

## Supplementary Material

The Supplementary Material for this article can be found online at the additional information tab for this paper.

### 1. Supplementary Figure

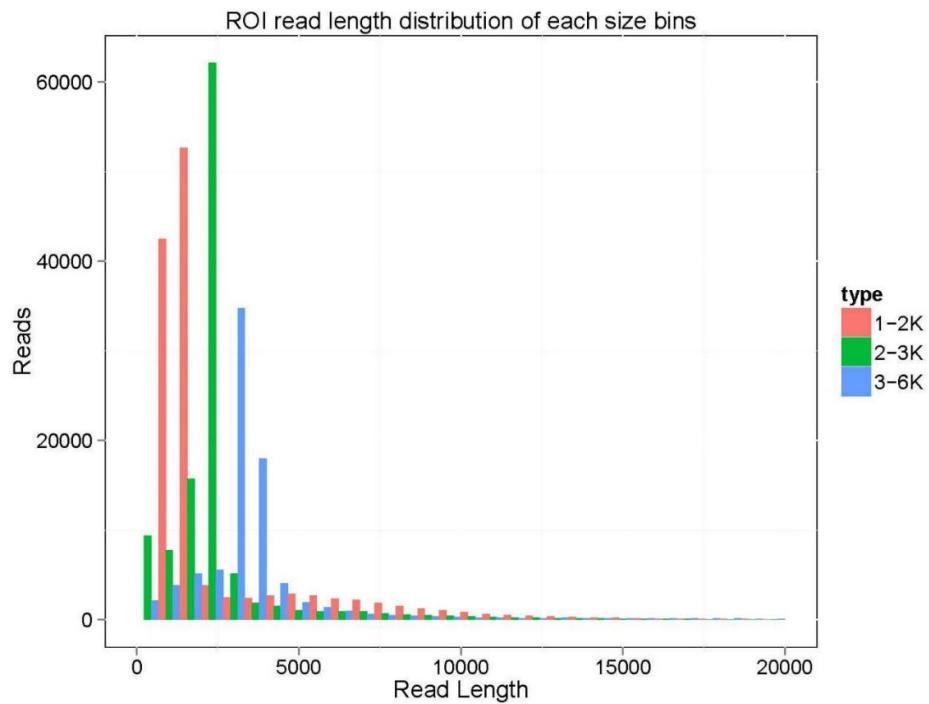


Figure S1. Summary of PacBio RS II single-molecule real-time (SMRT) sequencing.

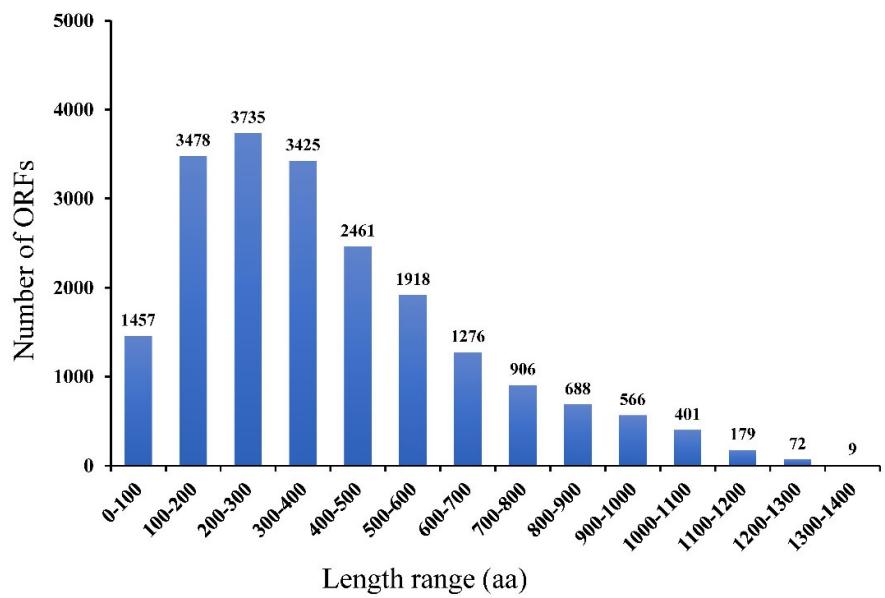
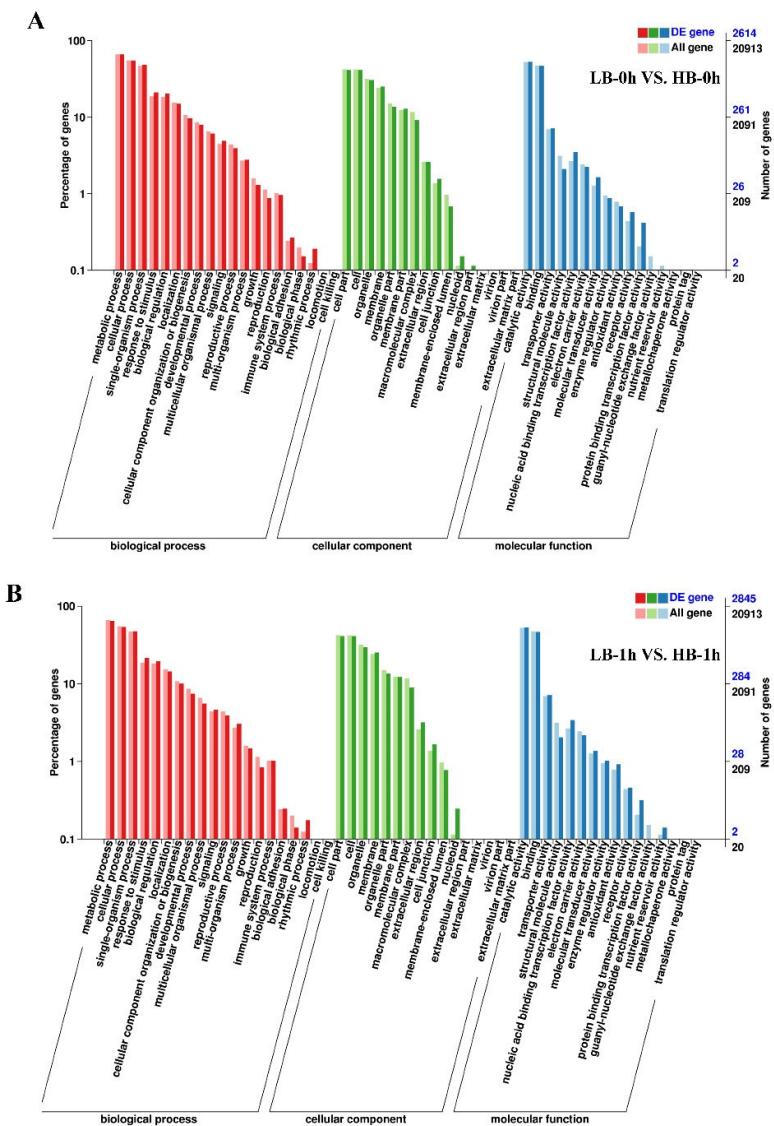


Figure S2. Length distribution of ORFs with new transcripts.



**Figure S3. Gene ontology (GO) categorization of pear flesh DEGs. (A) GO analysis of DEGs between low browning (LB) and high browning (HB) at 0h in three main categories. (B) GO analysis of DEGs between low browning (LB) and high browning (HB) at 1h in three main categories.**

**2. Supplementary Table**

**Table S1. Primers designed for qRT-PCR validation**

Category	Transcript ID	Forward primer (5'-3')	Reverse primer (5'-3')
reference gene	YLS8	TGAGGTGCTGGCTTCTGT	TGACCGTTGATGGATCGTA
	Pbr012851.1	GAGGTTCCGGTTGCATTGT	GGTGACTTCGGAATGGCTTC
	Pbr024635.1	ACAGGTGGACGGAGAGAAC	TGAGTCCGCAGAGAAACACT
	Pbr030350.1	AAGCCTCAACCTCCAAACA	TCTCCTGTGTACCCAACC
	Pbr036272.1	GAAGCACGGGACATAAACG	GTGAGACTTGTACGTTCCGC
	Pbr039972.1	GAACCGATGCCATCCAAGTC	ACATAACCCACGTCAACCAGT
	Pbr027985.1	CGGTGCTGTTCAAGGTCTA	CTTCGTTGAAATCGCTCACC
	Pbr017290.1	CAGAACATGTTGACCGGC	GTTGTAGTCGAAGCTCTGCG
	Pbr006709.1	TTGTCGGGGCTGATTCTCTT	AAGCCTCCCCTCAAATGGA
	Pbr022422.1	GCAGGTTCTGTGGATGTCCA	GAACGAGTTGCCATGATGGC
	Pbr022425.1	CTGCTCCACAGATGAAGACC	TCAAGATGTTGAGCTGATCC
	PB.10426	AGACTTTCTCTTGGCGTGT	GCGACGTTTAGTACTCTGGAG
	Pbr35186.1	ATGAATCATGCCGAATGCC	AGCAATCGTGAAGTGAAGC
mRNA	Pbr000321.4	ACCACAACTACCCTCTCCG	TCTTGCTAGGCCCTGGTCTA
	Pbr000322.1	CCCACCTGACCTAACCAAGT	TCCGGAGTGAAGGCTTGAT
	Pbr029378.1	GCTTCGGAAACTTCGGTCAG	TGCTCGAAAGATCATTGG
	Pbr017091.1	GGTGGACAGTATAAGGGCGA	TCCGGCTTCTCCCTGAAT
	Pbr026287.1	ACCCACTTGAGCCTTACCTC	TCCTCCAAGCATGACCATT
	Pbr022402.1	TCTGCAACCAACTGTCAACG	GCACACTCTCAGCACAAAGA
	Pbr019531.1	CTCGCTCTCTGGATTGCAC	ATGCACTGACATGTTCCG
	Pbr020145.1	CCAAGCCTCATCACTGGACT	TGAGAAAGGCAGAGGTCGTC
	Pbr033501.1	AGGCTGTTGGAAAGGAGTT	CCACCCACCCCTTCTCGTA
	Pbr011477.1	TACGCCAAGACCAATTCC	CAACGCTGACCAACTTGTGT
	Pbr013248.1	CCCTAGAGTCACCGTCACTG	GCCCTTCCACCGCAAAATT
	Pbr008363.1	ACAACAACGTCACTCCATGC	AGAGGCATTGAGGGTCTGAC
	PB.14775	TCCACAGCCTCAGGAAGAAC	TGCCAGCAGTACCGAAACTA
	Pbr001923.1	GTGTGAACGAAGCAGGCATT	GCTGGATTCAACATGGGTCC
	PB.156.1	GTTCTCCTCTCCTGGGCTT	GCGCCTTGTGTTGTCCTGTTA
	PB.6461.1	TTTGATGGATTCTGTGGCGG	GAGCACCCAAACCCAAATGT
lncRNA	PB.7534.1	TCCGAGTCACACCCTTCAA	TGCAGCCACGAAATTAGTCA
	PB.13941.1	TCTACCACTGCTGTGGAA	CGTCGGCCTAAAATCCTGG
	PB.8714.2	GGGAGTTGGAGAATTGAACCG	TGTTCATCGGGTACTCTCG
	PB.15038.3	TGGATGTGGGTGTGAAGGT	AGGAGGGCAAGAAGAAGCTT

**Table S2. The PacBio SMRT sequencing information of sand pear fruit**

Category	1-2kb	2-3kb	3-6kb	All
Number of SMRT cells	2	2	2	6
Number of reads of Insert	155,736	111,958	83,566	351,260
Mean Read Length of Insert	2,235	2,545	3,784	2,702
Number of five prime reads	87,476	73,389	49,472	210,337
Number of three prime reads	96,759	78,137	52,693	227,589
Number of poly-A reads	93,638	77,232	52,288	223,158
Number of filtered short reads	20,218	6,291	1,092	27,601
Number of non-full-length reads	62,831	42,706	41,892	147,429
Number of full-length reads	72,687	62,961	40,582	176,230
Number of full-length non-chimeric reads	72,370	62,718	40,545	175,633
Average full-length non-chimeric read length	1,416	2,170	3,160	2,087

**Table S3 ICE cluster result statistics**

Category	0-1kb	1-2kb	2-3kb	3-6kb	over 6kb	All
Number of consensus isoforms	2,277	30,396	23,745	12,118	586	69,122
Average consensus isoforms read length	882	1,509	2,440	3,455	9,052	2,213
Number of polished high-quality isoforms	2,084	26,080	15,627	4,003	3	47,797
Number of polished low-quality isoforms	193	4,316	8,118	8,115	583	21,325
Percent of polished high-quality isoforms (%)	91.52%	85.80%	65.81%	33.03%	0.51%	69.15%

**Table S4 Samples sequencing data evaluation**

Samples	BMK-ID	ReadSum	BaseSum	GC(%)	Q30(%)
LB-0h-1	T46	25,507,834	7,630,726,334	47.93	90.6
LB-0h-2	T47	24,472,937	7,315,878,272	48.03	90.56
LB-0h-3	T48	32,836,015	9,827,758,228	47.6	91
LB-1h-1	T49	23,778,107	7,116,394,766	47.7	90.91
LB-1h-2	T50	25,380,678	7,597,335,016	47.5	90.71
LB-1h-3	T51	25,032,532	7,491,386,194	47.69	90.55
HB-0h-1	T67	24,972,213	7,469,504,092	47.94	91.16
HB-0h-2	T68	29,059,610	8,687,022,644	47.81	90.9
HB-0h-3	T69	24,952,821	7,464,325,128	47.84	90.91
HB-1h-1	T70	25,302,776	7,566,162,862	48.26	91.26
HB-1h-2	T71	27,750,072	8,301,809,460	47.91	91.28
HB-1h-3	T72	24,857,794	7,437,296,976	47.67	91.24

**Table S5. Annotation of new transcript datasets to eight public databases**

Annotated Database	Annotated Number	300<=length<1000	length>=1000
COG_Annotation	8,255	184	8,071
GO_Annotation	12,584	313	12,271
KEGG_Annotation	9,115	252	8,863
KOG_Annotation	12,346	264	12,082
Pfam_Annotation	17,425	351	17,074
Swissprot_Annotation	14,586	320	14,266
eggNOG_Annotation	20,562	468	20,094
nr_Annotation	17,531	409	17,122
All_Annnotated	21,074	490	20,584

COG, clusters of orthologous groups; GO, gene ontology; KEGG, kyoto encyclopedia of genes and genomes; KOG.euKaryotic Ortholog Groups; Pfam,Protein family;Swiss-Prot, a well-annotated and manually checked protein database; eggNOG, evolutionary genealogy of genes: non-supervised orthologous groups; nr, a NCBI non-redundant protein database.

**Table S6. Summary of annotated differentially expressed genes between LB and HB sand pear flesh at 0h and 1h**

DEG Set	Total	COG	eggNOG	NR	Pfam	Swiss-Prot	GO	KEGG
LB VS HB at 0h	4738	1906	4542	4651	3946	3492	2614	1697
LB VS HB at 1h	5192	2078	4980	5106	4340	3843	2845	1777

**Table S7. KEGG pathway enrichment of LB vs. HB at 0h and 1h in RNA-Seq**

LB-0h vs. HB-0h					
#Kegg_pathway	ko_id	Cluter_frequency	Genome_frequency	P-value	Corrected_P-value
Phenylalanine, tyrosine and tryptophan biosynthesis	ko00400	24 out of 969 2.47678018575851%	79 out of 8004 0.987006496751624%	1.16E-05	0.00137
Protein processing in endoplasmic reticulum	ko04141	71 out of 969 7.32714138286894%	412 out of 8004 5.14742628685657%	0.001101928	0.13003
Galactose metabolism	ko00052	22 out of 969 2.2703818369453%	92 out of 8004 1.14942528735632%	0.001160464	0.13693
Plant hormone signal transduction	ko04075	73 out of 969 7.53353973168215%	443 out of 8004 5.53473263368316%	0.003202532	0.3779
Flavonoid biosynthesis	ko00941	18 out of 969 1.85758513931889%	78 out of 8004 0.974512743628186%	0.004732565	0.55844
Taurine and hypotaurine metabolism	ko00430	8 out of 969 0.825593395252838%	24 out of 8004 0.299850074962519%	0.005547423	0.6546
Ubiquinone and other terpenoid-quinone biosynthesis	ko00130	14 out of 969 1.44478844169247%	56 out of 8004 0.699650174912544%	0.00574476	0.67788
Circadian rhythm - plant	ko04712	17 out of 969 1.75438596491228%	77 out of 8004 0.962018990504748%	0.009433727	1
Biosynthesis of amino acids	ko01230	64 out of 969 6.6047471620227%	401 out of 8004 5.00999500249875%	0.011347821	1
Histidine metabolism	ko00340	10 out of 969 1.03199174406605%	38 out of 8004 0.474762618690655%	0.012719188	1
Alanine, aspartate and glutamate metabolism	ko00250	18 out of 969 1.85758513931889%	86 out of 8004 1.07446276861569%	0.013399834	1
Sulfur metabolism	ko00920	13 out of 969 1.34158926728586%	56 out of 8004 0.699650174912544%	0.014426297	1
Brassinosteroid biosynthesis	ko00905	7 out of 969 0.722394220846233%	23 out of 8004 0.28735632183908%	0.0158985	1
Biotin metabolism	ko00780	9 out of 969 0.928792569659443%	35 out of 8004 0.43728135932034%	0.020551053	1
Butanoate metabolism	ko00650	8 out of 969 0.825593395252838%	31 out of 8004 0.387306346826587%	0.027711116	1
Stilbenoid, diarylheptanoid and gingerol biosynthesis	ko00945	9 out of 969 0.928792569659443%	38 out of 8004 0.474762618690655%	0.034150411	1
Porphyrin and chlorophyll metabolism	ko00860	14 out of 969 1.44478844169247%	71 out of 8004 0.887056471764118%	0.043120371	1
Glycosaminoglycan degradation	ko00531	5 out of 969 0.515995872033024%	20 out of 8004 0.249875062468766%	0.085000441	1
Plant-pathogen interaction	ko04626	45 out of 969 4.64396284829721%	304 out of 8004 3.79810094952524%	0.086413695	1
Fatty acid biosynthesis	ko00061	16 out of 969 1.65118679050568%	93 out of 8004 1.16191904047976%	0.091707858	1
Degradation of aromatic compounds	ko01220	4 out of 969 0.412796697626419%	15 out of 8004 0.187406296851574%	0.09815157	1

Glycine, serine and threonine metabolism	ko00260	20 out of 969 2.0639834881321%	123 out of 8004 1.53673163418291%	0.102680466	1
Flavone and flavonol biosynthesis	ko00944	1 out of 969 0.103199174406605%	1 out of 8004 0.0124937531234383%	0.121064468	1
Steroid biosynthesis	ko00100	10 out of 969 1.03199174406605%	58 out of 8004 0.72463768115942%	0.157510022	1
Glycerolipid metabolism	ko00561	17 out of 969 1.75438596491228%	109 out of 8004 1.36181909045477%	0.163487036	1
Ascorbate and aldarate metabolism	ko00053	11 out of 969 1.13519091847265%	67 out of 8004 0.837081459270365%	0.181586946	1
Tropane, piperidine and pyridine alkaloid biosynthesis	ko00960	8 out of 969 0.825593395252838%	46 out of 8004 0.574712643678161%	0.186437366	1
Glyoxylate and dicarboxylate metabolism	ko00630	17 out of 969 1.75438596491228%	113 out of 8004 1.41179410294853%	0.202881024	1
Inositol phosphate metabolism	ko00562	16 out of 969 1.65118679050568%	106 out of 8004 1.32433783108446%	0.207996706	1
Terpenoid backbone biosynthesis	ko00900	12 out of 969 1.23839009287926%	77 out of 8004 0.962018990504748%	0.21671763	1
N-Glycan biosynthesis	ko00510	11 out of 969 1.13519091847265%	70 out of 8004 0.87456271864068%	0.221737206	1
Thiamine metabolism	ko00730	5 out of 969 0.515995872033024%	27 out of 8004 0.337331334332834%	0.222355712	1
Peroxisome	ko04146	21 out of 969 2.1671826625387%	147 out of 8004 1.83658170914543%	0.240003271	1
RNA transport	ko03013	40 out of 969 4.12796697626419%	296 out of 8004 3.69815092453773%	0.249119349	1
One carbon pool by folate	ko00670	7 out of 969 0.722394220846233%	44 out of 8004 0.549725137431284%	0.279491581	1
Synthesis and degradation of ketone bodies	ko00072	2 out of 969 0.206398348813209%	9 out of 8004 0.112443778110945%	0.298868642	1
beta-Alanine metabolism	ko00410	10 out of 969 1.03199174406605%	68 out of 8004 0.849575212393803%	0.305389627	1
Vancomycin resistance	ko01502	1 out of 969 0.103199174406605%	3 out of 8004 0.0374812593703148%	0.321033043	1
Ether lipid metabolism	ko00565	8 out of 969 0.825593395252838%	54 out of 8004 0.674662668665667%	0.327685671	1
Glycerophospholipid metabolism	ko00564	20 out of 969 2.0639834881321%	151 out of 8004 1.88655672163918%	0.368948395	1
Nitrogen metabolism	ko00910	8 out of 969 0.825593395252838%	57 out of 8004 0.712143928035982%	0.385168603	1
Vitamin B6 metabolism	ko00750	4 out of 969 0.412796697626419%	26 out of 8004 0.324837581209395%	0.387726228	1
C5-Branched dibasic acid metabolism	ko00660	2 out of 969 0.206398348813209%	11 out of 8004 0.137431284357821%	0.391795011	1
Propanoate metabolism	ko00640	7 out of 969 0.722394220846233%	50 out of 8004 0.624687656171914%	0.402557314	1
Cysteine and methionine metabolism	ko00270	17 out of 969 1.75438596491228%	130 out of 8004 1.62418790604698%	0.405899293	1
Pyruvate metabolism	ko00620	20 out of 969 2.0639834881321%	155 out of 8004 1.93653173413293%	0.416093694	1
Other glycan degradation	ko00511	5 out of 969 0.515995872033024%	35 out of 8004 0.43728135932034%	0.420274327	1

Lysine degradation	ko00310	7 out of 969 0.722394220846233%	52 out of 8004 0.649675162418791%	0.44404852	1	
Arachidonic acid metabolism	ko00590	4 out of 969 0.412796697626419%	28 out of 8004 0.349825087456272%	0.444927332	1	
Photosynthesis - antenna proteins	ko00196	4 out of 969 0.412796697626419%	28 out of 8004 0.349825087456272%	0.444927332	1	
Sphingolipid metabolism	ko00600	6 out of 969 0.619195046439629%	46 out of 8004 0.574712643678161%	0.489324737	1	
Cyanoamino acid metabolism	ko00460	17 out of 969 1.75438596491228%	137 out of 8004 1.71164417791104%	0.496077698	1	
alpha-Linolenic acid metabolism	ko00592	13 out of 969 1.34158926728586%	105 out of 8004 1.31184407796102%	0.510511975	1	
Tryptophan metabolism	ko00380	6 out of 969 0.619195046439629%	47 out of 8004 0.587206396801599%	0.511132536	1	
Selenocompound metabolism	ko00450	3 out of 969 0.309597523219814%	23 out of 8004 0.28735632183908%	0.539238042	1	
Pantothenate and CoA biosynthesis	ko00770	7 out of 969 0.722394220846233%	57 out of 8004 0.712143928035982%	0.544673716	1	
Valine, leucine and isoleucine degradation	ko00280	9 out of 969 0.928792569659443%	74 out of 8004 0.924537731134433%	0.54770813	1	
Carbon fixation in photosynthetic organisms	ko00710	15 out of 969 1.54798761609907%	125 out of 8004 1.56171914042979%	0.556281822	1	
Riboflavin metabolism	ko00740	2 out of 969 0.206398348813209%	15 out of 8004 0.187406296851574%	0.557677763	1	
RNA degradation	ko03018	24 out of 969 2.47678018575851%	201 out of 8004 2.51124437781109%	0.562333848	1	
SNARE interactions in vesicular transport	ko04130	9 out of 969 0.928792569659443%	75 out of 8004 0.937031484257871%	0.564631865	1	
Phenylalanine metabolism	ko00360	20 out of 969 2.0639834881321%	170 out of 8004 2.12393803098451%	0.590775878	1	
Tyrosine metabolism	ko00350	7 out of 969 0.722394220846233%	60 out of 8004 0.749625187406297%	0.60123832	1	
Citrate cycle (TCA cycle)	ko00020	11 out of 969 1.13519091847265%	95 out of 8004 1.18690654672664%	0.610458529	1	
ABC transporters	ko02010	5 out of 969 0.515995872033024%	45 out of 8004 0.562218890554723%	0.649783989	1	
Purine metabolism	ko00230	38 out of 969 3.92156862745098%	329 out of 8004 4.11044477761119%	0.649933222	1	
Zeatin biosynthesis	ko00908	3 out of 969 0.309597523219814%	28 out of 8004 0.349825087456272%	0.676115678	1	
Lipoic acid metabolism	ko00785	1 out of 969 0.103199174406605%	9 out of 8004 0.112443778110945%	0.687144396	1	
Fatty acid elongation	ko00062	5 out of 969 0.515995872033024%	47 out of 8004 0.587206396801599%	0.688187864	1	
Glycosphingolipid biosynthesis - globo series	ko00603	2 out of 969 0.206398348813209%	19 out of 8004 0.237381309345327%	0.688782489	1	
Fatty acid metabolism	ko01212	18 out of 969 1.85758513931889%	164 out of 8004 2.04897551224388%	0.708555557	1	
Biosynthesis of unsaturated fatty acids	ko01040	8 out of 969 0.825593395252838%	77 out of 8004 0.962018990504748%	0.729965756	1	
Diterpenoid biosynthesis	ko00904	4 out of 969 0.412796697626419%	40 out of 8004 0.499750124937531%	0.730556741	1	

Folate biosynthesis	ko00790	2 out of 969 0.206398348813209%	21 out of 8004 0.262368815592204%	0.741371124	1
Carbon metabolism	ko01200	46 out of 969 4.74716202270382%	411 out of 8004 5.13493253373313%	0.742462499	1
Valine, leucine and isoleucine biosynthesis	ko00290	4 out of 969 0.412796697626419%	41 out of 8004 0.512243878060969%	0.748500921	1
Glycosylphosphatidylinositol(GPI)-anchor biosynthesis	ko00563	4 out of 969 0.412796697626419%	41 out of 8004 0.512243878060969%	0.748500921	1
Endocytosis	ko04144	24 out of 969 2.47678018575851%	222 out of 8004 2.7736131934033%	0.755466516	1
Regulation of autophagy	ko04140	5 out of 969 0.515995872033024%	51 out of 8004 0.637181409295352%	0.756084531	1
Arginine and proline metabolism	ko00330	13 out of 969 1.34158926728586%	125 out of 8004 1.56171914042979%	0.761598448	1
Protein export	ko03060	8 out of 969 0.825593395252838%	80 out of 8004 0.999500249875062%	0.768455239	1
Starch and sucrose metabolism	ko00500	41 out of 969 4.23116615067079%	374 out of 8004 4.67266366816592%	0.778861502	1
Isoquinoline alkaloid biosynthesis	ko00950	3 out of 969 0.309597523219814%	33 out of 8004 0.412293853073463%	0.780461359	1
Nicotinate and nicotinamide metabolism	ko00760	2 out of 969 0.206398348813209%	23 out of 8004 0.28735632183908%	0.786164092	1
2-Oxocarboxylic acid metabolism	ko01210	9 out of 969 0.928792569659443%	91 out of 8004 1.13693153423288%	0.788159244	1
Fructose and mannose metabolism	ko00051	10 out of 969 1.03199174406605%	103 out of 8004 1.28685657171414%	0.815086554	1
Glycolysis / Gluconeogenesis	ko00010	18 out of 969 1.85758513931889%	178 out of 8004 2.22388805597201%	0.826014599	1
Phenylpropanoid biosynthesis	ko00940	33 out of 969 3.40557275541796%	318 out of 8004 3.97301349325337%	0.854360309	1
Pyrimidine metabolism	ko00240	23 out of 969 2.37358101135191%	230 out of 8004 2.8735632183908%	0.864907356	1
Phosphatidylinositol signaling system	ko04070	8 out of 969 0.825593395252838%	90 out of 8004 1.12443778110945%	0.867867638	1
Pentose and glucuronate interconversions	ko00040	15 out of 969 1.54798761609907%	158 out of 8004 1.97401299350325%	0.875269361	1
Ribosome biogenesis in eukaryotes	ko03008	14 out of 969 1.44478844169247%	151 out of 8004 1.88655672163918%	0.88904331	1
Proteasome	ko03050	9 out of 969 0.928792569659443%	103 out of 8004 1.28685657171414%	0.890554449	1
Fatty acid degradation	ko00071	6 out of 969 0.619195046439629%	73 out of 8004 0.912043978010994%	0.89083771	1
Carotenoid biosynthesis	ko00906	5 out of 969 0.515995872033024%	63 out of 8004 0.787106446776612%	0.893412431	1
Glutathione metabolism	ko00480	13 out of 969 1.34158926728586%	144 out of 8004 1.79910044977511%	0.902514708	1
Amino sugar and nucleotide sugar metabolism	ko00520	19 out of 969 1.96078431372549%	203 out of 8004 2.53623188405797%	0.911150256	1
Linoleic acid metabolism	ko00591	2 out of 969 0.206398348813209%	32 out of 8004 0.399800099950025%	0.913407179	1
Cutin, suberine and wax biosynthesis	ko00073	3 out of 969 0.309597523219814%	44 out of 8004 0.549725137431284%	0.915043145	1

Base excision repair	ko03410	5 out of 969 0.515995872033024%	70 out of 8004 0.87456271864068%	0.93757962	1
Sulfur relay system	ko04122	1 out of 969 0.103199174406605%	23 out of 8004 0.28735632183908%	0.948818047	1
Ribosome	ko03010	55 out of 969 5.67595459236326%	556 out of 8004 6.94652673663168%	0.960723211	1
Monoterpeneoid biosynthesis	ko00902	1 out of 969 0.103199174406605%	26 out of 8004 0.324837581209395%	0.965290522	1
Pentose phosphate pathway	ko00030	5 out of 969 0.515995872033024%	79 out of 8004 0.987006496751624%	0.970018166	1
Spliceosome	ko03040	31 out of 969 3.19917440660475%	343 out of 8004 4.28535732133933%	0.972680428	1
Basal transcription factors	ko03022	4 out of 969 0.412796697626419%	70 out of 8004 0.87456271864068%	0.976613748	1
Sesquiterpenoid and triterpenoid biosynthesis	ko00909	2 out of 969 0.206398348813209%	46 out of 8004 0.574712643678161%	0.980851902	1
mRNA surveillance pathway	ko03015	16 out of 969 1.65118679050568%	211 out of 8004 2.63618190904548%	0.988222741	1
Ubiquitin mediated proteolysis	ko04120	16 out of 969 1.65118679050568%	221 out of 8004 2.76111944027986%	0.993750018	1
RNA polymerase	ko03020	6 out of 969 0.619195046439629%	110 out of 8004 1.37431284357821%	0.994201848	1
Oxidative phosphorylation	ko00190	23 out of 969 2.37358101135191%	295 out of 8004 3.68565717141429%	0.994264857	1
Aminoacyl-tRNA biosynthesis	ko00970	5 out of 969 0.515995872033024%	111 out of 8004 1.38680659670165%	0.998365644	1
Phagosome	ko04145	6 out of 969 0.619195046439629%	148 out of 8004 1.84907546226887%	0.999835082	1
DNA replication	ko03030	3 out of 969 0.309597523219814%	114 out of 8004 1.42428785607196%	0.999947772	1
Photosynthesis	ko00195	12 out of 969 1.23839009287926%	243 out of 8004 3.0359820089955%	0.999965781	1
Nucleotide excision repair	ko03420	3 out of 969 0.309597523219814%	118 out of 8004 1.47426286856572%	0.999966969	1
Homologous recombination	ko03440	1 out of 969 0.103199174406605%	83 out of 8004 1.03698150924538%	0.999978971	1
Mismatch repair	ko03430	1 out of 969 0.103199174406605%	88 out of 8004 1.09945027486257%	0.999989051	1

LB-1h vs. HB-1h					
#Kegg_pathway	ko_id	Cluter_frequency	Genome_frequency	P-value	Corrected_P-value
Galactose metabolism	ko00052	27 out of 1027 2.62901655306719%	92 out of 8004 1.14942528735632%	2.04E-05	0.0024
Phenylalanine, tyrosine and tryptophan biosynthesis	ko00400	22 out of 1027 2.14216163583252%	79 out of 8004 0.987006496751624%	0.000270298	0.0319
Ubiquinone and other terpenoid-quinone biosynthesis	ko00130	17 out of 1027 1.65530671859786%	56 out of 8004 0.699650174912544%	0.000440318	0.05196
Plant-pathogen interaction	ko04626	56 out of 1027 5.45277507302824%	304 out of 8004 3.79810094952524%	0.002879213	0.33975
Protein processing in endoplasmic reticulum	ko04141	72 out of 1027 7.01071080817916%	412 out of 8004 5.14742628685657%	0.00327907	0.38693
Alanine, aspartate and glutamate metabolism	ko00250	20 out of 1027 1.94741966893866%	86 out of 8004 1.07446276861569%	0.005325293	0.62838
Histidine metabolism	ko00340	11 out of 1027 1.07108081791626%	38 out of 8004 0.474762618690655%	0.006569975	0.77526
Glycosaminoglycan degradation	ko00531	7 out of 1027 0.68159688412853%	20 out of 8004 0.249875062468766%	0.009552215	1
Sulfur metabolism	ko00920	14 out of 1027 1.36319376825706%	56 out of 8004 0.699650174912544%	0.009553884	1
Biotin metabolism	ko00780	10 out of 1027 0.973709834469328%	35 out of 8004 0.43728135932034%	0.010349933	1
Biosynthesis of amino acids	ko01230	67 out of 1027 6.5238558909445%	401 out of 8004 5.00999500249875%	0.012518492	1
Porphyrin and chlorophyll metabolism	ko00860	16 out of 1027 1.55793573515093%	71 out of 8004 0.887056471764118%	0.016109332	1
Steroid biosynthesis	ko00100	13 out of 1027 1.26582278481013%	58 out of 8004 0.72463768115942%	0.029597701	1
Circadian rhythm - plant	ko04712	16 out of 1027 1.55793573515093%	77 out of 8004 0.962018990504748%	0.033072866	1
Degradation of aromatic compounds	ko01220	5 out of 1027 0.486854917234664%	15 out of 8004 0.187406296851574%	0.034233485	1
Flavonoid biosynthesis	ko00941	16 out of 1027 1.55793573515093%	78 out of 8004 0.974512743628186%	0.036844833	1
Stilbenoid, diarylheptanoid and gingerol biosynthesis	ko00945	9 out of 1027 0.876338851022395%	38 out of 8004 0.474762618690655%	0.04716511	1
Photosynthesis - antenna proteins	ko00196	7 out of 1027 0.68159688412853%	28 out of 8004 0.349825087456272%	0.058773187	1
Sphingolipid metabolism	ko00600	10 out of 1027 0.973709834469328%	46 out of 8004 0.574712643678161%	0.062990798	1
Taurine and hypotaurine metabolism	ko00430	6 out of 1027 0.584225900681597%	24 out of 8004 0.299850074962519%	0.077604124	1
Ascorbate and aldarate metabolism	ko00053	13 out of 1027 1.26582278481013%	67 out of 8004 0.837081459270365%	0.081340102	1
Glyoxylate and dicarboxylate metabolism	ko00630	20 out of 1027 1.94741966893866%	113 out of 8004 1.41179410294853%	0.082420896	1
Butanoate metabolism	ko00650	7 out of 1027 0.68159688412853%	31 out of 8004 0.387306346826587%	0.093358594	1
Cysteine and methionine metabolism	ko00270	22 out of 1027 2.14216163583252%	130 out of 8004 1.62418790604698%	0.104103969	1

Riboflavin metabolism	ko00740	4 out of 1027 0.389483933787731%	15 out of 8004 0.187406296851574%	0.11587282	1
Thiamine metabolism	ko00730	6 out of 1027 0.584225900681597%	27 out of 8004 0.337331334332834%	0.12313498	1
Inositol phosphate metabolism	ko00562	18 out of 1027 1.75267770204479%	106 out of 8004 1.32433783108446%	0.12870001	1
Arachidonic acid metabolism	ko00590	6 out of 1027 0.584225900681597%	28 out of 8004 0.349825087456272%	0.140688197	1
Glycerolipid metabolism	ko00561	18 out of 1027 1.75267770204479%	109 out of 8004 1.36181909045477%	0.155214933	1
Glycine, serine and threonine metabolism	ko00260	20 out of 1027 1.94741966893866%	123 out of 8004 1.53673163418291%	0.155980595	1
Brassinosteroid biosynthesis	ko00905	5 out of 1027 0.486854917234664%	23 out of 8004 0.28735632183908%	0.163755457	1
Peroxisome	ko04146	23 out of 1027 2.23953261927945%	147 out of 8004 1.83658170914543%	0.180942223	1
Fatty acid biosynthesis	ko00061	15 out of 1027 1.46056475170399%	93 out of 8004 1.16191904047976%	0.20759005	1
Glycosphingolipid biosynthesis - ganglio series	ko00604	2 out of 1027 0.194741966893866%	7 out of 8004 0.087456271864068%	0.223517184	1
Plant hormone signal transduction	ko04075	62 out of 1027 6.03700097370983%	443 out of 8004 5.53473263368316%	0.245251315	1
Amino sugar and nucleotide sugar metabolism	ko00520	29 out of 1027 2.82375851996105%	203 out of 8004 2.53623188405797%	0.294776533	1
Synthesis and degradation of ketone bodies	ko00072	2 out of 1027 0.194741966893866%	9 out of 8004 0.112443778110945%	0.324487842	1
One carbon pool by folate	ko00670	7 out of 1027 0.68159688412853%	44 out of 8004 0.549725137431284%	0.33249583	1
Vancomycin resistance	ko01502	1 out of 1027 0.0973709834469328%	3 out of 8004 0.0374812593703148%	0.337690533	1
Phenylalanine metabolism	ko00360	24 out of 1027 2.33690360272639%	170 out of 8004 2.12393803098451%	0.339366427	1
Fructose and mannose metabolism	ko00051	15 out of 1027 1.46056475170399%	103 out of 8004 1.28685657171414%	0.340497574	1
Selenocompound metabolism	ko00450	4 out of 1027 0.389483933787731%	23 out of 8004 0.28735632183908%	0.340627862	1
Valine, leucine and isoleucine degradation	ko00280	11 out of 1027 1.07108081791626%	74 out of 8004 0.924537731134433%	0.349031221	1
SNARE interactions in vesicular transport	ko04130	11 out of 1027 1.07108081791626%	75 out of 8004 0.937031484257871%	0.3661016	1
Phosphatidylinositol signaling system	ko04070	13 out of 1027 1.26582278481013%	90 out of 8004 1.12443778110945%	0.368260772	1
Tryptophan metabolism	ko00380	7 out of 1027 0.68159688412853%	47 out of 8004 0.587206396801599%	0.398662061	1
Endocytosis	ko04144	30 out of 1027 2.92112950340798%	222 out of 8004 2.7736131934033%	0.409316823	1
C5-Branched dibasic acid metabolism	ko00660	2 out of 1027 0.194741966893866%	11 out of 8004 0.137431284357821%	0.421786179	1
Phenylpropanoid biosynthesis	ko00940	42 out of 1027 4.08958130477118%	318 out of 8004 3.97301349325337%	0.444917186	1
Glycosphingolipid biosynthesis - globo series	ko00603	3 out of 1027 0.292112950340798%	19 out of 8004 0.237381309345327%	0.447974619	1

Nitrogen metabolism	ko00910	8 out of 1027 0.778967867575463%	57 out of 8004 0.712143928035982%	0.451103583	1
Other glycan degradation	ko00511	5 out of 1027 0.486854917234664%	35 out of 8004 0.43728135932034%	0.472670518	1
Tyrosine metabolism	ko00350	8 out of 1027 0.778967867575463%	60 out of 8004 0.749625187406297%	0.511509866	1
beta-Alanine metabolism	ko00410	9 out of 1027 0.876338851022395%	68 out of 8004 0.849575212393803%	0.514749301	1
Glycolysis / Gluconeogenesis	ko00010	23 out of 1027 2.23953261927945%	178 out of 8004 2.22388805597201%	0.519931798	1
Biosynthesis of unsaturated fatty acids	ko01040	10 out of 1027 0.973709834469328%	77 out of 8004 0.962018990504748%	0.535158114	1
Terpenoid backbone biosynthesis	ko00900	10 out of 1027 0.973709834469328%	77 out of 8004 0.962018990504748%	0.535158114	1
Carbon fixation in photosynthetic organisms	ko00710	16 out of 1027 1.55793573515093%	125 out of 8004 1.56171914042979%	0.545005908	1
Tropane, piperidine and pyridine alkaloid biosynthesis	ko00960	6 out of 1027 0.584225900681597%	46 out of 8004 0.574712643678161%	0.549070618	1
Sesquiterpenoid and triterpenoid biosynthesis	ko00909	6 out of 1027 0.584225900681597%	46 out of 8004 0.574712643678161%	0.549070618	1
Carotenoid biosynthesis	ko00906	8 out of 1027 0.778967867575463%	63 out of 8004 0.787106446776612%	0.569578711	1
Fatty acid elongation	ko00062	6 out of 1027 0.584225900681597%	47 out of 8004 0.587206396801599%	0.571081626	1
Nicotinate and nicotinamide metabolism	ko00760	3 out of 1027 0.292112950340798%	23 out of 8004 0.28735632183908%	0.580994698	1
Diterpenoid biosynthesis	ko00904	5 out of 1027 0.486854917234664%	40 out of 8004 0.499750124937531%	0.596259601	1
alpha-Linolenic acid metabolism	ko00592	13 out of 1027 1.26582278481013%	105 out of 8004 1.31184407796102%	0.599577573	1
Fatty acid degradation	ko00071	9 out of 1027 0.876338851022395%	73 out of 8004 0.912043978010994%	0.604283443	1
Glycosylphosphatidylinositol(GPI)-anchor biosynthesis	ko00563	5 out of 1027 0.486854917234664%	41 out of 8004 0.512243878060969%	0.618973193	1
Carbon metabolism	ko01200	51 out of 1027 4.96592015579357%	411 out of 8004 5.13493253373313%	0.626830006	1
RNA transport	ko03013	36 out of 1027 3.50535540408958%	296 out of 8004 3.69815092453773%	0.663679772	1
Vitamin B6 metabolism	ko00750	3 out of 1027 0.292112950340798%	26 out of 8004 0.324837581209395%	0.666358362	1
Monoterpene biosynthesis	ko00902	3 out of 1027 0.292112950340798%	26 out of 8004 0.324837581209395%	0.666358362	1
Glycerophospholipid metabolism	ko00564	18 out of 1027 1.75267770204479%	151 out of 8004 1.88655672163918%	0.668901433	1
Lysine degradation	ko00310	6 out of 1027 0.584225900681597%	52 out of 8004 0.649675162418791%	0.672344057	1
N-Glycan biosynthesis	ko00510	8 out of 1027 0.778967867575463%	70 out of 8004 0.87456271864068%	0.691093704	1

Cyanoamino acid metabolism	ko00460	16 out of 1027 1.55793573515093%	137 out of 8004 1.71164417791104%	0.696003568	1
ABC transporters	ko02010	5 out of 1027 0.486854917234664%	45 out of 8004 0.562218890554723%	0.701833907	1
Ether lipid metabolism	ko00565	6 out of 1027 0.584225900681597%	54 out of 8004 0.674662668665667%	0.708261107	1
Lipoic acid metabolism	ko00785	1 out of 1027 0.0973709834469328%	9 out of 8004 0.112443778110945%	0.709619774	1
Protein export	ko03060	9 out of 1027 0.876338851022395%	80 out of 8004 0.999500249875062%	0.71406847	1
Fatty acid metabolism	ko01212	19 out of 1027 1.85004868549172%	164 out of 8004 2.04897551224388%	0.71955588	1
Purine metabolism	ko00230	39 out of 1027 3.79746835443038%	329 out of 8004 4.11044477761119%	0.730207714	1
Arginine and proline metabolism	ko00330	14 out of 1027 1.36319376825706%	125 out of 8004 1.56171914042979%	0.747522802	1
Pantothenate and CoA biosynthesis	ko00770	6 out of 1027 0.584225900681597%	57 out of 8004 0.712143928035982%	0.756922193	1
Starch and sucrose metabolism	ko00500	44 out of 1027 4.28432327166504%	374 out of 8004 4.67266366816592%	0.758620084	1
Folate biosynthesis	ko00790	2 out of 1027 0.194741966893866%	21 out of 8004 0.262368815592204%	0.771596584	1
Valine, leucine and isoleucine biosynthesis	ko00290	4 out of 1027 0.389483933787731%	41 out of 8004 0.512243878060969%	0.789744289	1
Citrate cycle (TCA cycle)	ko00020	10 out of 1027 0.973709834469328%	95 out of 8004 1.18690654672664%	0.793498068	1
Linoleic acid metabolism	ko00591	3 out of 1027 0.292112950340798%	32 out of 8004 0.399800099950025%	0.797361837	1
Sulfur relay system	ko04122	2 out of 1027 0.194741966893866%	23 out of 8004 0.28735632183908%	0.814058401	1
Pentose and glucuronate interconversions	ko00040	17 out of 1027 1.65530671859786%	158 out of 8004 1.97401299350325%	0.816474564	1
Cutin, suberine and wax biosynthesis	ko00073	4 out of 1027 0.389483933787731%	44 out of 8004 0.549725137431284%	0.833890279	1
Glutathione metabolism	ko00480	15 out of 1027 1.46056475170399%	144 out of 8004 1.79910044977511%	0.841645452	1
2-Oxocarboxylic acid metabolism	ko01210	9 out of 1027 0.876338851022395%	91 out of 8004 1.13693153423288%	0.842185466	1
Pyruvate metabolism	ko00620	16 out of 1027 1.55793573515093%	155 out of 8004 1.93653173413293%	0.857654584	1
Oxidative phosphorylation	ko00190	32 out of 1027 3.11587147030185%	295 out of 8004 3.68565717141429%	0.871471566	1
RNA degradation	ko03018	21 out of 1027 2.04479065238559%	201 out of 8004 2.51124437781109%	0.872613453	1
Propanoate metabolism	ko00640	4 out of 1027 0.389483933787731%	50 out of 8004 0.624687656171914%	0.899167023	1
Ubiquitin mediated proteolysis	ko04120	21 out of 1027 2.04479065238559%	221 out of 8004 2.76111944027986%	0.950220039	1
Pentose phosphate pathway	ko00030	6 out of 1027 0.584225900681597%	79 out of 8004 0.987006496751624%	0.950382814	1
RNA polymerase	ko03020	9 out of 1027 0.876338851022395%	110 out of 8004 1.37431284357821%	0.953810741	1

Base excision repair	ko03410	5 out of 1027 0.486854917234664%	70 out of 8004 0.87456271864068%	0.955937968	1
Spliceosome	ko03040	34 out of 1027 3.31061343719572%	343 out of 8004 4.28535732133933%	0.962273905	1
Proteasome	ko03050	8 out of 1027 0.778967867575463%	103 out of 8004 1.28685657171414%	0.962718488	1
Regulation of autophagy	ko04140	3 out of 1027 0.292112950340798%	51 out of 8004 0.637181409295352%	0.967553188	1
mRNA surveillance pathway	ko03015	19 out of 1027 1.85004868549172%	211 out of 8004 2.63618190904548%	0.968097755	1
Pyrimidine metabolism	ko00240	21 out of 1027 2.04479065238559%	230 out of 8004 2.8735632183908%	0.96896054	1
Zeatin biosynthesis	ko00908	1 out of 1027 0.0973709834469328%	28 out of 8004 0.349825087456272%	0.97876269	1
Ribosome biogenesis in eukaryotes	ko03008	12 out of 1027 1.16845180136319%	151 out of 8004 1.88655672163918%	0.979169321	1
Homologous recombination	ko03440	5 out of 1027 0.486854917234664%	83 out of 8004 1.03698150924538%	0.986411879	1
Isoquinoline alkaloid biosynthesis	ko00950	1 out of 1027 0.0973709834469328%	33 out of 8004 0.412293853073463%	0.989341285	1
Aminoacyl-tRNA biosynthesis	ko00970	6 out of 1027 0.584225900681597%	111 out of 8004 1.38680659670165%	0.997139502	1
Nucleotide excision repair	ko03420	6 out of 1027 0.584225900681597%	118 out of 8004 1.47426286856572%	0.998554739	1
Ribosome	ko03010	50 out of 1027 4.86854917234664%	556 out of 8004 6.94652673663168%	0.998608394	1
Phagosome	ko04145	8 out of 1027 0.778967867575463%	148 out of 8004 1.84907546226887%	0.999199931	1
Basal transcription factors	ko03022	2 out of 1027 0.194741966893866%	70 out of 8004 0.87456271864068%	0.999270427	1
DNA replication	ko03030	5 out of 1027 0.486854917234664%	114 out of 8004 1.42428785607196%	0.999395544	1
Mismatch repair	ko03430	3 out of 1027 0.292112950340798%	88 out of 8004 1.09945027486257%	0.999478854	1
Photosynthesis	ko00195	11 out of 1027 1.07108081791626%	243 out of 8004 3.0359820089955%	0.999997301	1

### **3. Supplementary Methods**

#### **3.1 LC-MS analyses**

The powder samples of sand pears (0.1 g) were extracted with 1 mL 80% methanol. The extraction was performed by incubation for 30 min under sonication with occasional shaking. Next, the slurry was centrifuged at 19,000g for 10 min, the supernatant was placed on 10mL glass centrifuge tube and concentrated to dry in vacuum at 37 °C. The residue was redissolved in 150 μl 80% methanol water solution and centrifuged for 10min at 14000g, then the supernatant was used for analysis.

The LC-MS/MS portion of the platform was based on a UHPLC system (1290 series, Agilent Technologies) equipped with an ACQUITY UPLC BEH Amide column (1.7 μm, 2.1 mm×100 mm, Wasters) and a triple quadruple mass spectrometer (5500 QTRAP, AB SCIEX) in the multiple reaction monitoring (MRM) mode. Metabolites were detected in electrospray negative-ionization and positive-ionization mode. The 2 μL samples were injected sequentially. The ACQUITY UPLC BEH Amide column (1.7 μm, 2.1 mm×100 mm, Wasters) was heated to 45 °C under a flow rate of 300 μL/min. A gradient was used to separate the compounds consisted of solvent A (20 mM ammonium acetate, 50% acetonitrile, pH 9.5) and solvent B (100% acetonitrile). The gradient started at 5% solvent A for 1 min and increasing linearly to 35% solvent A over 11 min, and then increased linearly to 60% solvent A over 2 min, and then decreasing to 5% of solvent A in 2 min, then keeping for extra 2 min. QC samples were injected during acquisition.

The MS conditions were set as follows: Source Temperature 550°C, Ion Source Gas1(GAS1): 40, Ion Source Gas2(GAS2): 50, Curtain Gas(CUR): 35, Ion Spray Voltage Floating (ISVF) : -4500V; The mass spectrometer was operated with a dwell time of 200 ms. To construct the metabolite MRM library, each metabolite standard (50 mg/mL) was first analyzed by LC-MS/MS to get the optimal MRM transition parameters. Then the retention time of each metabolite was determined by measuring the corresponding MRM (Q1/Q3) transition individually. A standard mixture sample (6 standard mixtures) that contains all of the 6 metabolites was measured together and used as the reference peaks for metabolite identification. Raw MRM data files were processed by peak finding, alignment, and filtering using MultiQuant software.