKMNTNDYNL  
 GQTVYLVI FLASELPGNLLSKRFGPERVIPVQIVLWSVICITQAGLKNRGQFIATRCLLG  
 MVQGGFIPDNILYLSYYTGAELTFRLSFFWCAIPLFQILGSL LASGI IEMRGIHNLAGW  
 QYLFII EGFLSLSVGVASFYLMRRGPTQTGESAFHKGKSLFTEYEEKIMVNRILRDDPSK  
 GDMSNRQPVT FKEI LYTLTEFDLWPLFIQGITAFI SLQTVGS YLSLILKSLNYSTFLSNI  
 LAIPGQALLL INLPLAALLSRKLKEKSLCVGIANVWVLPFIVSLVALPTDTPNWI KYILL  
 TGILGLPYTHSILAGWVSEISNSVRSRTVGTALYNMSAQVGAI IASNM YRNDDKPY YTRG  
 NKILLGFTCFNICMAVATKFYYISRNKYKDRKWSMTKEEQINYLDTTKDKGMKRLDYRF  
 IH

SEQ ID NO: 122

YJL133W

>sp|P10566|MRS3\_YEAST Mitochondrial RNA-splicing protein MRS 3  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=MRS3  
 PE=1 SV=4  
 MVENSSNNSTRPIPAIPMDLPDYEALPTHAPLYHQLIAGAFAGIMEHSVMFPIDALKTR  
 IQSANAKSLSAKNMLSQISHISTSEGTLALWKGVSILGAGPAHAVYFGTYEFCKKNLI  
 DSSDTQTHHPFKTAISGACATTASDALMNPFDTIKQRIQLNTSASVWQTTKQIYQSEGLA  
 AFYYSYPTTLVMNI PFAA FN FVI YESSTKFLNPSNEYNPLIHCLCGSISGSTCAAITTPL  
 DCIKTVLQIRGSQTVSLEIMRKADTF SKAASAIYQVYGWKGFWRGWKPRIVANMPATAIS  
 WTAYECAKHFLMTY

SEQ ID NO: 123

YJL219W

>sp|P40885|HXT9\_YEAST Hexose transporter HXT9 OS=Saccharomyces  
 cerevisiae (strain ATCC 204508 / S288c) GN=HXT9 PE=1 SV=1  
 MSGVNNTSANDLSTTESNSNSVANAPSVKTEHNDSKNLSNLDATEPPIDL PQKPLSAYTT  
 VAILCLMIAFGGFI FGWDTGTISGFVNLSDFIRRFQKNDKGTYYLSKVRMGLI VSIFNI  
 GCAIGGIVLSKVGDIYGRRI GLITVTAI YVVGILIQITSINKWYQYFIGRIISGLGVGGI  
 AVLSPMLISEVAPKQIRGTLVQLYQLMCTMGI FLGYCTNYGTKNYHNATQWRVGLGLCFA  
 WTTFMVSGMMFVPESPRYLIE VGDDEEAKRSLSKSNKVSVD DPALLAEYDTIKAGIELEK  
 LAGNASWSELLSTKTKVFQRVLMGVMIQSLQQLTGDNYFFYYGTTI FKS VGLKDSFQTSI  
 IIGVVNFFSSFIAVYTIERFGRRTCLLWGAASMLCCFAVFASVGVTKLWPQGSSHQDITS  
 QGAGNCMIVFTMFFIFSFATTWAGGCYVIVSETFPLRVKSRGMAIATAANW MWGFLISFF  
 TPFITGAINFYGYVFLGCLVFAYFYVFFFVPETKGLTLEEVNTMWLEGPVPAWKSASWVP  
 PERRTADYDADAIDHDDRPIYKRFFSS

SEQ ID NO: 124

YKL016C

>sp|P30902|ATP7\_YEAST ATP synthase subunit d, mitochondrial

OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=ATP7  
 PE=1 SV=2  
 MSLAKSAANKLDWAKVISSLRITGSTATQLSSFKKRNDARRQLLELQSQPTEVDFSHYR  
 SVLKNTSVIDKIESYVKQYKPKIDASKQLQVIESFEKHAMTNAKETESLVSKEKLDLQS  
 TLDNIQ SARPFDELTVDDLTKIKPEIDAKVEEMVKKGKWDVPGYKDRFGNLMVM

SEQ ID NO: 125  
 YKL050C

>sp|P35736|YKF0\_YEAST Uncharacterized protein YKL050C  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / 5288c) GN=YKL050C  
 PE=1 SV=1  
 MSLISALQTTDVESVQTSPEQITERKAVRVSTLQESLHSSSEMHRAPETPRSI SNSVHKL  
 KTIYSTYQQSGQPLSKEAIFRAKQKYGILNTPANYKTLGLGDSKSESVDLAARLASKRTK  
 VSPDDCVETAIEQKARGEAFKVTFSKIPLTPPEDVPI TVNLGLKGRRDFLTRLAQKALA  
 FSPSLDNSMKGTSDSSSVKRRFSGAPIGNEFDANLVNPQHPAGFKSLDLKVLDAEERR  
 AISRVNDRLYPQKVNFKNGLQSSDQSGVSKANKEVFKKGTLEKLEHSAEQFLESHAGNER  
 QRLSDQQYMCAGAADAVKDLDPKTLEDPDFAAREAQKKLYIKQVASPVVLNEAQKLANR  
 KLQDIDSRDYMMLFGNQAYNKLAVNIALQHYSVKQEKKKIYLGGLWMTPEEVNAVAK  
 KLISPVVNEIDERASQRDVKDIERRSRVLDQYEDGNSMERAKEQNDGQLLLAMASKQ  
 QQEKEAKKAEQRYDQFVQKMNILQOKEKELENARENRENLRNELQERLSKNLSGEND  
 ELNDWNDACERDLKNSSIEHYAVRSHFDNLGNSERGYDELLEERSKIQVEIERLVASIA  
 EHKTAIHGFGGETADAGGAIPAVQKQKI PTRKDLLDATVNDPLVISAEMAKEEAEMATEEC  
 MLKELQVDEMI IIRNIMLRECEKLEEEKETAKRSRRGTEESKNNSNFSRDVIMSTPDNN  
 EKVTPIGKSASPDKVKSRLFSTYNTGKDIDSSASARSITGVSGVLDGPKTPTSNKENE  
 LIDDEVKSYKVHQAVDGTGEDSIANKRDKSSRPAANSRGGITIEQFLFNKNADKQGLSKT  
 ESVTMKREPVVDQMSKKGHDFTHCNDNGRRSFSGFSQGS IENDYSNEVTDDQDDQEGSE  
 IRVRDSNDSNTSPKESFFKEVI

SEQ ID NO: 126  
 YKL120W

>sp|P32332|OAC1\_YEAST Mitochondrial oxaloacetate transport protein  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=OAC1  
 PE=1 SV=1  
 MSSDNSKQDKQIEKTAAQKISKFGSVFVAGGLAACIAVTVTNPIELIKIRMQLQGEMSASA  
 AKVYKNPIQGMVIFKNEGIKGLQKGLNAAIYQIGLNGSRLGFYEPISRSSLNQLFFPDQ  
 EPHKVQSVGVNVFSGAASGI IGAVIGSPLFLVKTRLQSYSEFIKIGEQTHTYTGWVWNLVT  
 IFKTEGVKGLFRGIDAAILRTGAGSSVQLPIYNTAKNILVKNDLMKDG PALHLTASTISG  
 LGVAVVMNPWDVILTRI YNQKGDLYKGPIDCLVKTVRIEGVTALYKGF AAQVFRIAPHTI  
 MCLTFMEQTMKLVYSIESRVLGHN

SEQ ID NO: 127  
 YKL146W

>sp|P36062|AVT3\_YEAST Vacuolar amino acid transporter 3  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=AVT3  
 PE=1 SV=1  
 MNGKEVSSGSGRTQSNNNKNNNGGSTGISHASGSPLTDGNGGNSNGNSRSR SRSRKSSG  
 TTGGLLKKPPLLNVNEAVHASVPDASHTSCNNGTLEVSINNPEPHVVD AVARHLIRNPSN  
 SLQLQGGDITRDLYKWTNDHPSSPSQYQYPSQPALSTSI PSQAPSFSNRKRSMFSASA  
 ASSSHLNNNSEANGNPLAAI GLAPAPMTH EIRAPGGFRRSFI IQKRRKHNVDAPI PNF  
 TRNFIEFLTLYGHFAGEDLSEEEEEEEEEETEEPEEEEALETETESTQLVSREHGRHPKSSSTV

KAVLLLLLSFVGTGVLFLPKAFHNGGWGFSALCLLSCALI SYGCFVSLITTKDKVGVVDGY  
 GDMGRILYGPCKMFAILSSIALSQIGFSAAYTVFTATNLQVFSENFFHLKPGSISLATYI  
 FAQVLI FVPLSLTRNIAKLSGTAL IADL FILLGLVYVYVYSI YYIAVNGVASDTMLMFNK  
 ADWSLFIGT AIFTFEGIGLLIPIQESMKHPKHFRPSLSAVMCI VAVIFISCGLLCYAAF  
 SDVKT VVLLNFPQDTSYTLTVQLLYALAILLSTPLQLFP AIRILENWT FPSNASGKYNPK  
 VKWLKNYFRCAIVVLT SILAWVGANDLDK FVSLVGSFACIPLI YIYPPLLHYKASILSGT  
 SRARLLLDLI VIVFGVAVMAYTSWQT IKMWSQ

SEQ ID NO: 128

YKL209C

>sp| P12866 ISTE6\_\_YEAST Alpha-factor-transporting ATPase  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / 5288c) GN=STE6  
 PE=1 SV=1

MNFLSFKTTKHYHIFRYVNIRNDYRLMIMI IGT VATGLVPAITSILTGRVFDLLSVFVA  
 NGSHQGLYSQLVQRSMAMVMAALGAASVPMWLSLTSWMHIGERQGFRI RSQILEAYLEEK  
 MEWYDNNEKLLGDFTQINRCVEELRSSSAEASAITFQNLVAICALLGTSFYYSWLSLTLII  
 LCSSP 1ITFFAVVFSRMIHVYSEKENSETSKAAQLLTWSMNAAQLVRLYCTQRLEKFK  
 EIILNCNTFFIKSCFFVAANAGILRFLTLTMFVQGFWFGSAMIKKGLNINDVITCFHSC  
 IMLGSTLNNTLHQIVVLQKGGVAMEKIMTLLKDGSKRNPLNKTVAHQFPLDYATSDLTFA  
 NVSFSYPSRPSEAVLKNVSLNFSAGQFTFIVGKSGSGKSTLSNLLLRFYDGYNGSISING  
 HNIQTIDQKLLIENITVVEQRCTLFNDTLRKNILLGSTDSVRNADCSTNENRHLIKDACQ  
 MALLDRFILDLPDGLLETIGTGGVTLSSGGQQORVAIARAFIRDTPILFLDEAVSALDIVH  
 RNLLMKAIRHWRKGT T I ILTHELSQIESDDYLYLMKEGEVVESGTQSELLADPTTTFST  
 WYHLQNDYSDAKTIVDTETEEKSIHTVESFNSQLETPKLGSCLSNLGYDETDQLSFYEAI  
 YQKRSNVRTRRVKVEEENIGYALKQKNTESSTGPQLLSIIQI IKRMIKSIRYKILILG  
 LLC SLIAGATNPVFSYTF SFLLEGIVPSTDGKTGSSHYLAKWSSLVLGVAAADGI FNFAK  
 GFLLDCCSEYWMDLRNEVMEKLRTRKNMDWFSGENNKASEISALVNLDRDLRSLVSEFL  
 SAMTSFVTVSTIGLIWALVSGWKL SLVCISMFLI IIFSAI YGGILQK CETDYKTSVAQL  
 ENCLYQIVTNIKTIKCLQAEHFHQLT YHDLKIKMQQIASKRAIATGFGISMTNMIVMCIQ  
 AIIYYYGLKLVMIHEYTSKEMFTTFTLLFTIMSCTSLVSQI PDISRGQRAASWI YRILD  
 EKHNTLEVENNNARTVGIAGHTYHGKEKKPIVSIQNLTFAYPSAPTA FVYKNMNFDMFCG  
 QTLGIIGESGTGKSTLVLLLTKLYNCEVGKIKIDGTDVNDWNLTSLRKEISVVEQKPLLF  
 NGTIRDNLTYGLQDEILEIEMYDALKYVGIHDFVISSPQGLDTRIDTLLSSGGQAQRLCI  
 ARALLRKSILILDECTSALDSVSSSIINEIVKKGPPALLTMVITHSEQMMRSCNSIAVL  
 KDGKVVVERGNFDTLYNNRGELFQIVSNQSS

SEQ ID NO: 129

YKR039W

>sp| P19145|GAP1\_\_YEAST General amino-acid permease GAP1  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=GAP1  
 PE=1 SV=2

MSNTSSYEKNNPDNLKHNGITIDSEFLTQEPITI PSNGSAV SIDETGSGSKWQDFKDSFK  
 RVKPIEVDPNLSEAEKVAI ITAQTP LKHHLKNRHLQMI AIGGAIGTGLLVGSGTALRTGG  
 PASLLIGWGSTGTMI YAMVMALGELAVI FPISGGFTTYATRFIDESFGYANNFN YMLQWL  
 VVLPLEIVSASITVNFWGTDPKYR DGFVALFWLAIVI INMFGVKGYGEAEFVFSFIKVIT  
 VVGFIIILGI ILNCGGGPTGGYIGGKYWHDPGAFAGDTPGAKFKGVCSVFVTAAFSFGASE  
 LVGLAASESVEPRKSVPKAAKQVFWRITLFYILSLLMIGLLVPYNDKSLIGASSVDAAAS  
 PFVIAIKTHGIKGLPSVNVVILIAVLSVGN SAIYACSR TMVALAEQRFLPEIFSYVDRK  
 GRPLVGI AVTSAFGLIAFVAASKKEGEVFNWLLALSGLSSLFTWGGICICHIRFRKALAA  
 QGRGLDELSFKSPTGVWGSYWGLFMVIMFIAQFYVAVFPVGDSPSAEGFFEAYLSFPLY  
 MVMYIGHKIYKR NWKLFIPA EKMDIDTGRREVLDLLKQEIAE EKAIMATKPRWYRIWNF  
 Wc

SEQ ID NO: 130

YLR411W

>sp|Q06686|CTR3\_YEAST Copper **transport** protein CTR3 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=CTR3 PE=1 SV=1  
 MNMGSSSTAACKATCKISMLWNWYTIDTCFIARSWRNDTKGKFAGSCIGCFALVVVAQW  
 LTRFSRQFDVELLKRQKIKHLASYSPEEYVVKCGEEDAKSDIEELQGFYNEPSWKTTLIS  
 LQKSFI YSFYVWGPRLNEPEDDLLKVLSCCTLITPVDLYPTFLDHMIRVTI FVLQWGL  
 SYIIMLLFMYNGYI I I SCLIGAI VGRFI FCYEPLGSLGANGSAQGTVSYDKESDDRKCC  
 L

SEQ ID NO: 131

YML038C

>sp|Q03697|YMD8\_YEAST Putative nucleotide-sugar transporter YMD8 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=YMD8 PE=1 SV=1  
 MNRTVFLAFVFGWYFCSIALSIYNRWMFDPKDLGIGYPVLVTTTFHQATLWLLSGIYIKL  
 RHKPVKNVLRKNNGFNWSFFLKFLPTAVASAGDIGLSNVSFQYVPLTIYTI IKSSSI AF  
 VLLFGCIFKLEKFWKLALSVI IMFVGVALMVFKPSDSTSTKNDQALVIFGSFLVLASSC  
 LSGLRWVYQTQLMLRNNPIQTNTAAVEESDGLFTENEDNVDNEPVVNLANNKMLENFGE  
 SKPHPIHTIHQLAPIMGITLLLLTSLLVKPPFGI FSSSIFRLDTSNGGVGTETTIVLSIVR  
 GIVLLILPGFAVFLLTICEFSILEQTPVLTVSI VGIVKELLTVIFGI IILSERLSGFYNW  
 LGMLI IMADVCIYNYFRYKQDLLQKYHSVSTQDNRNELKGFQDFEQLGSKKIAPYSISVD  
 LTNQEYELDMIAQNVSRSQQV

SEQ ID NO: 132

YMR166C

>sp|Q03829|YM39\_YEAST Uncharacterized mitochondrial carrier YMR166C OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=YMR166C PE=1 SV=1  
 MNSWNLSSSIPI IHTPHDHPPTSEGTDPQPNNNRDKDKLHKKRGDSDEDLSPIWHCVVSG  
 GIGGKIGDSAMHSLDVTQKTRQQGAPNVKKYRNMISAYRTIWLEEGVRRGLYGGYMAAMLG  
 SFPSAAI FFGTYEYTKRTMIEDWQINDTITHLSAGFLGDFISSFVYVPSEVLKTRLQLQG  
 RFNPNPFQSGYNYSNLRNAIKTVIKEEGFRSLFFGYKATLARDLPFSALQFAFYEKFRQL  
 AFKIEQKDRDGEISIPNEILTGACAGGLAGIITTPMDVVKTRVQTQQPPSQQSNKYSV  
 HPHVTNGRPAALSNSISLSLRVYQSEGLVGGFSGVGRFVWTSVQSSIMLLLYQMTLRG  
 LSNAPPTD

SEQ ID NO: 133

YMR27 9C

>sp|Q03263|**Y**M8M\_YEAST Uncharacterized transporter YMR279C OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=YMR27 9C PE=1 **SV=1**  
 MFSIFKKKTSVQGTDSSEIDEKITVKAKDKVVVSTEDDEEVTIVSSTKSTQVTNDSPWQDP  
 TYFSSFGKELMFIATCMLAQLLNQAGQTHALCIMNVLSKSFNSEANNQAWLMASFLAAG  
 SFILISGRLGDIYGLKKMLIVGYVIVIVWSIISGLSKYSNSDAFFITSRAFQGVGIAFIL  
 PNIMGLVGHVYKVSFRKNIVISFIGACAPTGGMFGGLFGGLIVTEDPNQWPWFYAFGI  
 ATFLSLLMAWYSI PNNVPTNIHGLSMDWTGSALAIIGLILFNFWNQAPIVGWDKPYIIV  
 LLIISVI FLVAFFVYESKYAEVPLLPRAMTKNRHMIMILLAVFLGWGSFGIWTFFYVVSFQ  
 LNLRHYSVWVTGGTYFVVFVIFGSMAAFFVAFSIKRLGPALLLFCFLMAFDAGSIMFSVLP  
 VEQSYWKLNFAMQAILCFGMDLSPASSIILSDGLPMQYQGMAGSLVNTVINYSASLCLG  
 MGGTVEHQINKSGNDLLKGYRAAVYLVGLASLGVVISVTYMLENLWNRHRKSEDRSLEA

SEQ ID NO: 134

YNL003C

>sp! P38 921 |PET8\_YEAST Putative mitochondrial carrier protein PETS  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=PET8  
 PE=1 SV=1  
 MNTFFLSLLSGAAAAGTSTDLVFFPIDTIKTRLQAKGGFFANGGYKGIYRGLGSASAVVASAP  
 GASLFFISYDYMVKVSRPYISKLYSQGSEQLIDTTTHMLSSSIGEICACLVVRVPAEVVKQ  
 RTQVHSTNSSWQTLQSILRNDNKEGLRKNLYRGWSTTIMREIPFTCIQFPLYEYLKKTWA  
 KANGQSQVEPWKGAICGSIAGGIAAATTTPLDFLKTRLMLNKTTASLGSVI IRIYREEGP  
 AVFFSGVGPRTMVISAGGAI FLGMYETVKSLLSKSFPTAGEMRA

SEQ ID NO: 135

YNL2 68W

>sp !P32487 jLYP1\_\_YEAST Lysine -specific permease OS=Saccharomyces  
 cerevisiae (strain ATCC 204508 / S288c) GN=LYP1 PE=1 SV=2  
 MGRFSNI ITS NKWDEKQNNIGE QSMQELPEDQIEHEMEAIDPSNKTTTPYSIDEKQYNTKK  
 KHGSLQGGAIADVNSITNSLTRLQVVS HETDINEDEEEAHYEDKHVKRALKQRHIGMIAL  
 GGTIGTGLFVIGISTPLSNAGPVGSLIAYIFMGTIVYFVTQSLGEMATFIPVTSSITVFSK  
 RFLSPA FGVSNGYMYWFNWAITYAVEVSVIGQVIEYWTDKVPLAAWIAIFWVI I TLMNFF  
 PVKVYGEFEFVWASVKVLAIMGYLIYALIIVCGGSHQGP IGFYWRNPGAWGPGI ISSDK  
 SEGRFLGWVSSLINAAFTYQGT ELVGITAGEAANPRKTVPRANKVVFRIVLFYIMSLFF  
 IGLLVPYND S RLSASSAVIASSPFVVISIQNAGTYALPDIFNAVVLITVVSAA NSNVYVGS  
 RVLYSLARTGNAPKQFGYVTRQGV P YLGVVCTAALGLLAFLVNNNANTAFNWLINISTL  
 AGLCAWLFISLAHIRFMQALKHRGISRDDLPFKAKLMPYGAYYAAFFVTVIIIFIQGFQAF  
 CPFKVSEFFTSYISLILLAVVFIGCQIYYKCRFIWKLEDIDIDSDRREIEAIIWEDDEPK  
 NLWEKFWAAVA

SEQ ID NO: 136

YNR055C

>sp| P53389 **HOL1\_YEAST** Protein HOL1 OS=Saccharomyces cerevisiae  
 (strain ATCC 204508 / S288c) GN=HOL1 PE=1 SV=1  
 MDKYTNRDHPDYIPGTFNIYSSQNL ENGI IYESKLKKTSSGVVLIPQPSYSPNDPLNWSS  
 WRKLAHFGLMAFI TAFTAATSNDAGAAQDSLNE IYGI SYDSMNTGAGVLF LG IGW5TLFL  
 APFANLYGRKITYI VCTTLGLFGALWFALAKRTSDTIWSQLFVGISESCAEAQVQLSLSD  
 IFFQHQLGSLVLT VYIMCTSIGTFLGPLIAGYISAFTNFRWGWVAVI ISGGLLIT IIFGC  
 EETYFDRGQYMTPLTSCQSGYEDGTTLQNSDNTAVSRRKRHLDAKLSTPGAMGEKGV DLS  
 ETAEFEVNNEEEVTIPETRELIDGSKEHLKPYPKRVA I LTKATNLKGYGFKQYFKYLKIN  
 LRMFLFPVWLSGMFWGIQDVFLTFYLT TQESAYYEPPWNYSDFGVAIMNVPTLIGAVIG  
 CICAGIVSDYFVLWMARHNRGILEAEFRLYFSIATAI IGPA GLLMFGIGTARQWPWQAI Y  
 VGLGFVGFAGCSGDIAMAYLMDCYPDMVLEGMVCTAI INNTISCI FTFTCSDWLAASGT  
 ENTYIALAVINFGITAFALPMYYYGKRIRLWTKRWYLQSVNLRDGV

SEQ ID NO: 137

YOL158C

>sp|Q08299 |**ENB1\_YEAST** Siderophore iron transporter ENB1  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / 5288c) GN=ENB1  
 PE=1 SV=1  
 MLETDHSRNDNLDDKSTVCYSEKTD SNVEKSTTSGLRRIDAVNKVLSDYSSFTAFGVTF S

SLKTALLVALFLQGYCTGLGGQISQS IQTYAANSFGKHSQVGSINTVKS IVASVVAVPYA  
 R I SDRFGR I ECW I FALVLYT I GE I ISAATPTFSGLFAGI VIQQFGYSGFRLLATALTGDL  
 SGLRDRTFAMNIFLIPVI INTWVSGNIVSSVAGNVAPYKWRWGYGIFCIIVPISTLILVL  
 PYVYAQYISWRSGKLPPLKLEKQTLRQTLWKFADDINLIGVILFTAFLVLVLLPLTIA  
 GGATSKWREGHIIAMI VVGGCLGFI FLIWELKFAKNPFI PRVYLGDP TI YVALLMEFVWR  
 LGLQIELEYLVTVLMAVAFGESTLSAQRIAQLYNFLQSC T N I VVGIMLHFYHPKVFVAVG  
 SLLGVIGMGLLYKYRVVYDGISGLIGAEIVVGIAGGMIRFPMWTLVHASTH NEMATVTG  
 LLMSVYQ IGDVAVGASI AGA I W T Q R L A K E L I Q R L G S S L G M A I Y K S P L N Y L K K Y P I G S E V R V  
 QMIESYSKIQRLLIIVSISFAAFNAVLCFFLRGFTVNKKQSLSAEEREKEKLEKIKQOSWL  
 RRVIGY

SEQ ID NO: 138

YOR100C

>sp | Q12289 | CRC1\_YEAST Mitochondrial carnitine carrier  
 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=CRC1  
 PE=1 SV=1

MSSDTSLSSESLLEESGLTKSRPPIKSNPVRENIKS FVAGGVGGVCAVFTGHPFDLI K  
 VRCQNGQANSTVHAITNIIKEAKTQVKGTLFTNSVKGFYKGVIPPLLGVTPIFAVSWFGY  
 DVGKKLVTFNNKQGGSNELTMGQMAAAGFISAIPTTLVTAPTERVKVVLQTSSKGSFIQA  
 AKTI VKEGGIASLFGSLATLARDGPGSALYFASYEISKNYLNSRQPRQDAGKDEPVNII  
 NVCLAGGIAGMSMWLAVFPIDTIKTKLQASSTRQNMLSATKEIYLRGGIKGFFPGLGPA  
 LLRSFPANAATFLGVEMTHSLFKKYGI

SEQ ID NO: 139

YOR153W

>sp | P33302 | PDR5\_YEAST Pleiotropic ABC efflux transporter of multiple  
 drugs OS=Saccharomyces cerevisiae (strain ATCC 204508 / 5288c)  
 GN=PDR5 PE=1 SV=1

MPEAKLNNNVNDVTSYSSASSSTENAADLHNYNGFDEHTEARIQKLARTLTAQSMQNSTQ  
 SAPNKSDAQSI FSSGVEGVNPI FSDPEAPGYDPKLDPNSENFSSAAWVKMAHL SAADPD  
 FYKPYSLGCAWKNL S A S G A S A D V A Y Q S T V V N I P Y K I L K S G L R K F Q R S K E T N T F Q I L K P M D  
 GCLNPGELLVVLGRPGSGCTLLKSISSNTHGFDLGADTKISYSGYSGDDIKKHFRGEV  
 YNAEADVHLPHLTVFETLVTVARLKTQNR I K G V D R E S Y A N H L A E V A M A T Y G L S H T R N T K  
 VGNDIVRGVSGGERKRV S I A E V S I C G S K F Q C W D N A T R G L D S A T A L E F I R A L K T Q A D I S N T  
 SATVAIYQCSQDAYDLFNKVCVLDGQYIYYGPADKAKKYFEDMGYVCP SRQT TADFLTS  
 VTSPSERTLNKDMKKGIHIPQTPKEMNDYWVKSPNYKELMKEVDQRLLNDD EASREAIK  
 EAHIAKQSKRARPPSPYTVSYMMQVKYLLIRNMWRLRNNIGFTLFMILGNCSMALILGSM  
 FFKIMKKGDTSTFYFRGSAMFFAILFNAFSSLEIFSLYEARPITEKHRTYSLYHPSADA  
 FASVLSEIPSKLIIAVCFNI IFYFLVDFRRNGGVFFFYLLINIVAVFMSHLFRCVGS LT  
 KTLSEAMVPASMLLLALSMTGFAIPKKKILRWSKWIWYINPLAYLFESLLINEFHGIKF  
 PCAEYVPRGPAYANISSTESVCTVVGAVPGQDYVLGDDFIRGTYQYYHKDKWRGFGIGMA  
 YVVFVVVYFLCEYNEGAKQKGEILVFP RSI VKRMKKRGVLTEKNANDPENVGERSDLS  
 SDRKMLQESSEEEEDTYGEIGLSKSEAI FHW R N L C Y E V Q I K A E T R R I L N N V D G W V K P G T L  
 TALMGASGAGKTTLLDCLAERTVMGVITGDILVNGIPRDKSFPRSIGYCQQQDLHLKTAT  
 VRESLRF SAYLRQPAEVSIEKNRYVEEVIKILEMEKYADAVVGVAGEGLNVEQRKRLTI  
 GVELTAKPKLLVFLDEPTSGLD S Q T A W S I C Q L M K K L A N H G Q A I L C T I H Q P S A I L M Q E F D R  
 L L F M Q R G G K T V Y F G D L G E G C K T M I D Y F E S H A H K C P A D A N P A E W M L E V V G A A P G S H A N Q D  
 Y Y E V W R N S E E Y R A V Q S E L D W M E R E L P K K G S I T A A E D K H E F S Q S I I Y Q T K L V S I R L F Q Q Y W  
 R S P D Y L W S K F I L T I F N Q L F I G F T F F I C A G T S L Q G L Q N Q M L A V F M F T V I F N P I L Q Q Y L P S F V  
 Q Q R D L Y E A R E R P S R T F S W I S F I F A Q I F V E V P W N L A G T I a y f . i Y Y Y P I G F Y S N A S A A G Q L  
 H E R G A L F W L F S C A F Y V Y V G S M G L L V I S F N Q V A E S A A N L A S L L F T M S L S F C G V M T T P S A M P  
 R F W I F M Y R V S P L T Y F I Q A L L A V G V A N D V K C A D Y E L L E F T P P S G M T C G Q Y M E P Y L Q L A K T

GYLIDENAI DTCSFCQISTINDYLANVNSFYSERWRNYGIFICYIAFN YIAGVFFYWLAR  
VPKKNGL5KK

SEQ ID NO: 140

YOR271C

>sp|Q12029|FSF1\_YEAST Probable mitochondrial transport protein FSF1  
OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=FSF1  
PE=1 SV=1  
MASSVPGPIDLPESRYDLSTYWGRIHCAEISDPTMLLTTEKDLAHAREI ISAYRHGELK  
ETTPEFWRAKKQLDSTVHPDTGKTVLLPFRMSSNVLSNLVVTVGMLTPGLGTAGTVFWQW  
ANQSLNVAVNSANANKSHPMSTSQLLTNYAAAVTASCVALGLNNLVPRLKNISPHSKLI  
LGRVLPFAAVVSAGIVNVFLMRGNEIRKGISVFD SNGDEVGKSKKAAFMAVGETALSRVI  
NATPTMVIPLILVRLQRGV LKGLVQTLANLGLISVTMFSALPFALGIFPQRQAIHL  
NKLEPELHGKKDKDGKPIEKVYFN RGI

SEQ ID NO: 141

YOR273C

>sp|Q12256|ITP04\_YEAST Polyamine transporter 4 OS=Saccharomyces  
cerevisiae (strain ATCC 204508 / 5288c) GN=TP04 PE=1 SV=1  
MPSSLTKTESNSDPRNTNIQQVPKALDKNVTNSGNLDSTSSSTGSITEDEKRSEPNADSNN  
MTGGEPIDPRDLWDGPDNDPNHNWSSLK KWTMTSAFLCLVVTMGSSLYVSSVPELV  
ERYHVSQTLALAGLTFYLLGLSTVIGAPLSEVFG RKPVYLFSLPVSMLFTMGVGLSNGHM  
RIILPLRFLSGV FASPALS VSGSTILDI FDVDQVSVAMTYFVLS PFLGPVLSPI MAGFAT  
EAKGWRWSEW IQLIAGGLILPFIALMPETHKGI ILRKRKRNRNIALKKFSREAQKEFLKT  
TVTITILRPLKMLVVEPIV FVFSVYVAFIFAILFGFFEAYAVIYRGVYHMSMGISGLPFI  
GIGVGLWIGAFFYLYIDRKYLPKPPAGTQPLTEKERTSKRTTPYRGARDAETGELLPVV  
PEKFLIACKFGSVALPIGLF¼'QAWTARS DVHMAPVAAGVPFGFGLILI FFSVLMYFSTC  
YPPLTVASCLAANLLRYVMSSVFLFTIQMYTKMKIKWASTLFALVCVMMI PIPWVFEK  
WGSKLRHKSQFGYAAMEKEAETE GIDDVNAV D GELNLTRMTTLRTMETDPSTREK PGER  
LSLRRTHTQPVPASFDRE D GQHAQNRNEPISNSLYSAIKDNEDGYSYTEMATDASARMV

SEQ ID NO: 142

YOR307C

>sp|P22215|SLY41\_YEAST Uncharacterized transporter SLY41  
OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=SLY41  
PE=1 SV=2  
MIQTQSTAIKRRNSVHKNLFDPSLYQIPEPPRGGFQHQKKEYSKETFSNQVFGYDITSLK  
KRFTQLFSPNIQGYLPEVDLRITI ICSIWYVTSSISSNLSKAILRTFNHPIALTELQFLV  
SAVLCVGFASIVNLFRLPRLKHTKFSKALNSFPDGILPEYLDGNFRSSILHKFLVPSKLV  
LMTTFPMGIFQFIGHITSHKAVSMIPVSLVHSV KALSPI ITVGYKFFEHRYNSMTYYT  
LLLLI FGVMTTCWSTHGSKRASDNKSGSS LIGLLF AFISMI IFVAQNI FAKNILTIRRKV  
GILPSSSTDDVTSKEGQPSLDKTRFSPLQVDKITILFYCSCIGFSLTLLPFLT GELMHGG  
SVINDLTLETVALVAIHGIAHFFQAM LAFQLIGLLSSINYSVANIMKRIVVISVALFWET  
KLNFFQVFGVILT IAGLYGYDKWGLSKKDGRQA

SEQ ID NO: 143

YOR332W

>sp|P22203|VATE\_YEAST V-type proton ATPase subunit E  
OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=VMA4  
PE=1 SV=4  
MSSAITALTPNQVNDELNKMQAFIRKEAEEKAKEIQLKADQEYEIEKTNIVRNETNNDIG  
NFKSKLKKAMLSQQITKSTIANKMRLKVL SAREQSLDGI FEETKEKLSGIANNRDEYKPI  
LQSLIVEALLKLLPEPKAIVKALERDV D LIESMKDDIMREYGEKAQRAPLEEIVISNDYLN

KDLVSGGVVSNASDKIEINNTLEERLKLSEEALPAIRLELYGPSKTRKFFD

SEQ ID NO: 144

YOR348C

>sp|P15380|PUT4\_YEAST Proline-specific permease OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=PUT4 PE=1 SV=2  
 MVNILPFHKNNRHSAGVVTCAADVSGDGGSGDTKKEEDVVQVTESPSSGSRNNHRSDNEK  
 DDAIRMEKISKNSASSNGTIREDLIMDVLEKSPSVVDGSEPHKQGLQSRHVQLI  
 GGAIGTGLLVGTSTLHTCGPAGLFISYIIISAVI YPIMCALGEMVCFPLPGDSDSAGST  
 ANLVTRYVDP SLGFATGWNYFYCYVILVAAECTAASGVVEYWTAVPKGVWITIFLCVVV  
 IILNFSAVKVYGESEFWFASIKILCIVGLI ILSFILFWGGPNHDLGFRYQHPGAF  
 AHH LTGGSLGNFTDI YTG IKGAFAFILGPELVCM TSAECADQRRNI AKASRRFVWRLI  
 FFYV LGTLAISVIVPYNDPTLVNALAQKPGAGSSPFVIGIQNAGIKVLPHI INGCILT  
 SAWSA ANAFMFASTRSLLTMAQTGQAPKCLGRINKWGVYVAVGVSFLCSCLAYLNVSS  
 STADV NWFNSNISTISGFLGWMCGCIAYLRFKAI FYNGLYDRLPFKTWGQPYTVWFSL  
 IVIGIIT ITNGYAFIFPKYWRVADFIAAYITLPIFLVLWFGHKLYTRTWQWLPVSEID  
 VTTGLVE IEEKSREIEEMRLPPTGFKDKFLDALL

SEQ ID NO: 145

YPL036W

>sp|P19657|IPMA2\_YEAST Plasma membrane ATPase 2 OS=Saccharomyces cerevisiae (strain ATCC 204508 / S288c) GN=PMA2 PE=1 SV=3  
 MSSTEAKQYKEKPSKEYLHASDGDPPANNSAASSSSSSSTSTSASSSAAA VPRKAAAASA  
 ADDSDSDEDIDQLIDELQSNYEGEGDES GEEVTRTDGVHAGQRVVPEKDLSTDPAYGLTSD  
 EVARRRKKYGLNQMAEENESLIVKFLMFFVGP IQFVMEAAA ILAAGLSDWVDVGVICALL  
 LLNASVGF IQEFQAGSIVDELKKTLAN TATVIRDGQLIEIPANEVVPGEILQLES GTIAP  
 ADGRIVTEDCFLQIDQSAITGESLAAEKHYGDEVFSSSTVKTGEAFMVVTATGDNTFVGR  
 AAALVGQASGVEGHFTEVLNGIGI ILLV LVIATLLLVWTACFYRTVGVIVSILRYTLGITI  
 IGVPVGLPAVVTTTMAVGAAYLAKKQAI VQKLSAIESLAGVEILCSDKTGTLTKNKL SLH  
 EPYTVEGVSPDDLMLTACLAASRKKKGLDAI DKAFLKSLIEYPKAKDALTKYKVLEFHPF  
 DPVSKKVTAVVESPEGERIVCVKGAPL FVLKTV EEDHPIPEDVHENYENKVAELASRGFR  
 ALGVARKRGEHWEILGVMPCMDPPRDDTAQTINEARNLGLRIKMLTGDAVGI AKETCRQ  
 LGLGTN IYNAERLGLGGGDMPGSELAD FVENADGFAEVFPQH KYRVEILQNRGYLVAM  
 TGDGVNDAPSLKKADTGI AVEGATDAARSAADIVFLAPGLSAI IDALKTSRQIFHRMYSY  
 VVYRIALS LHLEIFLGLWIAILNNSLDINLIVFAI FADVATLTIAYDNAPYAPEPVKWN  
 LPR LWMSIILGIVLAIGSWITLTTMFLPNGGI IQNFGAMNGVMFLQISLTENWLI FVTR  
 AAGPFWSSIPSWQLAGAVFAVDIIATMFTLFGWSENWTDIVSVVRVWIWSIGIFCVLGG  
 FYYIMSTSQA FDRLMNGKSLKEKKSTRS VEDFMAAMQRVSTQHEKSS

SEQ ID NO: 146

YDL198C

MPHTDKKQSG LARLLGSASA GIMEIAVFHP VDTISKRLMS NHTKITSQGE LNRVIFRDHF  
 SEPLGKRLFT LFPGLGYAAS YKVLQRVYKY GGQPFANEFL NKHYKKDFDN LFGEKTGKAM  
 RSAAAGSLIG IGEIVLLPLD VLKIKRQTNP ESFKGRGFIK ILRDEGLFNL YRGWGWTAAAR  
 NAPGSFALFG GNAFAKEYIL GLKDYSQATW SQNFISSIVG ACSSLIVSAP LDVIKTRIQN  
 RNFDNPESGL RIVKNTLKNE GVTAFFKGLT PKLLTTGPKL VFSFALAQSL IPRFDNLLSK

SEQ ID NO: 147

YFL054C

MSYESGRSSS SSESTRPPTL KEEPNGKIAW EESVKKSRN NENDSTLLRR KLGETRKAIE  
 TGGSSRNKLS ALTPLKKVVD ERKDSVQPQV PSMGFTYSLP NLKTLNSFSD AEQARIMQDY  
 LSRGVNQNS NNYVDPLYRQ LNPTMGSSRN RPVWSLNQPL PHVLDRLGLAA KMIQKNMDAR  
 SRASSRRGST DISRGGSTTS VKDWKRLLRG AAPGKKLGD I EAQTQRDNTV GADV KPTKLE



PENPQKPSNT	HIENVSRRKK	RTSHNVNFSL	GDESYASSIA	DAESRKLKMN	QTLDGSTPVY
TKLPEELIEE	ENKSTSALDG	NEIGASEDED	ADIMTFPNFW	AKIRYHMREP	FAEFLGTLVL
VIFGVGGNLQ	ATVTKGSGGS	YESLSFAWGF	GCMLGVYVAG	GISGGH.TNPA	VTISMAIFRK
FPWKKVPVYI	VAQIIGAYFG	GAMAYGYFWS	SITEFEGGPH	IRTTATGACL	FTDPKSYVTW
RNAFFDEFIG	ASILVGCLMA	LLDDSNAPPG	NGMTALIIGF	LVAAIMALG	YQTSFTINPA
RDLGPRIFAS	MIGYGPHAFH	LTHWWWTWGA	WGGPIAGGIA	GALIYDIFIF	TGCESPVNYP
DNGYIENRVG	KLLHAEFHQN	DGTVSEDESG	NSNSNTGSKK	SVPTSS	

SEQ ID NO: 148

*Oryza sativa* sequence encoding EUGT11

MDSGYSSSYA	AAAGMHVIC	PWLAFGHLLP	CLDLAQRLAS	RGHRVSFVST	PRNISRLPPV
RPALAPLVAE	VALPLPRVEG	LPDGAESTND	VPHDRPDMVE	LHRRAFDGLA	APFSEFLGTA
CADWVIVDVF	HHWAAAAALE	HKVPCAMMLL	GSAHMIAAIA	DRRLERAETE	SPAAAGQGRP
AAAPTFEVAR	MKLIRTKGSS	GMSLAERFSL	TLRSSSLVVG	RSCVEFEPET	VPLLSTLRGK
PITFLGLMPP	LHEGRREDGE	DATVRWLDLQ	PAKSVVYVAL	GSEVPLGVEK	VHELALGLEL
AGTRFLWALR	KPTGVSDADL	LPAGFEERTR	GRGVVATRWV	PQMSILAHAA	VGAFLLTHCGW
NSTIEGLMFG	HPLIMLPIFG	DQGPNARLIE	AKNAGLQVAR	NDGDGSFDRE	GVAAAIRAVA
VEEESKVFQ	AKAKKLQEI V	ADMACHERYI	DGFIQQLRSY	KD	

SEQ ID NO: 149

*Synechococcus* sp. GGPPS

MVAQTFNLDT	YLSQRQQQVE	EALSAALVPA	YPERIYEAMR	YSLLAGGKRL	RPILCLAACE
LAGGSVEQAM	PTACALEMIH	TMSLIHDDL	AMDNDDFRRG	KPTNHKVFGE	DIAILAGDAL
LAYAFEHI.A3	QTRGVPPQLV	LQVIARIGHA	VAATGLVGGQ	VVDLESEGKA	ISLETLEYIH
SHKTGALLEA	SVVSGGILAG	ADEELLARLS	HYARDIGLAF	QIVDDILDVT	ATSEQLGKTA
GKDQAAAKAT	YPSLLGLEAS	RQKAEELIQS	AKEALRPYGS	QAEPLALAD	FITRRQH

SEQ ID NO: 150

*Zea mays* truncated CDPS

MAQHTSESAA	VAKGSSLTPI	VRTDAESRRT	RWPTDDDDAE	PLVDEIRAML	TMSDGDISV
SAYDTAWVGL	VPRLDGGEGP	QFPAAVRWIR	NNQLPDGSWG	DAALFSAYDR	LINTLACVVT
LTRWSLEPEM	RGRGLSFLGR	NMWKLATEDE	ESMPIGFELA	FPSLIELAKS	LGVHDFPYDH
QALQGIYSSR	EIKMKRIPKE	VMHTVPTSIL	HSLEGMPGLD	WAKLLKLQSS	DGSFLFSPAA
TAYALMNTGD	DRCFSYIDRT	VKKFNGGVPN	VYPVDLFEHI	WAVDRLERLG	ISRYFQKEIE
QCMDYVNRHW	TEDGICWARN	SDVKEVDDTA	MAFRLLRLHG	YSVSPDVFKN	FEKDGEFFAF
VGQSNQAVTG	MYNLNRSQI	SFPGEDVLHR	AGAFSYEFLR	RKEAEGALRD	KWIISKDLPG
EVVYTLDFPW	YGNLPRVEAR	DYLEQYGGGD	DVWIGKTYLR	MPLVNNDVYL	ELARMDFNHC
QALHQLEWQG	LKRWYTENRL	KDFGVAQEDA	LRAYFLAAAS	VYEPCRAAER	LAWARAAILA
NAVSTHLRNS	PSFRERLEHS	LRCRPSEETD	GSWFNSSSGS	DAVLVKAVLR	LTDSLAREAQ
PIHGGDPEDI	IHKLLRSAWA	EWVREKADAA	DSVCNGSSAV	EQEGSRMVHD	KQTCLLLARM
LEISAGRAAG	EAASEDGDRR	IIQLTGSICD	SLKQKMLVSQ	DPEKNEEMMS	HVDDELKLRI
REFVQYLLRL	GEKKTGSSET	RQTFLSIVKS	CYYAAHCPPH	VVDRHISRVI	FEPVSAAK

SEQ ID NO: 151

*Arabidopsis thaliana* KS (similar to GenBank AEE36246.1)

MSINLRSSGC	SSPISATLER	GLDSEVQTRA	NNVSFEQTKE	KIRKMLEKVE	LSVSAYDTSW
VAMVPSPSSQ	NAPLFPQCVK	WLLDNQHEDG	SWGLDNHDHQ	SLKKDVLSS	LASILALKKK
GIGERQINKG	LQFIELNSAL	VTDETIQKPT	GFDIIFPGMI	KYARDLNLTI	PLGSEVVDDM
IRKRDLKLC	DSEKFSKGRE	AYLAYVLEGT	RNLKDWDLI V	KYQRKNGSLF	DSPATTAADF
TQFGNDGCLR	YLCSLLQKFE	AAVPSVYPPD	QYARLSI IVT	LESLGIDRDF	KTEIKSILDE
TYRYWLRGDE	EICLDLATCA	LAFRLLLAHG	YDVSYPDKP	FAEESGFSDT	LEGYVKNFTS

VLELFKAAQS YPHESALKKQ CCWTKQYLEM ELSSWVKTSV RDKYLKKEVE DALAFPSYAS  
 LERSDHRRKI LNGSAVENTR VTKTSYRLHN ICTSDILKLA VDDFNFCQSI HREEMERLDR  
 WIVENRLQEL KFARQKLAYC YFSGAATLFS PELSDARISW AKGGVLTTVV DDFFDVGGSK  
 EELENLIHLV EKWDLNGVPE YSSEHVEIIF SVLRDTILET GDKAFTYQGR NVTHHIVKIW  
 LDLLKSMLE AEWSSDKSTP SLEDYMENAY ISFALGPIVL PATYLIGPPL PEKTVDSHQY  
 NQLYKLVSTM GRLLNDIQGF KRESAEGKLN AVSLHMKHER DNRSKEVIIIE SMKGLAERKR  
 EELHKLVL EE KGSVVPRECK EAF LKMSKVL NLFYRKDDGF TSNDLMSLVK SVIYEPVSLQ  
 KESLT

SEQ ID NO: 152

s. rebaudiana KOI

MDAVTGLLTV PATAITIGGT AVALAVALIF WYLKSYTSAR RSQSNHLPRV PEVPGVPLLG  
 NLLQLKEKKP YMTFTRWAAT YGPIYSIKTG ATSMVVSSN EIAKEALVTR FQSISTRNLS  
 KALKVLTADK TMVAMSDYDD YHKTVKRHIL TAVLGPNAQK KHRIHRDIMM DNISTQLHEF  
 VKNNPEQEEV DLRKIFQSEL FGLAMRQALG KDVESLYVED LKITMNRDEI FQVLVVDPM  
 GAIDVDWRDF FPYLKWPVNK KFENTIQQMY IRREAVMKS LKEHKKRIAS GEKLN SYIDY  
 LLSEAQTLTD QQLLMSLWEP IIESSDTTMV TTEWAMYELA KNPQLQDRLY RDIKSVCGSE  
 KITEEHLSQL PYITAIFHET LRRHSPVPII PLRHVHEDTV LGGYHVPAGT ELAVNIYGCN  
 MDKNVWENPE EWNPERFMKE NETIDFQKTM AFGGGKRVCA GSLQALLTAS IGIGRMVQEF  
 EWKLDKMTQE EVNTIGLTTQ MLRPLRAIK PRI

SEQ ID NO: 153

A. thaliana ATR2

MSSSSSSSTS MIDLMAAIK GEPVIVSDPA NA5AYE5VAA ELSSMLIENR QFAMIVTTSI  
 AVLIGCIVML VWRSSGSGNS KRVEPLKPLV IKPREEEIDD GRKKVTIFFG TQTGTAEGFA  
 KALGEEAKAR YEKTRFKIVD LDDYAADDDE YEEKLKKEDV AFFFLATYGD GEPTDNAARF  
 YKWFTEGNDR GEWLKNLKYG VFGLGNRQYE HFNKVAKVVD DILVEQGAQR LVQVGLGDD  
 QCIEDDFTAW REALWPELDT ILREEGDTAV ATPYTAAVLE YRVSIHSDSED AKFNDITLAN  
 GNGYTVFDAQ HPYKANVAVK RELHTPESDR SCIHLEFDIA GSGLTMKLG D HVGVLCDNLS  
 ETVDEALRLL DMSPDYFSL HAEKEDGTP I SSSLPPFP CNLRTALTRY ACLLSSPKKS  
 ALVALAAHAS DPTEAERLKH LASPAGKDEY SKWVVESQRS LLEVMAEFPS AKPPLGVFFA  
 GVAPRLQPRF YSISSSPKIA ETRIHVTCAL VYEKMPGTGRI HKGVCSTWMK NAVPYEKSEK  
 LFLGRPIFVR QSNFKLPSDS KVPIIMIGPG TGLAPFRGFL QERLALVESG VELGPSVLFF  
 GCRNRRMDFI YEEELQRFVE SGALAELSVA FSREGPTKEY VQHKMMDKAS DIWNMISQGA  
 YLYVCGDAKG MARDVHRSLH TIAQEQGSM STKAEGFVKN LQTSGRYL RD VW

SEQ ID NO: 154

Stevia **rebaudiana** KAHel

MEASYLYISI LLLLASYLFT TQLRRKSANL PPTVFPSIPI IGHLYLLKKP LYRTLAKIAA  
 KYGPILQLQL GYRRVLVISS PSAAEECFNT NDVIFANRPK TLF GKIVGGT SLGSLSYGDQ  
 WRNLRRVASI EILSVHRLNE FHDIRVDENR LLIRKLRS S SPVTLITVFY ALTLNVIMRM  
 ISGKRYFDSG DRELEEEGKR FREILDETL LAGASNVGDY LPILNWLGVK SLEKLIALQ  
 KKRDDFFQGL IEQVRKSRGA KVGKGRK TMI ELLLSLQESE PEYYTDAMIR SFVLGLLAAG  
 SDTSAGTMEW AMSLLVNHPH VLKKAQAEID RVIGNNRLID ESDIGNIPYI GCIINETLRL  
 YPAGPLLPFH ESSADCVISG YNIPRGTMLI VNQWAIHHPD KVWDDPETFK PERFQGLEGT  
 RDGFKLMPFG SGRRCPEGEG LAIRLLG MTL GSVIQCDFWE RVGDEMVDMT EGLGVTLPKA  
 VPLVAKCKPR SEMTNLLSEL

SEQ ID NO: 155

Stevia **rebaudiana** CPR8

MQSNSVKISP LDLVTALFSG KVLDTSNASE SGESAMLPTI AMIMENRELL MILTTSVAVL  
 IGCVVVLVWR RSSTKKSAL E PPVIVVPKRV QEEVEVDDGKK KVTVFFGTQT GTAEGFAKAL  
 VEEAKARYEK AVFKVIDLDD YAADDDEYEE KLKKE SLAFF FLATYGDGEP TDNAARFYK W

FTEGCAKGEW LNKLQYGVFG LGNRQYEHFN KIAKVDDGL VEQGAKRLVP VGLGDDDQCI  
 EDDFTAWKEL VWPELDQLLR DEDDTTVATP YTAQVAEYRV VFHEKPDALS EDYSYTNHGA  
 VHDAQHPCRS NVAVKKELHS PESDRSCTHL EFDISNTGLS YETGDHVG VY CENLSEVVND  
 AERLVGLPPD TYSSIHTDSE DGSPLGGASL PPPFPCTLR KALTCYADVL SSPKKSALLA  
 LAAHATDPSE ADRLKFLASP AGKDEYSQWI VASQRSLLLEV MEAFPSAKPS LGVFFASVAP  
 RLQPRYYSIS SSPKMAPDRI HVTICALVYEK TPAGRIHKG V CSTWMKNAVP MTESQDCSWA  
 PIYVRTSNFR LPSPDKVPVI MIGPGTGLAP FRGFLQERLA LKEAGTDLGL SILFFGCRNR  
 KVDFIYENEL NNFVETGALS ELIVAFSREG PTKEYVQH KM SEKASDIWNL LSEGAYLYVC  
 GDAKGMAKDV HRTLHTIVQE QGSLDSSKAE LYVKNLQMSG RYL RDVW

SEQ ID NO: 156

Stevia rebaudiana UGT85C2

MDAMATTEKK PHVIFIPFPA QSHIKAMLKL AQLLHHKGLQ ITFVNTDFIH NQFLESSGPH  
 CLDGAPGFRF ETIPDGVSHS PEASIPİRES LLRSIETNFL DRFIDLVT KL PDPPTCIISD  
 GFLSVFTIDA AKKLGIPVMM YWTLAACGFM GFYHIHSLIE KGFAPLKDAS YLTNGYLDTV  
 IDWVPGMEGI RLKDFPLDWS TDLNDKVL MF TTEAPQRSHK VSHHIFHTFD ELEPSIIKTL  
 SLRYNHIYTI GPLQLLLDQI PEEKKQTGIT SLHGYSLVKE EPECFQWLQS KEPNSVVYVN  
 FGSTTVMSLE DMTEFGWGLA NSNHYFLWI I RSNLVIGENA VLPPELEEH I KKRGFIA SWC  
 SQEKVLKHP S VGGFLTHCGW GSTIESLSAG VPMICWPYSW DQLTNCRYIC KEWEVGLEMG  
 TKVKRDEVKR LVQELMGEGG HKMRNKAKDW KEKARIAIAP NGSSSLNIDK MVKEITVLAR  
 N

SEQ ID NO: 157

S. rebaudiana UGT74G1 (GenBank A ~~LR06920.1~~)

MAEQQKIKKS PHVLLIPFPL QGHINPFIQF GKRLISKGVK TTLVTTIHTL NSTLNHSNTT  
 TTSIEIQAIS DGCDEGGFMS AGESYLETFK QVGSKSLADL IKKLQSEGTT IDAIIYDSMT  
 EWVLDVAIEF GIDGGSFFTQ ACVVNSLYYH VHKGLISLPL GETVSVPGFP VLQRWETPLI  
 LQNHQIQSP WSQMLFGQFA NIDQARWVFT NSFYKLEEEV LEWTRKIWNL KVIGPTLPSM  
 YLDKRLDDDK DNGFNLYKAN HHECMNWLDD KP KESVVYVA FGSLVKHGPE QVEEITRALI  
 DSDVNFLWVI KHKEEGKLPE NLSEVIKTGK GLIVAWCKQL DVLAHESVGC FVTHCGFNST  
 LEAISLGV PV VAMPQFSDQT TNAKLLDEIL GVGVRKADE NGIVRRGNLA SCIKMIMEEE  
 RGV IIRKNAV KWKDLAKVAV HEGGSSDNDI VEFVSELIKA

SEQ ID NO: 158

S. rebaudiana UGT76G1

MENKTETTVR RRRRIILFPV PFQGHINPIL QLANVLYSKG FSITIFHTNF NKPKTSNYPH  
 FTFRFILDND PQDERISNLP THGPLAGMRI PIINEHGADE LRRELELLML ASEDEEVSC  
 LITDALWYFA QSVADSLNLR RLVLMTSSLF NFHAHVSLPQ FDELGYLDPD DKTRLEEQAS  
 GFPM LKVKDI KSAYSNWQIL KEILGKMIKQ TKASSGVIWN SFKELEESEL ETVIREIPAP  
 SFLIPLPKHL TASSSSLLDH DRTVFQWLDQ QPPSSVLYVS FGSTSEVDEK DFLEIARGLV  
 DSKQSFLWVV RPGFVKGSTW VEPLPDGFLG ERGRIVKWVP QQEVLAHGAI GAFWTHSGWN  
 STLESVCEGV PMIFSDFGLD QPLNARYMSD VLKGVYLEN GWERGEIANA IRRVMVDEEG  
 EYIRQARVL KQKADVSLMK GGSSYESLES LVSYISSL

SEQ ID NO: 159

S. rebaudiana UGI91D2e-b

MATSDSIVDD RKQLHVATFP WLAFGHILPY LQLSKLIAEK GHKVSFLSTT RNIQRLSSHI  
 SPLINVVQLT LPRVQELPED AEATTDVHPE DIPYLK KASD GLQPEVTRFL EQHSPDWI IY  
 DYTHYWLPSI AASLGISR AH FSVTTPWAIA YMGPSADAMI NGSDGRTTVE DLTPPKWFP  
 FPTKVCWRKH DLARLV PYKA PGISDGYRMG MVLKGS DCLL SKCYHEFGTQ WLPLLET LHQ  
 VPVVPVGLLP PEIPGDEKDE TWVSIKKWLD GKQKGSVVYV ALGSEALVSQ TEVVELALGL  
 ELSGLPFVWA YRKPKGPAKS DSVELPDGFV ERTRDRGLVW TSWAPQLRIL SHESVCGFLT  
 HCGSGSIVEG LMFHGHLIML PIFGDQPLMA RLLEDKQVGI EIPRNEEDGC LTKESVARSL

RSYWEKEGE IYKANARELS KIYNDTKVEK EYVSQFVDYL **EKNARAVai D** HES

SEQ ID NO: 160

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atggctacct tgttgaaca ttttcaagct atgccattcg ctattccaat tgcttt ggct
gctttgtctt ggttgttttt gttctacatc aaggtttctt tcttctccaa caaatccgct
caagctaaat tgccaccagt tccagttggt ccaggtttgc cagttattgg taatttggtg
caattgaaag aaaagaagcc ataccaaac ttcactagat gggctgaaga atatggtcca
atctactcta ttagaactgg tgcttctact atggttgtct tgaacactac tcaagttgcc
aaagaagcta tggttaccag atacttgtct atctctacca gaaagttgtc caacgccttg
aaaattttga cgcgtgataa gtgcatgggt gccatttctg attacaacga tttccacaag
atgatcaaga gatatatctt gtctaacggt ttgggtccat ctgccccaaa aagacataga
tctaacagag ataccttgag agccaacggt tgttctagat tgcattccca agttaagaac
tctccaagag aagctgtcaa ctttagaaga gttttcgaat ggaattatt cggtatcgct
ttgaaacaag ccttcggtaa ggatattgaa aagccaatct acgtcgaaga attgggtact
actttgtcca gagatgaaat cttcaagggt ttgggtcttg acattatgga aggtgccatt
gaagttgatt ggagagattt tttccatac ttgcgttga ttccaacac cagaatggaa
actaagatcc aaagattata ctttagaaga aaggcgtta tgaccgcctt gattaacgaa
caaaagaaaa gaattgcctc cggtgaaaga atcaactgct acatcgattt cttggtgaaa
gaaggtaga ccttgaccat ggaccaaac tctatggtgt tgtgggaaac cgttattgaa
actgctgata ccacaatggt tactactgaa tgggctatgt acgaagttgc taaggattct
aaaagacaag acagattata ccaagaaatc caaaaggtct gcggttctga aatggttaca
gaagaatact tgtcccaatt gccatacttg aatgctgttt tccacgaaac tttgagaaaa
cattctccag ctgctttggt tccattgaga tatgctcatg aagatactca attgggtggt
tattacattc cagccggtac tgaaattgcc attaacatct acggttgcaa catggacaaa
caccaatggg aatctccaga agaatggaag ccagaaagat ttttggatcc taagtttgac
ccaatggact tgtacaaaac tatggctttt ggtgctggta aaagagtttg cgctggttct
ttacaagcta tgttgattgc ttgtccaacc atcggtagat tggttcaaga atttgaatgg
aagttgagag atggtgaaga agaaaacggt gatactgttg gtttgaccac ccataagaga
tatccaatgc atgctatttt gaagccaaga tcttaa
    
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SEQ ID NO: 161

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MATLLEHFQA MPFAIPIALA ALSWLFLFYI KVSFFSNKSA QAKLPPVPV PGLPVIGNLL
QLKEKKPYQT FTRWAEYGP IYSIRTGAST MVVLNTTQVA KEAMVTRYLS ISTRKLSNAL
KILTADKCMV AISDYNDFHK MIKRYILSNV LGPSAQKRHR SNRDTLRANV CSRLHSQVKN
SPREAVNFRR VFEWELFGIA LKQAFGKDIE KPI YVEELGT TLSRDEIFKV LVLDIMEGAI
EVDWRDFFPY LRWIPNTRME TKIQRLYFRR KAVMTALINE QKKRIASGEE INCYIDFLLK
EGKTLTMDQI SMLLWETVIE TADTTMVTTE WAMYEVAKDS KRQDRLYQEI QKVCSEMVT
EEYLSQLPYL NAVFHETLRK HSPAALVPLR YAHEDTQLGG YYIPAGTEIA INIYGCNMDK
HQWESPEEWK PERFLDPKFD PMDLYKTMAF GAGKRVCAGS LQAMLIACPT IGRLVQEFEW
KLRDGEENV DTVGLTTHKR YPMHAILKPR S
    
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**[00164]** Having described the invention in detail and by reference to specific embodiments thereof, it will be apparent that modifications and variations are possible without departing from the scope of the invention defined in the appended claims. More specifically, although some aspects of the present invention are identified herein as particularly advantageous, it is contemplated that the present invention is not necessarily limited to these particular aspects of the invention.

**WHAT IS CLAIMED IS:**

1. A recombinant host capable of synthesizing a steviol glycoside, comprising a gene encoding a transporter polypeptide and/or a gene encoding a transcription factor polypeptide that regulates expression of at least one transporter gene;  

wherein expression of the gene encoding the transporter polypeptide and/or the gene encoding the transcription factor polypeptide that regulates expression of at least one transporter gene is modified and the recombinant host transports at least a portion of the synthesized steviol glycoside from the host into a culture medium.
2. The recombinant host of claim 1, wherein the gene encoding the transporter polypeptide is an endogenous gene.
3. The recombinant host of any one of claims 1 or 2, wherein the transporter polypeptide comprises an ATP-binding cassette (ABC) transporter, a major facilitator superfamily (MFS) transporter, an amino acid/auxin permease (AAP) family transporter, an ATPase transporter, a sulfate permease (SuIP) family transporter, a lysosomal cystine transporter (LCT) family transporter, a Ca<sup>2+</sup>:cation antiporter (CaCA) family transporter, an amino acid-polyamine-organocation (APC) superfamily transporter, a multidrug/oligosaccharidyl-lipid/polysaccharide (MOP) transporter, a ZRT/IRT-like protein (ZIP) metal transporter family transporter, a mitochondrial protein translocase (MPT) family transporter, a voltage-gated ion channel (VIC) family transporter, a monovalent cation:proton antiporter-2 (CPA2) family transporter, a ThrE family of putative transmembrane amino acid efflux transporter, an oligopeptide transporter (OPT) family transporter, a K<sup>+</sup> transporter (Trk) family transporter, a bile acid:Na symporter (BASS) family transporter, a drug/metabolite transporter (DMT) superfamily transporter, a mitochondrial carrier (MC) family transporter, an auxin efflux carrier (AEC) family transporter, an ammonia channel transporter (Ami) family transporter, a metal ion (Mn<sup>2+</sup>-iron) transporter (Nramp) family transporter, a transient receptor potential Ca<sup>2+</sup> channel (TRP-CC) family transporter, an arsenical resistance-3 (ACR3) family transporter, a nucleobase:cation symporter-1 (NCS1) family transporter, an inorganic phosphate transporter (PiT) family transporter, an arsenite-antimonite (ArsAB) efflux family transporter, an MSP family of transporter, a glycerol uptake (GUP) family transporter, a metal ion transport

(MIT) family transporter, a copper transport (Ctr) family transporter, or a cation diffusion facilitator (CDF) family transporter.

4. The recombinant host of any one of claims 1-3, wherein modified expression comprises:
  - (a) overexpressing the gene encoding the transporter polypeptide and/or the gene encoding the transcription factor polypeptide; or
  - (b) deleting the gene encoding the transporter polypeptide and/or the gene encoding the transcription factor polypeptide.
5. The recombinant host of any one of claims 1-4, wherein the gene encoding the transporter polypeptide and/or the gene encoding the transcription factor polypeptide has an activity that is increased.
6. The recombinant host of any one of claims 1-5, wherein one or more of the genes encoding the transporter polypeptide and/or one or more of the gene encoding the transcription factor polypeptide is overexpressed.
7. The recombinant host of claim 6, wherein the transporter polypeptide and/or transcription polypeptide comprise YAL067C set forth in SEQ ID NO:14, YBL089W set forth in SEQ ID NO:15, YBL099W set forth in SEQ ID NO:16, YBR008C set forth in SEQ ID NO:86, YBR021W set forth in SEQ ID NO:87, YBR043C set forth in SEQ ID NO:88, YBR180W set forth in SEQ ID NO:13, YBR241C set forth in SEQ ID NO:17, YBR287W set forth in SEQ ID NO:89, YBR294W set forth in SEQ ID NO:18, YBR295W set forth in SEQ ID NO:90, YBR296C set forth in SEQ ID NO:91, YCL038C set forth in SEQ ID NO:92, YCL069W set forth in SEQ ID NO:19, YCR011C set forth in SEQ ID NO:93, YCR028C set forth in SEQ ID NO:20, YCR075C set forth in SEQ ID NO:21, YDL054C set forth in SEQ ID NO:94, YDL100C set forth in SEQ ID NO:95, YDL128W set forth in SEQ ID NO:22, YDL185W set forth in SEQ ID NO:23, YDL194W set forth in SEQ ID NO:24, YDL210W set forth in SEQ ID NO:25, YDL245C set forth in SEQ ID NO:96, YDL247W set forth in SEQ ID NO:97, YDR011W set forth in SEQ ID NO:98,

YDR061W set forth in SEQ ID NO:26, YDR093W set forth in SEQ ID NO:27,  
YDR292C set forth in SEQ ID NO:99, YDR338C set forth in SEQ ID NO:28,  
YDR406W set forth in SEQ ID NO:29, YDR497C set forth in SEQ ID NO:100,  
YDR536W set forth in SEQ ID NO:30, YEL006W set forth in SEQ ID NO:101,  
YEL027W set forth in SEQ ID NO:102, YEL031W set forth in SEQ ID NO:31,  
YEL065W set forth in SEQ ID NO:103, YER019C-A set forth in SEQ ID NO:104,  
YER053C set forth in SEQ ID NO:105, YER119C set forth in SEQ ID NO:106,  
YER166W set forth in SEQ ID NO:32, YFL011W set forth in SEQ ID NO:33,  
YFL028C set forth in SEQ ID NO:107, YFR045W set forth in SEQ ID NO:108,  
YGL006W set forth in SEQ ID NO:34, YGL013C set forth in SEQ ID NO:35,  
YGL084C set forth in SEQ ID NO:109, YGL104C set forth in SEQ ID NO:110,  
YGL114W set forth in SEQ ID NO:111, YGL167C set forth in SEQ ID NO:112,  
YGL255W set forth in SEQ ID NO:36, YGR125W set forth in SEQ ID NO:37,  
YGR181W set forth in SEQ ID NO:38, YGR217W set forth in SEQ ID NO:39,  
YGR224W set forth in SEQ ID NO:40, YGR257C set forth in SEQ ID NO:113,  
YGR281W set forth in SEQ ID NO:41, YHL016C set forth in SEQ ID NO:42,  
YHL035C set forth in SEQ ID NO:114, YHL036W set forth in SEQ ID NO:115,  
YHR002W set forth in SEQ ID NO:116, YHR096C set forth in SEQ ID NO:117,  
YIL006W set forth in SEQ ID NO:118, YIL088C set forth in SEQ ID NO:43, YIL120W  
set forth in SEQ ID NO:119, YIL121W set forth in SEQ ID NO:120, YIL166C set forth  
in SEQ ID NO:121, YJL093C set forth in SEQ ID NO:44, YJL094C set forth in SEQ  
ID NO:45, YJL108C set forth in SEQ ID NO:46, YJL133W set forth in SEQ ID  
NO:122, YJL212C set forth in SEQ ID NO:47, YJL219W set forth in SEQ ID NO:123,  
YJR106W set forth in SEQ ID NO:48, YJR160C set forth in SEQ ID NO:49, YKL016C  
set forth in SEQ ID NO:124, YKL050C set forth in SEQ ID NO:125, YKL064W set  
forth in SEQ ID NO:50, YKL120W set forth in SEQ ID NO:126, YKL146W set forth in  
SEQ ID NO:127, YKL209C set forth in SEQ ID NO:128, YKR039W set forth in SEQ  
ID NO:129, YKR050W set forth in SEQ ID NO:51, YKR105C set forth in SEQ ID  
NO:52, YKR106W set forth in SEQ ID NO:53, YLR411W set forth in SEQ ID NO:130,  
YLR447C set forth in SEQ ID NO:54, YML038C set forth in SEQ ID NO:131,  
YML116W set forth in SEQ ID NO:55, YMR034C set forth in SEQ ID NO:56,  
YMR056C set forth in SEQ ID NO:57, YMR166C set forth in SEQ ID NO:132,  
YMR253C set forth in SEQ ID NO:58, YMR279C set forth in SEQ ID NO:133,  
YNL003C set forth in SEQ ID NO:134, YNL065W set forth in SEQ ID NO:59,  
YNL070W set forth in SEQ ID NO:60, YNL083W set forth in SEQ ID NO:61,  
YNL095C set forth in SEQ ID NO:62, YNL121C set forth in SEQ ID NO:63,  
YNL142W set forth in SEQ ID NO:64, YNL268W set forth in SEQ ID NO:135,

YNR055C set forth in SEQ ID NO:136, YOL020W set forth in SEQ ID NO:65, YOL075C set forth in SEQ ID NO:66, YOL077W-A set forth in SEQ ID NO:67, YOL122C set forth in SEQ ID NO:68, YOL158C set forth in SEQ ID NO:137, YOR079C set forth in SEQ ID NO:69, YOR087W set forth in SEQ ID NO:70, YOR092W set forth in SEQ ID NO:71, YOR100C set forth in SEQ ID NO:138, YOR130C set forth in SEQ ID NO:72, YOR153W set forth in SEQ ID NO:139, YOR222W set forth in SEQ ID NO:73, YOR271C set forth in SEQ ID NO:140, YOR273C set forth in SEQ ID NO:141, YOR291W set forth in SEQ ID NO:74, YOR306C set forth in SEQ ID NO:75, YOR307C set forth in SEQ ID NO:142, YOR316C set forth in SEQ ID NO:76, YOR332W set forth in SEQ ID NO:143, YOR334W set forth in SEQ ID NO:77, YOR348C set forth in SEQ ID NO:144, YPL036W set forth in SEQ ID NO:145, YPL078C set forth in SEQ ID NO:78, YPL270W set forth in SEQ ID NO:79, YPL274W set forth in SEQ ID NO:80, YPR003C set forth in SEQ ID NO:81, YPR01 1C set forth in SEQ ID NO:82, YPR058W set forth in SEQ ID NO:83, YPR128C set forth in SEQ ID NO:84, or YPR201W set forth in SEQ ID NO:85.

8. The recombinant host of any one of claims 1-7, wherein YBR043C set forth in SEQ ID NO:88, YDL100C set forth in SEQ ID NO:95, YDL054C set forth in SEQ ID NO:94, YDL128W set forth in SEQ ID NO:22, YDL198C set forth in SEQ ID NO:146, YDR061W set forth in SEQ ID NO:26, YDR536W set forth in SEQ ID NO:30, YEL027W set forth in SEQ ID NO:102, YFL054C set forth in SEQ ID NO:147, YGL167C set forth in SEQ ID NO:112, YGR181W set forth in SEQ ID NO:38, YHL016C set forth in SEQ ID NO:42, YIL166C set forth in SEQ ID NO:121, YJL093C set forth in SEQ ID NO:44, YJR106W set forth in SEQ ID NO:48, YKL120W set forth in SEQ ID NO:126, YKL146W set forth in SEQ ID NO:127, YKR039W set forth in SEQ ID NO:129, YMR034C set forth in SEQ ID NO:56, YMR166C set forth in SEQ ID NO:132, YOL122C set forth in SEQ ID NO:68, YOR079C set forth in SEQ ID NO:69, YPL270W set forth in SEQ ID NO:79, and/or YPR01 1C set forth in SEQ ID NO:82 are overexpressed.
9. The recombinant host of any one of claims 1-8, further comprising:
  - (a) one or more genes encoding a sucrose transporter and a sucrose synthase;



- (b) a gene encoding a geranylgeranyl diphosphate synthase (GGPPS) polypeptide;
- (c) a gene encoding an ent-copalyl diphosphate synthase (CDPS) polypeptide;
- (d) a gene encoding a kaurene synthase (KS) polypeptide;
- (e) a gene encoding a kaurene oxidase (KO) polypeptide;
- (f) a gene encoding a steviol synthase (KAH) polypeptide;
- (g) a gene encoding a cytochrome P450 reductase (CPR) polypeptide;
- (h) a gene encoding a UGT85C2 polypeptide;
- (i) a gene encoding a UGT76G1 polypeptide;
- o) a gene encoding a UGT74G1 polypeptide;
- (k) a gene encoding a UGT91D2 functional homolog; and/or
- (l) a gene encoding a EUGT1 1 polypeptide;

wherein at least one of the genes is a recombinant gene; and

wherein the host is capable of producing one or more of RebA, RebB, RebD and/or RebM.

- 10. The recombinant host of claim 9, wherein at least one of the genes is codon optimized for expression in the host.
- 11. The recombinant host of claim 10, wherein at least one of the genes is codon optimized for expression in *Saccharomyces cerevisiae*.
- 12. The recombinant host of claim 9, wherein:
  - (a) the GGPPS polypeptide comprises a polypeptide having at least 70% identity to an amino acid sequence set forth in SEQ ID NO: 149;
  - (b) the CDPS polypeptide comprises a polypeptide having at least 70% identity to an amino acid sequence set forth in SEQ ID NO:150;

- (c) the KO polypeptide comprises a polypeptide having at least 70% identity to an amino acid sequence set forth in SEQ ID NO: 152;
  - (d) the KS polypeptide comprises a polypeptide having at least 40% identity to an amino acid sequence set forth in SEQ ID NO: 151;
  - (e) the KAH polypeptide comprises a polypeptide having at least 60% identity to an amino acid sequence set forth in SEQ ID NO: 154;
  - (f) the CPR polypeptide comprises a polypeptide having at least 70% identity to an amino acid sequence set forth in SEQ ID NO: 153 and/or a polypeptide having at least 65% identity to an amino acid sequence set forth in SEQ ID NO: 155;
  - (g) the UGT85C2 polypeptide comprises a polypeptide having at least 55% identity to an amino acid sequence set forth in SEQ ID NO: 156;
  - (h) the UGT76G1 polypeptide comprises a polypeptide having at least 50% identity to an amino acid sequence set forth in SEQ ID NO: 158;
  - (i) the UGT74G1 polypeptide comprises a polypeptide having at least 55% identity to an amino acid sequence set forth in SEQ ID NO: 157;
  - (j) the a UGT91 D2 functional homolog comprises a UGT91D2e-b polypeptide having at least 90% identity to the amino acid sequence set forth in SEQ ID NO: 159; and
  - (k) the EUGT1 1 polypeptide comprises a polypeptide having at least 65% identity to an amino acid sequence set forth in SEQ ID NO: 148.
13. The recombinant host of any one of claims 1-12, wherein the recombinant host comprises a microorganism that is a plant cell, a mammalian cell, an insect cell, a fungal cell, or a bacterial cell.
14. The recombinant host of claim 13, wherein the bacterial cell comprises *Escherichia* bacteria cells, *Lactobacillus* bacteria cells, *Lactococcus* bacteria cells, *Cornibacterium* bacteria cells, *Acetobacter* bacteria cells, *Acinetobacter* bacteria cells, or *Pseudomonas* bacterial cells.
15. The recombinant host of claim 13, wherein the fungal cell is a yeast cell.

16. The recombinant host of claim 15, wherein the yeast cell is a cell from *Saccharomyces cerevisiae*, *Schizosaccharomyces pombe*, *Yarrowia lipolytica*, *Candida glabrata*, *Ashbya gossypii*, *Cyberlindnera jadinii*, *Pichia pastohs*, *Kluyveromyces lactis*, *Hansenula polymorpha*, *Candida boidinii*, *Arxula adenivorans*, *Xanthophyllomyces dendrorhous*, or *Candida albicans* species.
17. The recombinant host of claim 16, wherein the yeast cell is a *Saccharomycete*.
18. The recombinant host of claim 17, wherein the yeast cell is a cell from the *Saccharomyces cerevisiae* species.
19. A method of producing a steviol glycoside, comprising:
  - (a) growing the recombinant host of any one of claims 1-18 in a culture medium, under conditions in which the genes discussed in any one of claims 1 to 18 are expressed,  
wherein the steviol glycoside is synthesized by the host; and
  - (b) optionally isolating the steviol glycoside.
20. The method of claim 19, wherein the steviol glycoside is RebA, RebB, RebD, and/or RebM, and wherein:
  - (a) RebA is capable of being synthesized in the recombinant host of any one of claims 1-18 expressing UGT85C2, UGT76G1, UGT74G1, and UGT91D2;
  - (b) RebB is capable of being synthesized in the recombinant host of any one of claims 1-18 expressing UGT85C2, UGT76G1, and UGT91D2;
  - (c) RebD is capable of being synthesized in the recombinant host of any one of claims 1-18 expressing UGT85C2, UGT76G1, UGT74G1, and UGT91D2 and/or EUGT1 1; and
  - (d) RebM is capable of being synthesized in the recombinant host of any one of claims 1-18 expressing UGT85C2, UGT76G1, UGT74G1, and UGT91D2 and/or EUGT1 1.

21. The method of claims 19 or 20, wherein a gene encoding YBR043C set forth in SEQ ID NO:88, YDL100C set forth in SEQ ID NO:95, YDL054C set forth in SEQ ID NO:94, YDL128W set forth in SEQ ID NO:22, YDL198C set forth in SEQ ID NO:146, YDR061W set forth in SEQ ID NO:26, YDR536W set forth in SEQ ID NO:30, YEL027W set forth in SEQ ID NO:102, YFL054C set forth in SEQ ID NO:147, YGL167C set forth in SEQ ID NO:112, YGR181W set forth in SEQ ID NO:38, YHL016C set forth in SEQ ID NO:42, YIL166C set forth in SEQ ID NO:121, YJL093C set forth in SEQ ID NO:44, YJR106W set forth in SEQ ID NO:48, YKL120W set forth in SEQ ID NO:126, YKL146W set forth in SEQ ID NO:127, YKR039W set forth in SEQ ID NO:129, YMR034C set forth in SEQ ID NO:56, YMR166C set forth in SEQ ID NO:132, YOL122C set forth in SEQ ID NO:68, YOR079C set forth in SEQ ID NO:69, YPL270W set forth in SEQ ID NO:79, and/or YPR011C set forth in SEQ ID NO:82 is overexpressed.
22. The method of any one of claims 19-21, wherein the steviol glycoside is produced at a concentration of between about 500 mg/L to about 10,000 mg/L.
23. A method of increasing production or transport of a steviol glycoside into a culture medium, comprising:
- (a) growing the recombinant host of any one of claims 1-18 in a culture medium, under conditions in which the genes discussed in any one of claims 1 to 18 are expressed, wherein the steviol glycoside is synthesized by the host; and
  - (b) optionally isolating the steviol glycoside.
24. The method of claim 23, wherein the steviol glycoside is RebA, RebB, RebD, and/or RebM.

Figure 1

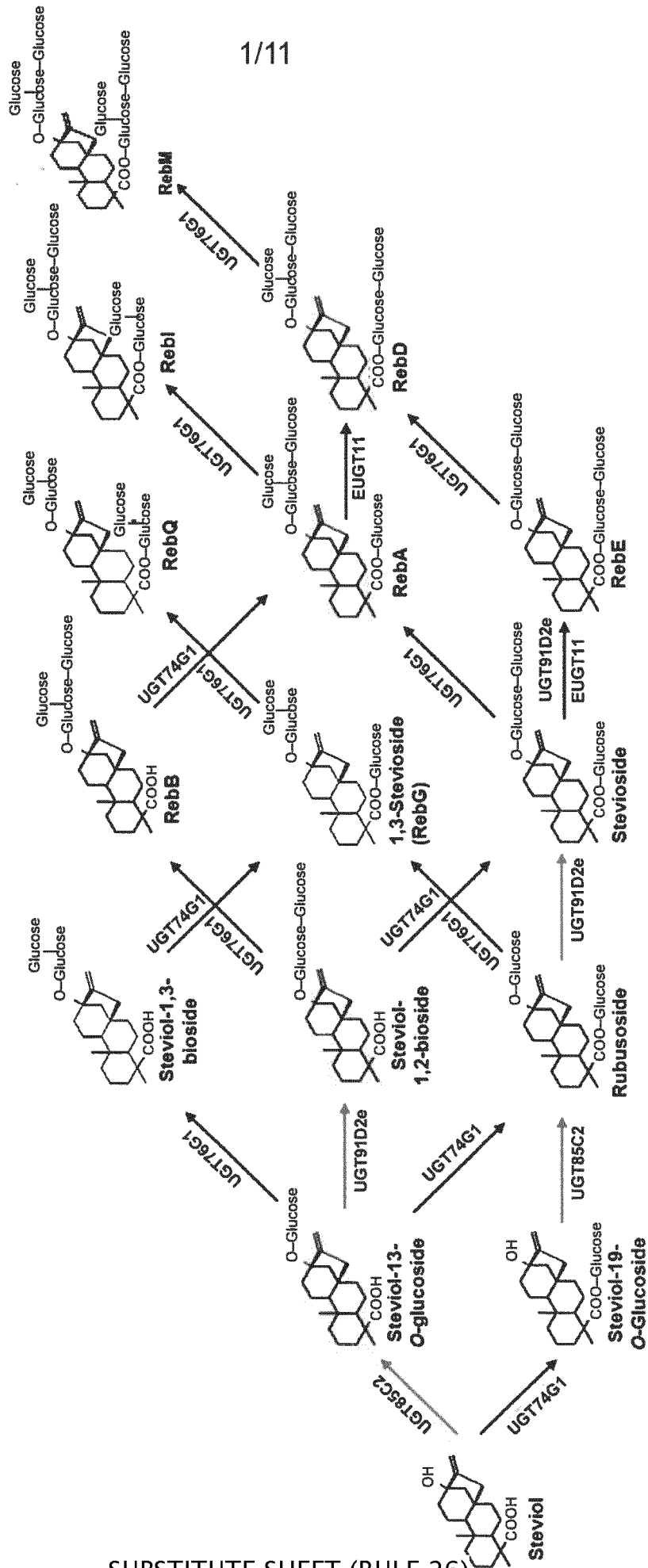
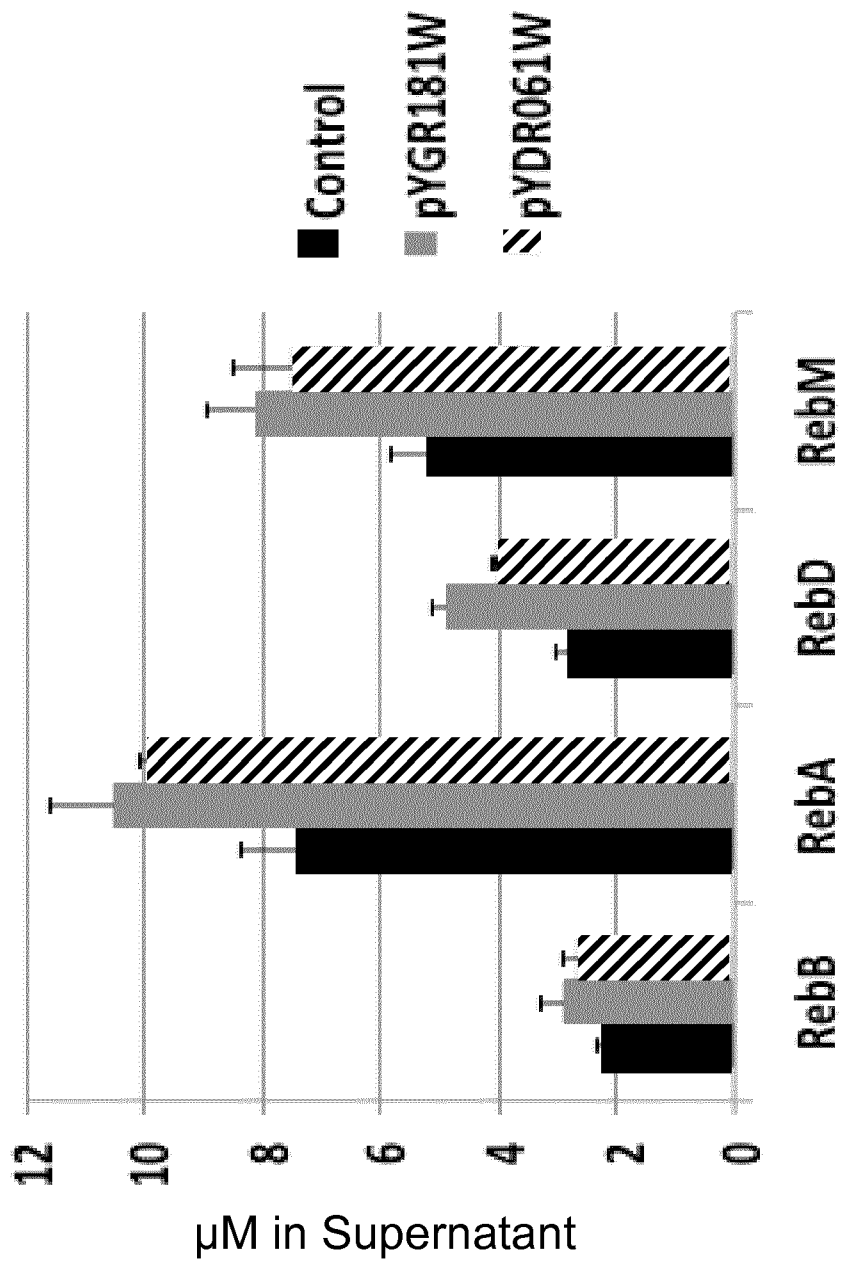
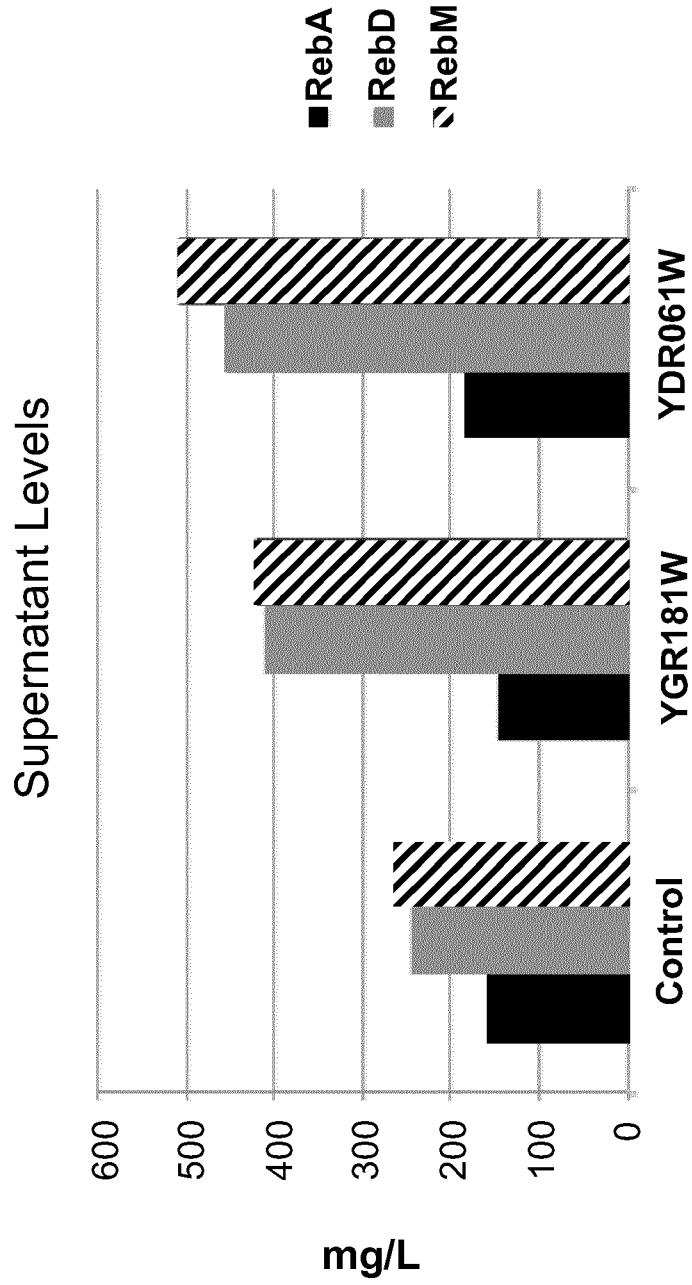


Figure 2



3/11

Figure 3A



4/11

Figure 3B

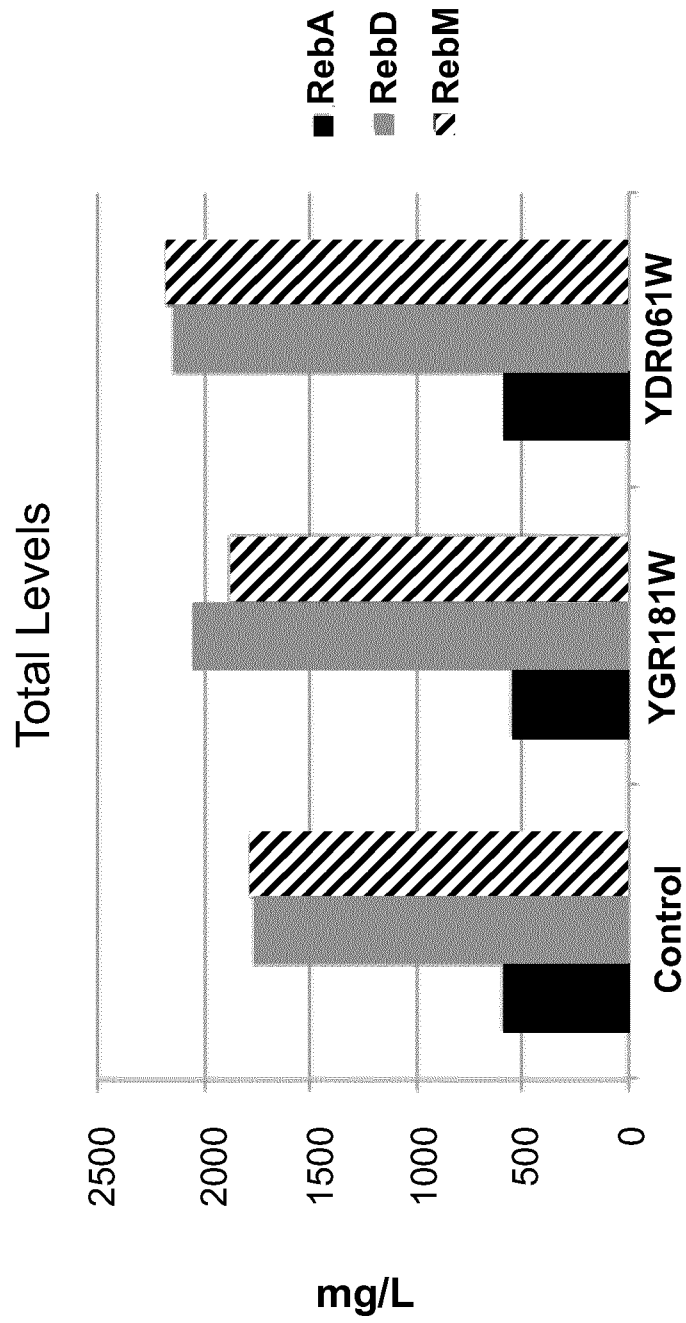
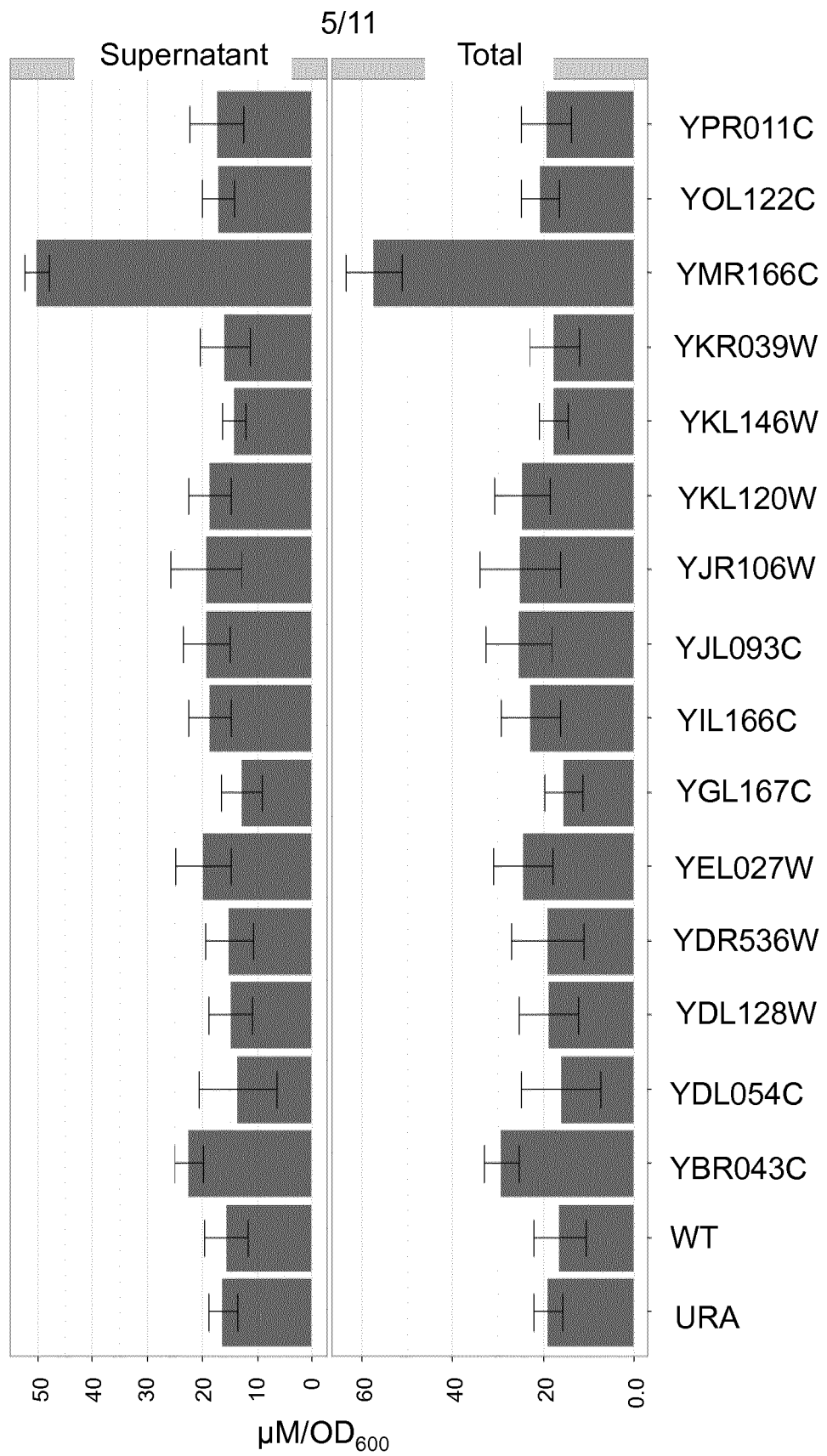
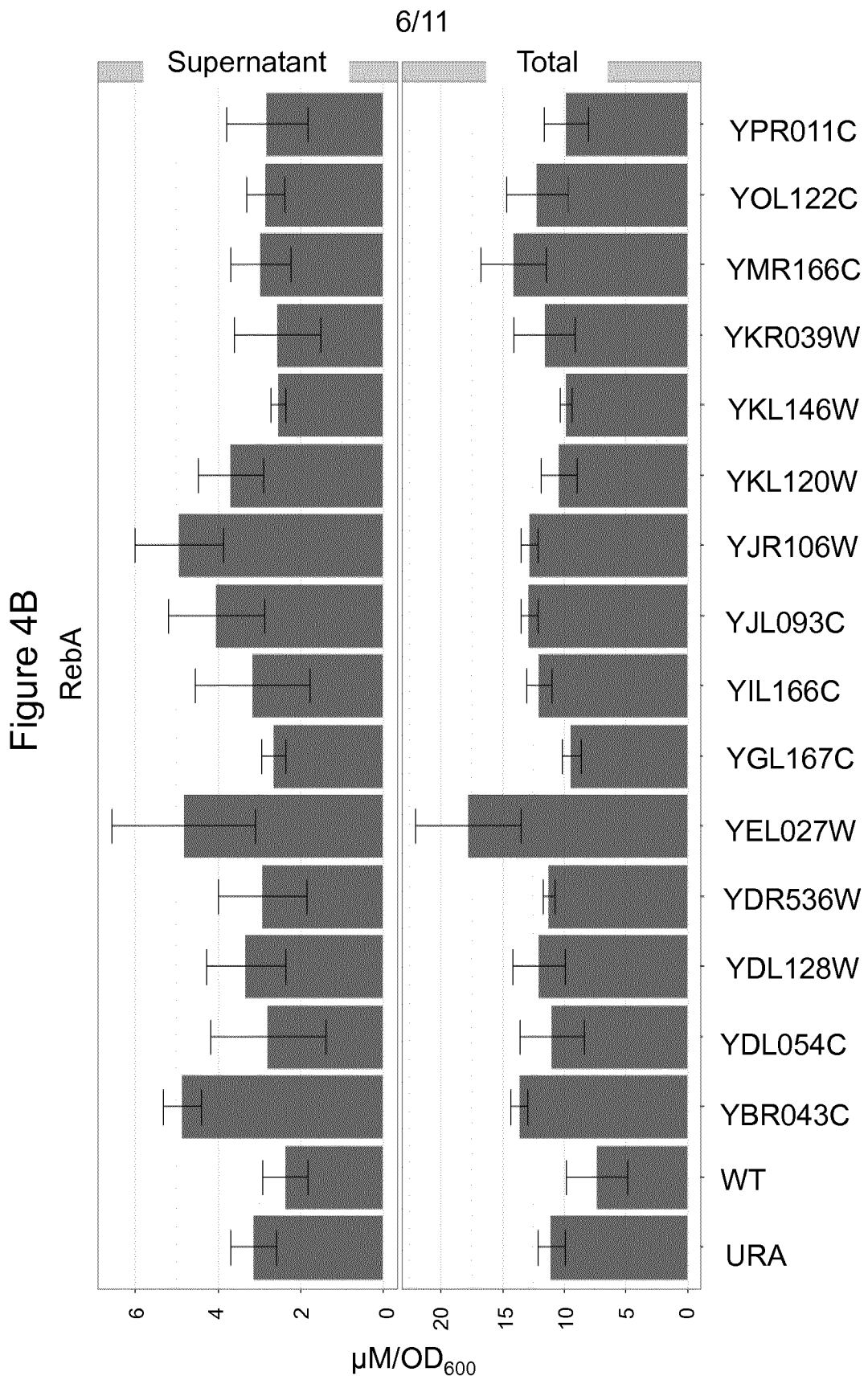
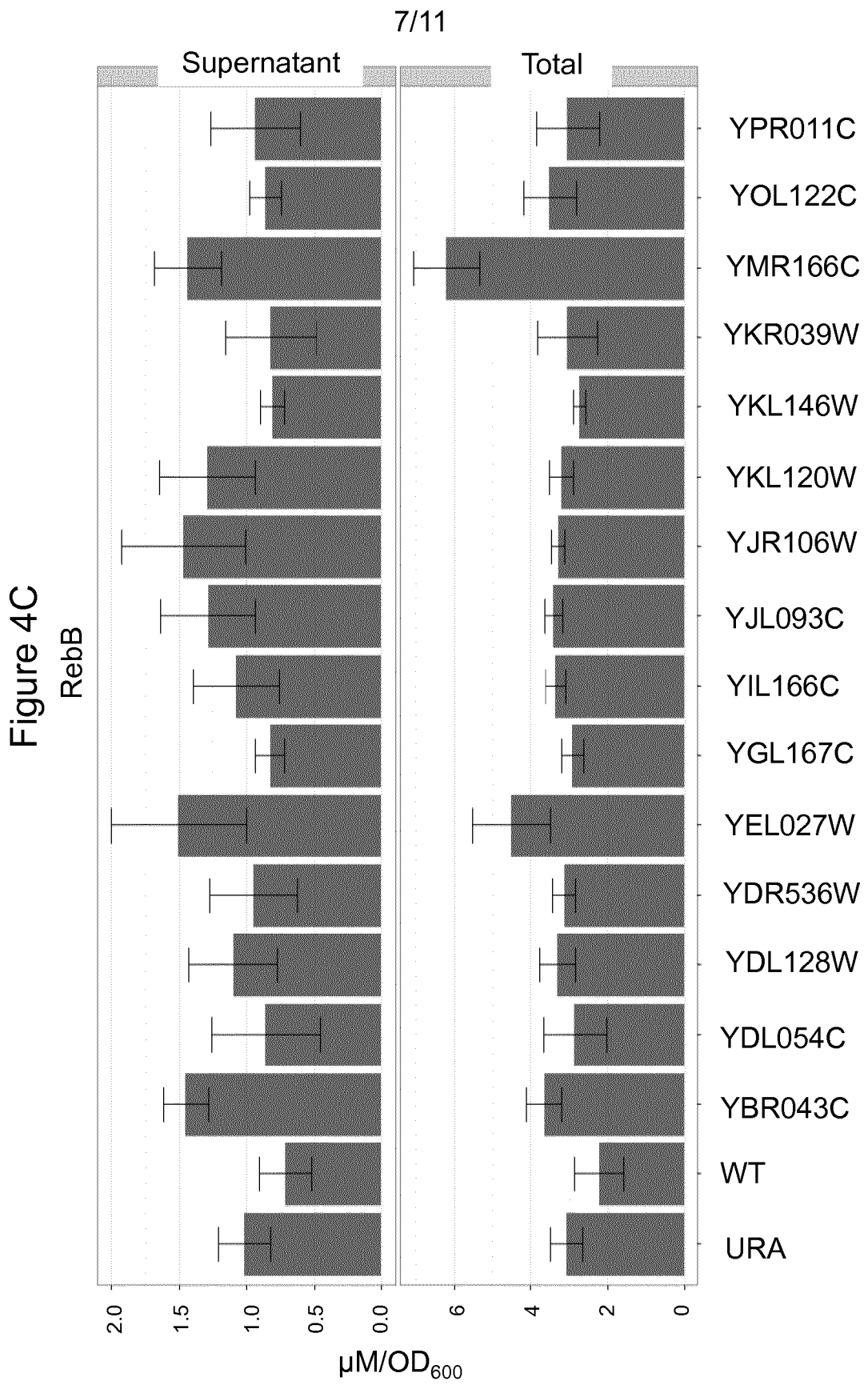


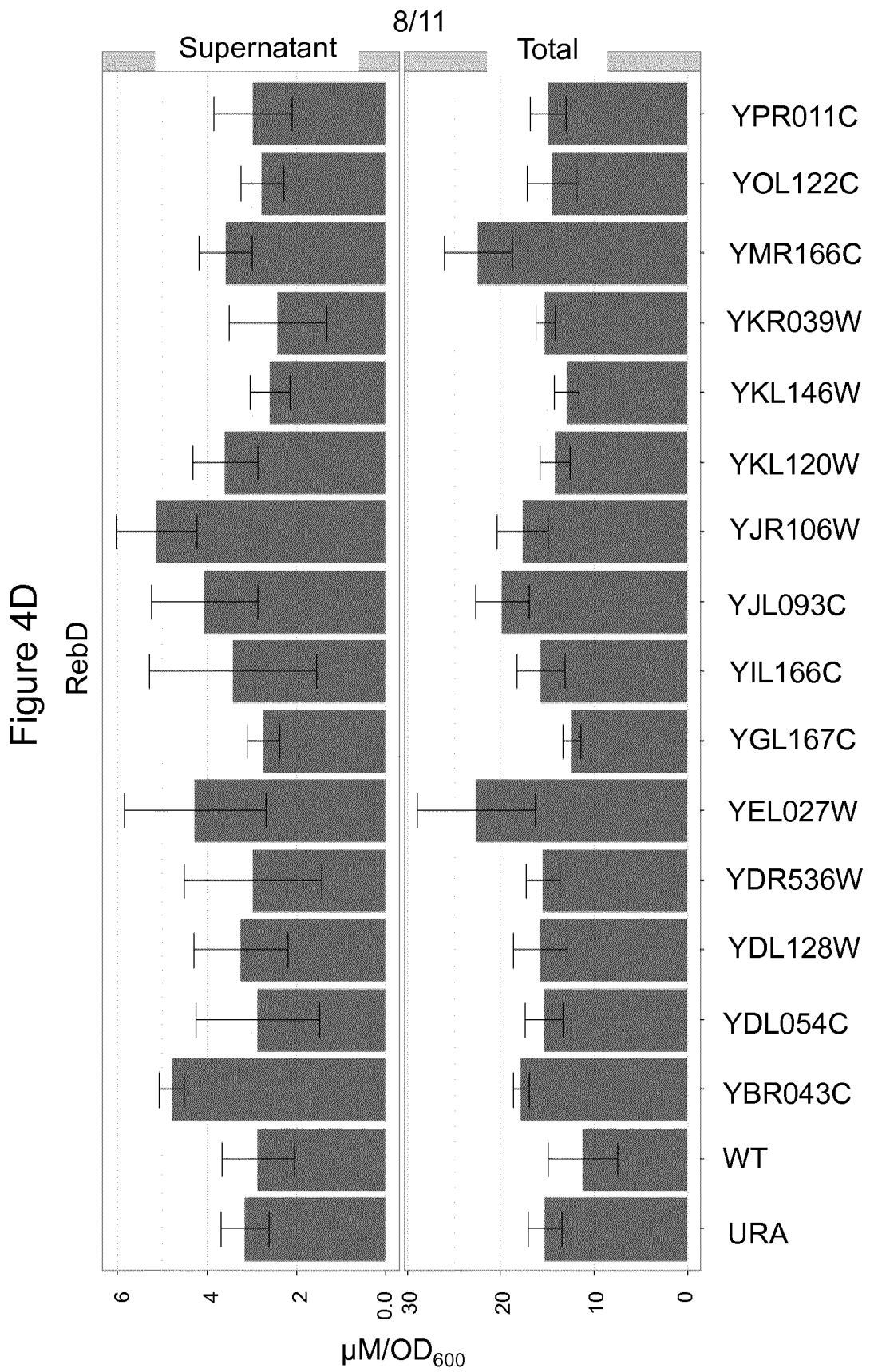


Figure 4A  
13-SMG









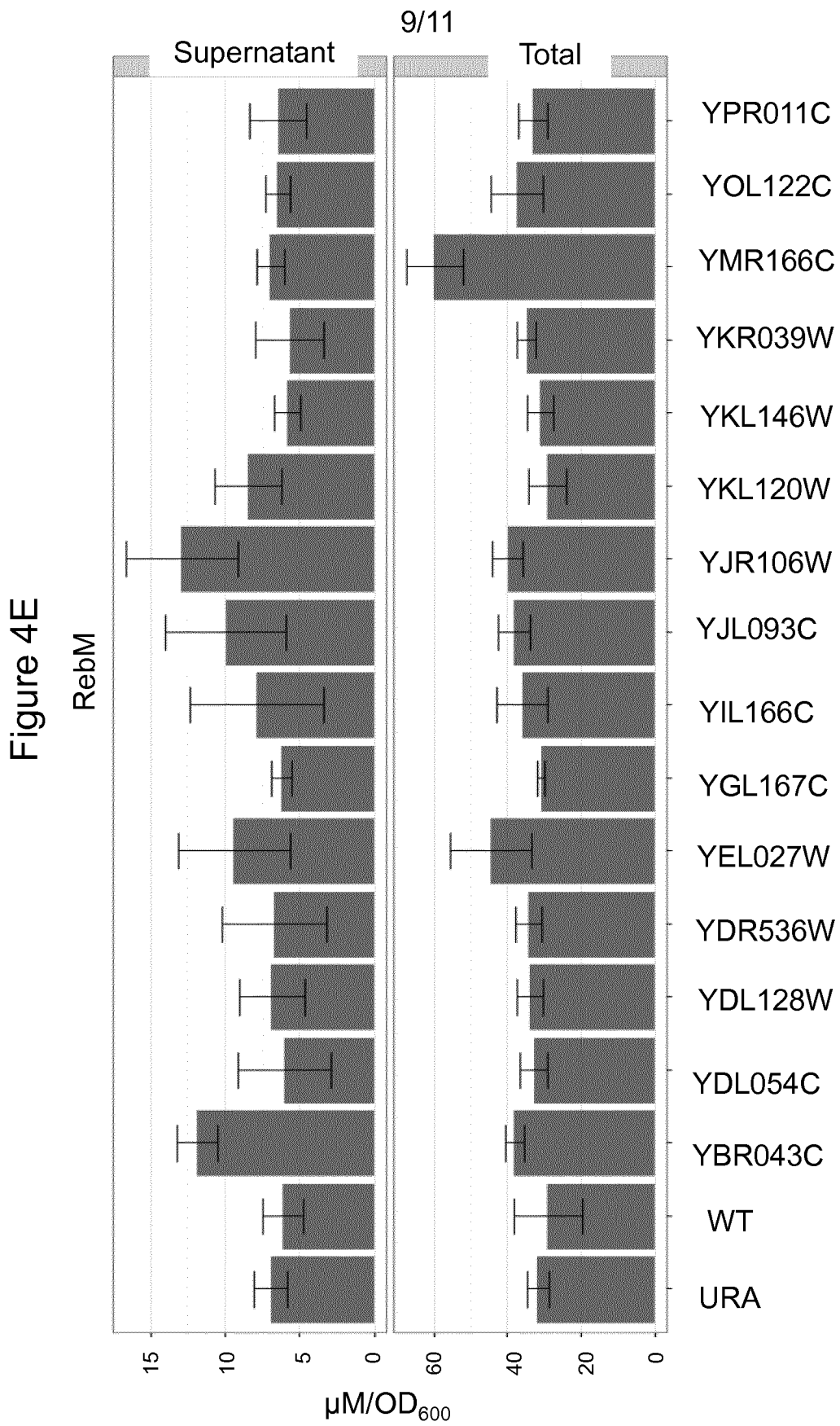


Figure 4E

Figure 5A

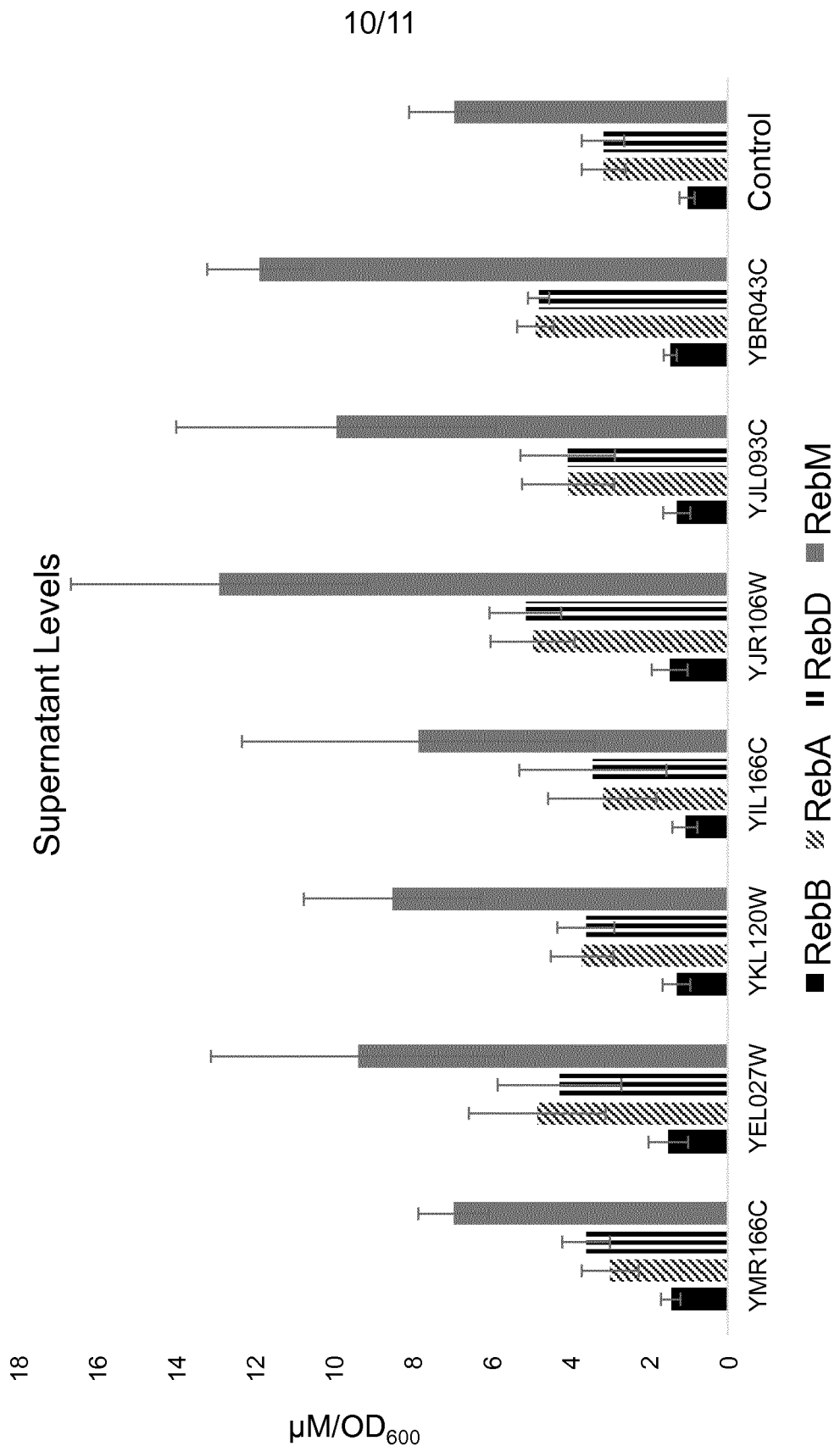
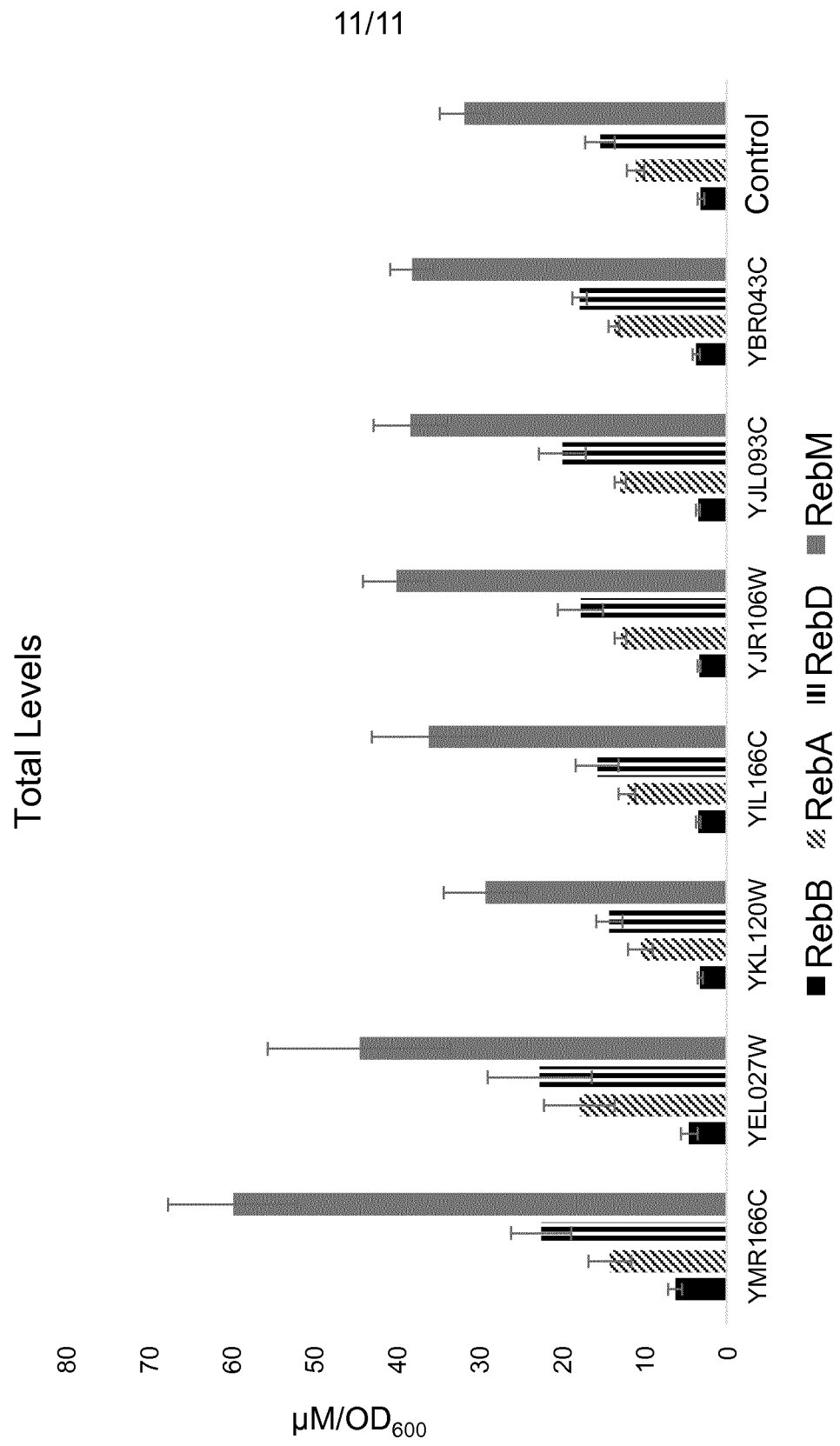


Figure 5B



INTERNATIONAL SEARCH REPORT

International application No  
PCT/EP2015/068314

A. CLASSIFICATION OF SUBJECT MATTER  
INV. A23L1/236 C12N15/81 C07H1/08  
ADD.

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)  
A23L C12N C07H

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)

EPO-Internal , EMBASE, WPI Data, Sequence Search

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	wo 2014/086890 AI (EVOLVA SA [CH] ) 12 June 2014 (2014-06-12) the whole document page 28 - page 39	1-24
Y	BRANDLE ET AL: "Stevi ol glycosi de biosynthesi s", PHYTOCHEMISTRY, PERGAMON PRESS, GB, vol . 68, no. 14, 1 July 2007 (2007-07-01) , pages 1855-1863 , XP022145443 , ISSN: 0031-9422 , DOI : 10.1016/J .PHYTOCHEM.2007.02.010 page 1860, paragraph 2 - page 1861 , paragraph 1	1-24

Further documents are listed in the continuation of Box C.

See patent family annex.

\* Special categories of cited documents :

"A" document defining the general state of the art which is not considered to be of particular relevance

"E" earlier application or patent but published on or after the international filing date

"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)

"O" document referring to an oral disclosure, use, exhibition or other means

"P" document published prior to the international filing date but later than the priority date claimed

"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention

"X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone

"Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art

"&" document member of the same patent family

Date of the actual completion of the international search

15 October 2015

Date of mailing of the international search report

20/01/2016

Name and mailing address of the ISA/

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Authorized officer

Spri nks , Matthew



## INTERNATIONAL SEARCH REPORT

International application No

PCT/EP2015/068314

C(Continuation). DOCUMENTS CONSIDERED TO BE RELEVANT		
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y	<p>YAZAKI ET AL: "ABC transporters involved in the transport of plant secondary metabolites"  FEBS LETTERS] ELSEVIER, AMSTERDAM, NL,  vol . 580, no. 4,  13 February 2006 (2006-02-13) , pages  1183-1191 , XP028030476,  ISSN: 0014-5793 , DOI :  10.1016/J . FEBSLET.2005 . 12 . 009  [retrieved on 2006-02-13]  abstract  page 1186, paragraph 4</p> <p style="text-align: center;">-----</p>	1-24
Y	<p>w0 2011/151326 A2 (VIB vzw [BE] ; UNIV  LEUVEN KATH [BE] ; THEVELEIN JOHAN [BE] ;  SAERENS SOF) 8 December 2011 (2011-12-08)  abstract; sequence 38  page 2 - page 3; sequence 38</p> <p style="text-align: center;">-----</p>	1-24
Y	<p>w0 2011/153144 A1 (GEVO INC [US] ; VIB vzw  [BE] ; UNIV LEUVEN KATH [BE] ; DUNDON  CATHERINE A) 8 December 2011 (2011-12-08)  abstract  paragraph [0014] ; table 1; sequence 38</p> <p style="text-align: center;">-----</p>	1-24

# INTERNATIONAL SEARCH REPORT

International application No.  
PCT/EP2015/068314

## Box No. II Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)

This international search report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1.  Claims Nos.:  
because they relate to subject matter not required to be searched by this Authority, namely:
  
2.  Claims Nos.:  
because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:
  
3.  Claims Nos.:  
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

## Box No. III Observations where unity of invention is lacking (Continuation of item 3 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:

see additional sheet

1.  As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.
  
2.  As all searchable claims could be searched without effort justifying an additional fees, this Authority did not invite payment of additional fees.
  
3.  As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos. :
  
4.  No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos. :

I-24(parti al ly)

### Remark on Protest

- The additional search fees were accompanied by the applicant's protest and, where applicable, the payment of a protest fee.
- The additional search fees were accompanied by the applicant's protest but the applicable protest fee was not paid within the time limit specified in the invitation.
- No protest accompanied the payment of additional search fees.

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. claims: 1-24(partially)

A recombinant host capable of synthesizing a steviol glycoside and transporting at least a portion of it into the culture medium, comprising a gene encoding a transporter polypeptide, the expression of which is modified, wherein the transporter polypeptide is an endogenous ABC transporter, and subject-matter relating thereto.

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2-32. claims: 1-24(partially)

As subject 1 but wherein the transporter polypeptide is selected from the list in claim 3, starting with MFS transporter.

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# INTERNATIONAL SEARCH REPORT

Information on patent family members

International application No <b>PCT/EP2015/068314</b>
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Patent document cited in search report	Publication date	Patent family member(s)	Publication date
WO 2014086890 A1	12-06-2014	EP 2928321 A1	14-10-2015
		US 2015342234 A1	03-12-2015
		WO 2014086890 A1	12-06-2014
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WO 2011151326 A2	08-12-2011	EP 2575499 A2	10-04-2013
		US 2013302868 A1	14-11-2013
		WO 2011151326 A2	08-12-2011
		WO 2011153144 A1	08-12-2011
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WO 2011153144 A1	08-12-2011	EP 2575499 A2	10-04-2013
		US 2013302868 A1	14-11-2013
		WO 2011151326 A2	08-12-2011
		WO 2011153144 A1	08-12-2011
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