

Supplementary Table S1: Previously reported metabolic and reproductive associations with selected candidate SNPs from core circadian regulating genes.

Gene/Gene name/ SNP	Functional consequence	Relevant Metabolic and Reproductive Associations	References
<b><i>CLOCK</i></b>			
<b><i>Clock Circadian Regulator</i></b>			
rs3749474	3' UTR	weight loss, obesity, BMI, infertility	(Garaulet et al., 2009, 2010; Loria-Kohen et al., 2016; Shen et al., 2015b)
rs4580704	intron variant	metabolic syndrome, obesity, overweight, BMI, type-2 diabetes, coronary heart disease related dyslipidemia, non-alcoholic fatty liver disease	(Bandín et al., 2013; Corella et al., 2016; Garaulet et al., 2009, 2010, 2011; Gomez-Delgado et al., 2015; Sookoian et al., 2007, 2008)
rs1464490	intron variant	obesity	(Corella et al., 2016; Garaulet et al., 2010)
rs6843722	intron variant	obesity	(Sookoian et al., 2007, 2008; Valladares et al., 2015)
rs6850524	intron variant	metabolic syndrome, obesity, idiopathic recurrent spontaneous abortion, infertility	(Hodžić et al., 2013, 2018; Sookoian et al., 2008, 2010; Ye et al., 2016)
rs4864548	non-coding transcript exon variant	metabolic syndrome, obesity, type-2 diabetes	(Krishnan et al., 2017; Scott et al., 2008; Sookoian et al., 2008, 2010; Uemura et al., 2016)
rs1801260	3' UTR	insulin metabolism, metabolic syndrome, fatty acids, obesity, LDL cholesterol	(Bandín et al., 2013; Galbete et al., 2012; Garaulet et al., 2009, 2010, 2011, 2012; Garcia-Rios et al., 2014; Scott et al., 2008; Shen et al., 2015b; Tsuzaki et al., 2010; Uemura et al., 2016)
<b><i>ARNTL</i></b>			
<b><i>Aryl Hydrocarbon Receptor Nuclear Translocator Like</i></b>			
rs2278749	intron variant	infertility	(Kovanen et al., 2010)
rs6486121	intron variant	type-2 diabetes, hypertension, HDL cholesterol	(Klarin et al., 2018; Woon et al., 2007)

rs7950226	intron variant	type-2 diabetes, gestational diabetes mellitus, hypertension, metabolic syndrome	(Kelly et al., 2012; Pappa et al., 2013; Woon et al., 2007)
rs11022775	intron variant	gestational diabetes mellitus, type-2 diabetes	(Kelly et al., 2012; Pappa et al., 2013; Woon et al., 2007)
<b><i>PER1</i></b>			
<b>Period Circadian regulator 1</b>			
rs2585405	missense variant	reproductive complications	(Chu et al., 2008)
rs3027178	synonymous variant	hepatocellular carcinoma	(Zhang et al., 2014)
<b><i>PER2</i></b>			
<b>Period Circadian regulator 2</b>			
rs2304672	5' UTR	plasma fatty acid composition, obesity	(Garaulet et al., 2010; Garcia-Rios et al., 2012)
rs56013859	intron variant	fasting blood glucose	(Englund et al., 2009)
rs7602358	intron variant	type-2 diabetes	(Kelly et al., 2012)
<b><i>PER3</i></b>			
<b>Period Circadian regulator 3</b>			
rs228669	synonymous variant	hepatocellular carcinoma	(Zhang et al., 2014)
rs2640908	synonymous variant	hepatocellular carcinoma	(Zhang et al., 2014; Zhao et al., 2012)
<b><i>CRY1</i></b>			
<b>Cryptochrome Circadian regulator 3</b>			
rs2287161	regulatory region variant	insulin resistance	(Dashti et al., 2014)
rs3809236	5' UTR	hepatocellular carcinoma	(Zhang et al., 2014)
rs12315175	intron variant	type-2 diabetes	(Kelly et al., 2012)
<b><i>CRY2</i></b>			
<b>Cryptochrome Circadian regulator 2</b>			
rs2292912	non coding transcript exon variant	type-2 diabetes	(Kelly et al., 2012)

rs11605924

intron variant

type-2 diabetes, fasting glucose  
glucose metabolism, HDL cholesterol(Barker et al., 2011; Dashti et al., 2015;  
Dupuis et al., 2010; Hu et al., 2010;  
Langlois et al., 2016; Manning et al.,  
2012; Mirzaei et al., 2014; Renström et  
al., 2015)**NPAS2****Neuronal PAS domain Protein 2**

rs2305160

missense variant

reproductive complications

(Yuan et al., 2014; Kovanen et al., 2010;  
Chu et al., 2008)  
(Englund et al., 2009)

rs11541353

missense variant

hypertension

Supplementary Table S2: Previously reported metabolic and reproductive associations with selected candidate SNPs for circadian-related and lipid-related genes.

Gene/Gene name/ SNPs	Functional consequence	Relevant Metabolic and Reproductive Associations	References
<b><i>SIRT1</i></b>			
<b>Sirtuin 1</b>			
rs12413112	intron variant	BMI	(Clark et al., 2012)
rs3758391	intergenic variant	type- 2 diabetes	(Cruz et al., 2010)
rs2273773	synonymous variant	BMI, hypertension, total cholesterol, hyperglycemia, obesity	(Berg et al., 2009; Kelic et al., 2014; Shimoyama et al., 2011, 2012; Zhong et al., 2015b)
rs10997860	intron variant	reproductive complications, placental abruption	(Workalemahu et al., 2013)
<b><i>CELSR2-PSRC1-SORT1</i> gene cluster</b>			
<b>Cadherin, EGF LAG Seven-pass G-type Receptor 2, Proline and Serine rich coiled-coil 1, Sortilin 1</b>			
rs646776	regulatory region variant	LDL cholesterol, HDL cholesterol, C-reactive protein, blood protein levels, coronary artery disease	(Arvind et al., 2014; Aulchenko et al., 2009; Chasman et al., 2008; Consortium, 2011; Devaney et al., 2011; Dumitrescu et al., 2011; Gigante et al., 2012; Grallert et al., 2012; Guo et al., 2015a; He et al., 2017; Hubacek et al., 2017; Jansen et al., 2014; Kanai et al., 2018; Kathiresan et al., 2008; Keebler et al., 2010; Keebler Mary E. et al., 2009; Lanktree et al., 2009; Lighart et al., 2016; Lu et al., 2010; Muendlein et al., 2009; Murray et al., 2009; Qi et al., 2011a, 2011b; Reilly et al., 2011; Saleheen et al., 2010; Sandhu et al., 2008; Shirts et al., 2011; Suhre et al., 2017; Sun et al., 2018; Surakka et al., 2015; Walia et al., 2014; Wallace et al., 2008; Zhou et al., 2015)

rs599839

regulatory region variant

LDL cholesterol, fasting glucose, coronary artery disease, total cholesterol, triglycerides

(Abe et al., 2015; Angelakopoulou et al., 2012; Breitling et al., 2015; Cho et al., 2009; Coronary Artery Disease Consortium et al., 2009; Ellis et al., 2011; Fujimaki et al., 2015; Gigante et al., 2012; Grallert et al., 2012; Guo et al., 2015a; He et al., 2017; Kathiresan et al., 2008; Kleber et al., 2010; Linsel-Nitschke et al., 2010; Ma et al., 2010; Ogawa et al., 2010; Roslin et al., 2009; Samani et al., 2008; Sandhu et al., 2008; Schunkert et al., 2011; Spracklen et al., 2017; Wallace et al., 2008; Wang et al., 2011a; Willer et al., 2008; Zhou et al., 2011, 2015)

**POMC****Proopiomelanocortin**

rs28932472

missense variant

LDL cholesterol, blood pressure

(Queiroz et al., 2015)

**PPARA****Peroxisome Proliferator Activated Receptor Alpha**

rs1800206

missense variant

type-2 diabetes, obesity, LDL cholesterol, dyslipidemia, hypertriglyceridemia

(Alsaleh et al., 2011; Andrulionytè et al., 2007; Bouchard-Mercier et al., 2011; Costa-Urrutia et al., 2017; Dong et al., 2015; Fan et al., 2015; Gu et al., 2015)

**PPARG****Peroxisome Proliferator Activated Receptor****Gamma**

rs7638903

intron variant

type-2 diabetes,

(Claussnitzer et al., 2014)

rs1801282

missense variant

type-2 diabetes, BMI, insulin resistance, total cholesterol, triglycerides, obesity, hypertension, metabolic syndrome, coronary artery disease

(Bego et al., 2011; Ben Ali et al., 2009; Bordini et al., 2017; Bystryva et al., 2017; Chan et al., 2013; Dedoussis et al., 2009; DIAbetes Genetics Replication And Meta-analysis (DIAGRAM) Consortium et al., 2014; Ding et al., 2012; Estevalet et al., 2011; Gaulton et al., 2008; Gouda et al., 2010; Hindy et

***PPARGC1A***  
**Peroxisome Proliferator  
Activated Receptor Gamma  
Coactivator 1-alpha**

rs12640088	intron variant	type- 2 diabetes	(Villegas et al., 2014)
rs8192678	missense variant	type-2 diabetes, Non-alcoholic steatohepatitis, insulin resistance, BMI	(Deeb and Brunzell, 2009; Franks et al., 2014; Lin et al., 2013; Povel et al., 2010; Queiroz et al., 2015; Tai et al., 2016; Zhu et al., 2017a)
rs251464	intron variant	BMI, type-2 diabetes	(Villegas et al., 2014)

***VDR***  
**Vitamin D Receptor**

rs17383291	intergenic variant	colorectal cancer	(Poynter et al., 2010)
rs2228570	start lost	BMI, insulin resistance, type-2 diabetes, triglycerides, HDL cholesterol, hypertension	(Jia et al., 2015; Koroglu et al., 2014; Li et al., 2014; Safar et al., 2018; Zhong et al., 2015a)

***FOXO1***  
**Forkhead Box O1**

rs10507486	intron variant	carotid atherosclerosis, type-2 diabetes	(Kedenko et al., 2014; Sookoian et al.,
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al., 2016; Hoffmann et al., 2018; Hsiao and Lin, 2015; Huang et al., 2016; Kilpeläinen et al., 2008; Li et al., 2015; Manning et al., 2012; Morris et al., 2012; Namvaran et al., 2011; Phani et al., 2016; Queiroz et al., 2015; Regieli et al., 2009; Sanghera et al., 2008, 2010; Saxena et al., 2007; Scott et al., 2007; Shi et al., 2012; Stancáková et al., 2009; Tan et al., 2014; Tellechea et al., 2009; Trombetta et al., 2013; Wang et al., 2015b; Wang and Liu, 2012; Wu et al., 2016; Zeggini et al., 2007; Zhao et al., 2017; Zhu et al., 2017a)

rs2297627

intron variant

carotid atherosclerosis

2009b)

(Kedenko et al., 2014; Muller et al., 2015; Müssig et al., 2009)

***MTNR1B*****Melatonin Receptor 1B**

rs10830963

intron variant

gestational diabetes mellitus, type-2 diabetes, fasting glucose, hemoglobin A1C, insulin resistance, birth weight, obesity-related traits

(Barker et al., 2011; Beaumont et al., 2018; Chambers et al., 2009; Comuzzie et al., 2012; de Luis et al., 2018; DIAbetes Genetics Replication And Meta-analysis (DIAGRAM) Consortium et al., 2014; Dupuis et al., 2010; Evangelou et al., 2018; Garaulet et al., 2015; Goni et al., 2014; Holzapfel et al., 2011; Horikoshi et al., 2016; Hu et al., 2010; Huopio et al., 2013; Ingelsson et al., 2010; Kanai et al., 2018; Keaton et al., 2018; Kettunen et al., 2012; Kong et al., 2015; Kwak et al., 2012; Lane et al., 2016; Langenberg et al., 2009; Liao et al., 2012; Liu et al., 2010; Loomis et al., 2018; Lu et al., 2017; Mao et al., 2012; Morris et al., 2012; Nettleton et al., 2007; Ohshige et al., 2011; Palmer et al., 2015; Prokopenko et al., 2009, 2014; Qi et al., 2017; Rasmussen-Torvik et al., 2010; Rasmussen-Torvik et al., 2012; Reinehr et al., 2011; Ren et al., 2014; Renström et al., 2011, 2015; Rönn et al., 2009; Salman et al., 2015; Simonis-Bik et al., 2010; Sparsø et al., 2009; Staiger et al., 2008; Stuebe et al., 2014; Tam et al., 2010; Tarnowski et al., 2017; Wang et al., 2011b; Wheeler et al., 2017; Wood et al., 2017; Wu et al., 2016; Wu and Pankow, 2018; Xue et al., 2018; Zhang et al., 2014b; Zhao et al., 2014, 2017;

Zheng et al., 2015)

***LEPR***  
**Leptin Receptor**  
rs1137101

missense variant  
triglycerides, gestational diabetes mellitus, fasting glucose, BMI, cardiovascular disease, preterm birth, type-2 diabetes, plasma lipids

(Angheben-Oliveira et al., 2017; Ben Ali et al., 2009; Furusawa et al., 2010; Gregoor et al., 2009; Kasim et al., 2016; Mahmoudi et al., 2016; Manriquez et al., 2018; Queiroz et al., 2015; Rojano-Rodriguez et al., 2016; Salem et al., 2016; Tabassum et al., 2012; Urbanek et al., 2012; Wu and Sun, 2017; Yang et al., 2016; Zayani et al., 2017)

***HMGCR***  
**3-Hydroxy-3-Methylglutaryl-CoA Reductase**  
rs2303152

intron variant  
intron variant  
preterm delivery  
HDL cholesterol, LDL cholesterol, triglycerides, fasting blood glucose

(Bream et al., 2013; Steffen et al., 2007)

(Kathiresan et al., 2008; Keebler Mary E. et al., 2009; Kim et al., 2011)

***DHCR24* 24-Hydrocholesterol Reductase**  
rs2274941

non coding transcript exon variant  
prematurity

(Steffen et al., 2007)

***DHCR7***  
**7-Dehydrocholesterol Reductase**

rs1630498  
rs2002064  
intron variant  
intron variant  
low birth weight, prematurity  
birth weight

(Bream et al., 2013; Steffen et al., 2007)  
(Steffen et al., 2007)

***PCSK9***  
**Proprotein Convertase**

**Subtilisin/Kexin Type 9**

rs11591147

missense variant

LDL cholesterol, total cholesterol, cardiovascular disease, fasting glucose, type-2 diabetes

(Chasman et al., 2012; Feng et al., 2017a; Guella et al., 2010; Guo et al., 2016; Kathiresan et al., 2007, 2008; Kettunen et al., 2012, 2016; Klarin et al., 2018; Nagy et al., 2017; Nelson et al., 2017; Pott et al., 2018; Qiu et al., 2017; Rasmussen-Torvik et al., 2012; Schmidt et al., 2017; Smith et al., 2010; Southam et al., 2017; Spracklen et al., 2017; Surakka et al., 2015; Tsai et al., 2015; van der Harst and Verweij, 2018)

***CETP*****Cholesteryl Ester Transfer Protein**

rs1800775

regulatory region variant

HDL cholesterol, LDL cholesterol, total cholesterol, triglycerides, coronary artery disease, metabolic syndrome, dyslipidemia

(Andaleon et al., 2018; Barbosa et al., 2012; Boes et al., 2009; Chasman et al., 2009; Ganesan et al., 2016; Guo et al., 2015b; Hamrefors et al., 2010; Hebbar et al., 2017; Hou et al., 2017; Hu et al., 2016; Kathiresan et al., 2007, 2008; Kenny et al., 2011; Ma et al., 2010; Murray et al., 2009; Nie et al., 2017; Ridker et al., 2009; Ronald et al., 2009; Sabatti et al., 2009; Saxena et al., 2007, 200; Shin et al., 2014; Wakil et al., 2016; Webb et al., 2017; Winkler et al., 2015; Wu et al., 2013)

***LCAT*****Lecithin-Cholesterol Acyltransferase**

rs1109166

3' UTR

gestational age, HDL cholesterol

(Spracklen et al., 2017; Steffen et al., 2007)

***LIPC*****Lipase C, Hepatic type**

rs6083

missense variant

gestational age, preterm prelabor rapture of

(Romero et al., 2010; Steffen et al., 2007;

rs1800588

intron variant

membranes, HDL cholesterol  
total cholesterol, HDL cholesterol,  
triglyceride levels, LDL cholesterol,  
cardiovascular disease, hypertriglyceridemia

Yang et al., 2010)  
(Ahmad et al., 2011; Ayyappa et al.,  
2013; Fan et al., 2009; Guardiola et al.,  
2015; Hamrefors et al., 2010; Hodoglugil  
et al., 2010; Kanai et al., 2018;  
Kathiresan et al., 2008; Liu et al., 2011;  
Lu et al., 2008, 2016a; Nie et al., 2017;  
Ríos-González et al., 2014; Rudkowska  
et al., 2013; Sabatti et al., 2009;  
Spracklen et al., 2017; Villard et al.,  
2013; White et al., 2015; Yang et al.,  
2010)

***LIPG*****Lipase G, endothelial type**

rs2156552

intergenic variant

HDL cholesterol, ischemic stroke

(Carty Cara L. et al., 2012; Gaulton et al.,  
2008; Jeemon et al., 2011; Kathiresan et  
al., 2008; Keebler Mary E. et al., 2009;  
Khetarpal et al., 2011; Klarin et al., 2018;  
Ma et al., 2010; Murray et al., 2009;  
Spracklen et al., 2017; Waterworth et al.,  
2010)

***LPL*****Lipoprotein Lipase**

rs328

stop variant

HDL cholesterol, triglycerides, blood  
pressure, coronary heart disease

(Ayyappa et al., 2017; Chen et al., 2009;  
De Castro-Orós Isabel et al., 2014;  
Drenos et al., 2009; Dumitrescu et al.,  
2011; Emamian et al., 2015; Garcia-Rios  
et al., 2011; Guardiola et al., 2015;  
Jeemon et al., 2011; Kathiresan et al.,  
2007, 2008; Keebler Mary E. et al., 2009;  
Kurano et al., 2016; Larifla et al., 2016;  
Legry et al., 2011; Murray et al., 2009;  
Pirim et al., 2015; Ronald et al., 2009;  
Sabatti et al., 2009; Sagoo et al., 2008;  
Shahid et al., 2017; Tan et al., 2012;  
Tang et al., 2010; Webster et al., 2009;

***ABCA1*****ATP Binding Cassette  
Subfamily A Member 1**

rs2066716

missense variant  
intron variantprematurity, gestational age, birth weight  
HDL cholesterol, overweight, obesity

White et al., 2015; Yeo et al., 2017; Yue et al., 2017; Zhou et al., 2013)

rs4149313

missense variant

gestational age

(Steffen et al., 2007)

(Jeemon et al., 2011; Kathiresan et al., 2008; Kong et al., 2015; Sabatti et al., 2009; Waterworth et al., 2010; Yao et al., 2016a, 2016b)

(Steffen et al., 2007)

***LDLR*****Low Density Lipoprotein  
Receptor**

rs6511720

intron variant

LDL cholesterol, total cholesterol, coronary artery disease, hypertension

(Chasman et al., 2008, 2009; Consortium, 2011; Ding and Kullo, 2009; Elbers et al., 2012; Feng et al., 2017b; Grallert et al., 2012; Gupta et al., 2010; Inouye et al., 2012; Kathiresan et al., 2008, 2009; Keebler Mary E. et al., 2009; Lettre et al., 2011; Linsel-Nitschke et al., 2010; Middelberg et al., 2011; Nelson et al., 2017; Rafiq et al., 2012; Ronald et al., 2009; Shetty Priya B. et al., 2015; Teslovich et al., 2010; van der Harst and Verweij, 2018; Willer et al., 2008; Ye et al., 2014)

***APOB*****Apolipoprotein B**

rs693

synonymous  
variant

LDL cholesterol, HDL cholesterol, triglycerides, total cholesterol, ischemic stroke, coronary heart disease

(Au et al., 2017; Aulchenko et al., 2009; Chen et al., 2016; Haas Blake E. et al., 2011; Hamrefors et al., 2010; Hubacek et al., 2017; Kathiresan et al., 2008; Murray et al., 2009; Niu et al., 2017; Park et al., 2011; Rodrigues et al., 2013; Sabatti et al., 2009; Sandhu et al., 2008; Saxena et

***APOA1*****Apolipoprotein A1**

rs28927680

3' UTR

triglycerides, total cholesterol

rs5070

intron variant

prematurity, HDL cholesterol

al., 2007; Shirts et al., 2011; Takeuchi et al., 2012; Walker et al., 2011; Xiao et al., 2017)

***APOE*****Apolipoprotein**

rs405509

regulatory region variant

prematurity, gestational age, birth weight, serum fasting insulin, overweight

rs7412

missense variant

gestational age, LDL cholesterol, total cholesterol, HDL cholesterol, coronary artery disease, premature coronary artery disease, dyslipidemia, type-2 diabetes

(Jeemon et al., 2011; Kathiresan et al., 2008; Sabatti et al., 2009)  
(Rudkowska et al., 2013; Steffen et al., 2007)

***APOC1*****Apolipoprotein C1**

rs4420638

intergenic variant

LDL cholesterol, HDL cholesterol, total cholesterol, triglycerides, coronary artery disease, c-reactive protein, BMI, type-2 diabetes

(Clark et al., 2009; Edwards et al., 2011; Komurcu-Bayrak et al., 2011; Steffen et al., 2007)  
(Alharbi et al., 2014; Ansari et al., 2017; Barbosa et al., 2012; Cahua-Pablo et al., 2016; Chasman et al., 2012; Futema et al., 2015; Hanh et al., 2016; Mazzotti et al., 2014; Ripatti et al., 2016; Smith et al., 2010; Smolková et al., 2015; Steffen et al., 2007; Surakka et al., 2015; Takeuchi et al., 2012; Tejedor et al., 2014; van der Harst and Verweij, 2018; Wu et al., 2013; Zhu et al., 2017b)

(Adeyemo et al., 2012; Aslibekyan et al., 2012; Burkhardt et al., 2008; Burkhardt Ralph et al., 2008; Chung et al., 2014; Dehghan Abbas et al., 2011; Deshmukh et al., 2012; Elliott et al., 2009; Hubacek et al., 2017; Kathiresan et al., 2008; Keller et al., 2013; Kenny et al., 2011; Kim et al., 2017b; Li et al., 2017; Ligthart et al., 2016; Liu et al., 2011; Lu et al., 2016b; Middelberg et al., 2011;

***PNPLA3***  
**Patatin Like Phospholipase  
 Domain Containing 3**

rs738409

missense variant

non-alcoholic fatty liver disease  
 pathogenesis, total cholesterol, triglycerides,  
 insulin resistance BMI, gestational diabetes,  
 metabolic syndrome

Mohlke et al., 2008; Nikpay et al., 2015;  
 Okada et al., 2011; Park et al., 2011;  
 Sandhu et al., 2008; Saxena et al., 2007;  
 Shirali et al., 2016, 201; Shirts et al.,  
 2011; Spracklen et al., 2017; Teslovich et  
 al., 2010; Varga et al., 2014; Wallace et  
 al., 2008; Waterworth et al., 2010; Willer  
 et al., 2013; Winkler et al., 2015; Zhao et  
 al., 2017)

(Akuta et al., 2016; Alam et al., 2017;  
 Ali et al., 2016; Atkinson et al., 2017;  
 Basantani et al., 2011; Bhatt et al., 2013;  
 Bo et al., 2015; Buch et al., 2015; Burza  
 et al., 2014; Cai et al., 2011; Chalasani et  
 al., 2010; Chamorro et al., 2014; Chan et  
 al., 2017; Cox et al., 2011; Falleti et al.,  
 2016; Fan et al., 2016; Flores et al.,  
 2016; Gao et al., 2017; Goran et al.,  
 2010; Graff et al., 2013; Guichelaar et  
 al., 2013; Guyot et al., 2013; Hassan et  
 al., 2013; Hernaez et al., 2013; Hotta et  
 al., 2010; Huang et al., 2015a, 3, 2017, 3;  
 Jiménez-Sousa et al., 2016; Johansson et  
 al., 2008; Kantartzis et al., 2009;  
 Kawaguchi et al., 2012; Kim et al., 2018,  
 3; Kitamoto et al., 2015; Klarin et al.,  
 2018; Kotronen et al., 2009; Kovac and  
 Rozman, 2015; Krawczyk et al., 2011,  
 2017; Kupcinskas et al., 2017; Lee et al.,  
 2014; Lin et al., 2011, 2014; Liu et al.,  
 2014a; Mancina et al., 2016; Mangge et  
 al., 2015; Miyaaki et al., 2018; Mondul  
 et al., 2015; Moritou et al., 2013; Oniki

**TM6SF2**  
**Transmembrane 6  
Superfamily Member 2**

rs58542926

missense variant

non-alcoholic fatty liver disease  
pathogenesis, triglycerides, lipid  
abnormalities, total cholesterol

et al., 2015; Pan et al., 2015; Petta et al., 2012, 2016; Pirazzi et al., 2012; Pontoriero et al., 2015; Rausch et al., 2016; Romeo et al., 2008, 2010a, 2010b; Rüegeger et al., 2015; Salameh et al., 2016; Santoro et al., 2010; Sato et al., 2014; Scheiner et al., 2015; Seko et al., 2018; Sevastianova et al., 2011; Shang et al., 2015; Shen et al., 2014, 2015a; Singal et al., 2014; Smagris et al., 2015; Sookoian et al., 2009a; Sookoian and Pirola, 2016; Speliotes et al., 2010, 2011; Stickel et al., 2011; Stojkovic et al., 2014; Tai et al., 2015; Takeuchi et al., 2013; Tang et al., 2015; Trepo et al., 2012; Trépo et al., 2011a, 2011b, 2014; Ueyama et al., 2016; Uygun et al., 2017; Valenti et al., 2010a, 2010b, 2012; Verrijken et al., 2013; Vespasiani-Gentilucci et al., 2016; Viganò et al., 2013; Viitasalo et al., 2015; Wagenknecht et al., 2011; Wang et al., 2016; Xia et al., 2016; Xu et al., 2015; Yasui et al., 2015; Zain et al., 2012; Zhang et al., 2014a, 2015)

(Boonvisut et al., 2016; Chen et al., 2015, 2; Dongiovanni et al., 2015; Ehrhardt et al., 2017; Eslam et al., 2016; Falleti et al., 2016; Goffredo et al., 2016; Kanai et al., 2018; Kanth et al., 2016; Kim et al., 2017a; Liu et al., 2014b; Petta et al., 2016; Pirola and Sookoian, 2015; Sookoian et al., 2015, 2016; Surakka et al., 2015; Tang et al., 2015; Wang et al.,

***NCAN*****Neurocan**

rs16996148

intergenic variant

LDL cholesterol, HDL cholesterol,  
triglycerides, coronary heart disease

2016)

(Kathiresan et al., 2008; Willer et al.,  
2008)***TRIB1*****G-Protein-Coupled Receptor-  
Induced Gene 2 Protein**

rs17321515

intron variant

LDL cholesterol, HDL cholesterol,  
triglycerides,  
coronary heart disease(Aung et al., 2011; Dastani et al., 2012;  
Hegele et al., 2009; Huang et al., 2016;  
Kathiresan et al., 2008; Keebler Mary E.  
et al., 2009; Mohlke et al., 2008; Ollila et  
al., 2012; Park et al., 2011; Wang et al.,  
2015a; Willer et al., 2008; Zhou et al.,  
2011, 2013)***ANGPTL3*****Angiopoietin Like 3**

rs12130333

intergenic variant

hypertriglyceridemia, triglycerides

(Hegele et al., 2009; Kathiresan et al.,  
2008; Wang et al., 2008)***IGF2*****Insulin like growth factor 2**

rs74050124

3' UTR

pregnancy complications

(Queiroz et al., 2015, 2)

\*rs74050124 and rs680 are in LD

Supplementary Table S3: p-values for the association between 2<sup>nd</sup> trimester lipid levels and SNPs in lipid and circadian genes.

<b>SNPs</b>	<b>Core circadian regulating genes</b>	<b>CHOL</b>	<b>HDL</b>	<b>LDL</b>	<b>TG</b>
rs3749474	<i>CLOCK</i>	0.86	$5.37 \times 10^{-2}$	0.79	$8.70 \times 10^{-4}$
rs4580704	<i>CLOCK</i>	0.31	$1.93 \times 10^{-3}$	0.63	$1.02 \times 10^{-2}$
rs1464490	<i>CLOCK</i>	0.85	$4.94 \times 10^{-2}$	0.75	$9.72 \times 10^{-4}$
rs6843722	<i>CLOCK</i>	0.73	$8.07 \times 10^{-3}$	0.83	$4.43 \times 10^{-4}$
rs6850524	<i>CLOCK</i>	0.44	$2.27 \times 10^{-4}$	0.54	$2.05 \times 10^{-3}$
rs4864548	<i>CLOCK</i>	0.89	$4.14 \times 10^{-2}$	0.73	$8.93 \times 10^{-4}$
rs1801260	<i>CLOCK</i>	0.42	0.40	$9.28 \times 10^{-2}$	0.92
rs2278749	<i>ARNTL</i>	0.67	$3.56 \times 10^{-3}$	0.82	$1.50 \times 10^{-3}$
rs6486121	<i>ARNTL</i>	$6.21 \times 10^{-3}$	0.93	$6.42 \times 10^{-3}$	0.50
rs7950226	<i>ARNTL</i>	0.62	$6.88 \times 10^{-3}$	0.85	0.21
rs11022775	<i>ARNTL</i>	0.56	0.25	0.30	$6.35 \times 10^{-2}$
rs2585405	<i>PER1</i>	$2.31 \times 10^{-2}$	0.80	$1.06 \times 10^{-2}$	0.76
rs3027178	<i>PER1</i>	0.17	0.42	0.14	$2.97 \times 10^{-3}$
rs2304672	<i>PER2</i>	0.65	0.45	0.69	0.99
rs56013859	<i>PER2</i>	0.11	$6.04 \times 10^{-2}$	$2.20 \times 10^{-2}$	$5.08 \times 10^{-2}$
rs7602358	<i>PER3</i>	$2.51 \times 10^{-2}$	$1.72 \times 10^{-3}$	0.18	0.12

rs228669	<i>PER3</i>	0.50	0.19	0.81	$1.24 \times 10^{-6}$
rs2640908	<i>PER3</i>	$3.77 \times 10^{-4}$	0.28	$1.78 \times 10^{-4}$	0.64
rs2287161	<i>CRY1</i>	0.53	0.95	0.81	0.95
rs3809236	<i>CRY1</i>	0.69	0.52	0.52	$3.56 \times 10^{-2}$
rs12315175	<i>CRY1</i>	$9.28 \times 10^{-2}$	0.38	0.41	0.15
rs2292912	<i>CRY2</i>	$4.84 \times 10^{-2}$	0.85	$3.92 \times 10^{-3}$	0.25
rs11605924	<i>CRY2</i>	N/A	N/A	N/A	N/A
rs2305160	<i>NPAS2</i>	0.21	0.23	0.11	$5.46 \times 10^{-2}$
rs11541353	<i>NPAS2</i>	0.84	0.30	0.22	$4.03 \times 10^{-2}$
<b>SNPs</b>	<b>Circadian-related and lipid-related genes</b>	<b>CHOL</b>	<b>HDL</b>	<b>LDL</b>	<b>TG</b>
rs12413112	<i>SIRT1</i>	0.79	$6.17 \times 10^{-2}$	0.96	$5.25 \times 10^{-3}$
rs3758391	<i>SIRT1</i>	$1.83 \times 10^{-2}$	0.39	$7.73 \times 10^{-3}$	$3.84 \times 10^{-2}$
rs2273773	<i>SIRT1</i>	$7.22 \times 10^{-2}$	0.78	$5.10 \times 10^{-2}$	0.40
rs10997860	<i>SIRT1</i>	N/A	N/A	N/A	N/A
rs646776	<i>CELSR2-PSRC1-SORT1</i>	$2.35 \times 10^{-5}$	$5.35 \times 10^{-4}$	$5.10 \times 10^{-11}$	0.19
rs599839	<i>CELSR2-PSRC1-SORT1</i>	$2.83 \times 10^{-5}$	$7.10 \times 10^{-4}$	$1.16 \times 10^{-10}$	$5.38 \times 10^{-2}$
rs28932472	<i>POMC</i>	0.13	0.38	0.65	0.39
rs1800206	<i>PPARA</i>	0.28	0.49	0.61	$4.35 \times 10^{-3}$

rs7638903	<i>PPARG</i>	0.92	0.16	0.88	0.29
rs1801282	<i>PPARG</i>	0.95	0.60	0.94	0.45
rs12640088	<i>PPARGC1A</i>	0.39	0.31	0.16	$1.10 \times 10^{-2}$
rs8192678	<i>PPARGC1A</i>	0.78	0.12	0.35	0.82
rs251464	<i>PPARGC1B</i>	0.24	0.95	$3.30 \times 10^{-2}$	$4.40 \times 10^{-2}$
rs17383291	<i>VDR</i>	0.45	0.21	0.56	$4.17 \times 10^{-2}$
rs2228570	<i>VDR</i>	0.58	$2.87 \times 10^{-2}$	0.44	$4.66 \times 10^{-2}$
rs10507486	<i>FOXO1</i>	0.55	0.47	0.25	0.15
rs2297627	<i>FOXO1</i>	0.27	0.87	0.16	0.10
rs10830963	<i>MTNR1B</i>	0.69	0.86	0.76	0.56
rs1137101	<i>LEPR</i>	0.10	0.39	0.44	0.70
rs2303152	<i>HMGCR</i>	0.56	0.20	0.24	0.51
rs12654264	<i>HMGCR</i>	N/A	N/A	N/A	N/A
rs2274941	<i>DHCR24</i>	N/A	N/A	N/A	N/A
rs1630498	<i>DHCR7</i>	0.73	0.92	0.60	0.57
rs2002064	<i>DHCR7</i>	0.86	0.92	0.65	0.68
rs11591147	<i>PCSK9</i>	0.47	0.27	0.98	0.75
rs1800775	<i>CETP</i>	0.10	0.12	0.54	$4.27 \times 10^{-2}$

rs1109166	<i>LCAT</i>	0.99	$3.15 \times 10^{-2}$	0.85	0.28
rs6083	<i>LIPC</i>	$8.86 \times 10^{-2}$	0.74	$2.75 \times 10^{-2}$	0.80
rs1800588	<i>LIPC</i>	0.22	$5.57 \times 10^{-2}$	$7.63 \times 10^{-3}$	$2.16 \times 10^{-4}$
rs2156552	<i>LIPG</i>	0.25	0.63	0.34	0.12
rs328	<i>LPL</i>	$6.65 \times 10^{-3}$	$5.52 \times 10^{-2}$	$1.13 \times 10^{-2}$	$1.33 \times 10^{-3}$
rs2066716	<i>ABCA1</i>	0.13	0.996	0.16	$1.81 \times 10^{-4}$
rs3890182	<i>ABCA1</i>	0.69	0.73	0.32	$1.63 \times 10^{-2}$
rs4149313	<i>ABCA1</i>	0.18	0.42	$2.74 \times 10^{-3}$	0.49
rs6511720	<i>LDLR</i>	0.92	0.55	0.92	0.35
rs693	<i>APOB</i>	$8.46 \times 10^{-2}$	0.83	$3.70 \times 10^{-3}$	0.70
rs28927680	<i>APOA1</i>	0.44	0.13	0.18	$6.66 \times 10^{-3}$
rs5070	<i>APOA1</i>	0.35	0.92	$4.83 \times 10^{-2}$	$1.24 \times 10^{-3}$
rs405509	<i>APOE</i>	$3.39 \times 10^{-3}$	$6.94 \times 10^{-2}$	$1.63 \times 10^{-3}$	0.31
rs7412	<i>APOE</i>	$2.81 \times 10^{-6}$	$6.35 \times 10^{-5}$	$< 1.0 \times 10^{-12}$	0.10
rs4420638	<i>APOC1</i>	0.14	0.52	$9.52 \times 10^{-3}$	0.23
rs738409	<i>PNPLA3</i>	0.33	$9.81 \times 10^{-5}$	0.54	$3.56 \times 10^{-6}$
rs58542926	<i>TM6SF2</i>	0.12	0.89	$4.22 \times 10^{-2}$	0.89
rs16996148	<i>NCAN</i>	0.76	0.49	0.81	0.38

rs17321515	<i>TRIB1</i>	0.55	$6.69 \times 10^{-3}$	0.89	0.26
rs12130333	<i>ANGPTL3</i>	0.71	0.18	0.33	$9.71 \times 10^{-2}$
rs74050124	<i>IGF2</i>	0.91	$3.38 \times 10^{-2}$	0.91	$4.73 \times 10^{-2}$

All data presented are unadjusted and represent the p-value for the association between each individual lipid (as the outcome) with a single candidate SNP.

Supplementary Table S4: Minor allele frequencies for candidate SNPs and differences in frequency by race.

			Minor Allele Frequency				
	SNPs	Minor Allele	Overall	Hispanic	White Non-Hispanic	Asian	p-value
<i>ABCA1</i>	rs2066716	A	0.17	0.16	0.18	0.18	0.57
<i>ABCA1</i>	rs3890182	A	0.09	0.10	0.08	0.11	0.41
<i>ABCA1</i>	rs4149313	G	0.26	0.25	0.30	0.18	0.003
<i>ANGPTL3</i>	rs12130333	T	0.20	0.19	0.19	0.23	0.40
<i>APOA1</i>	rs28927680	G	0.10	0.09	0.11	0.09	0.35
<i>APOA1</i>	rs5070	A	0.47	0.47	0.49	0.44	0.45
<i>APOB</i>	rs693	T	0.38	0.38	0.37	0.40	0.76
<i>APOC1</i>	rs4420638	G	0.13	0.14	0.13	0.10	0.34
<i>APOE</i>	rs405509	C	0.47	0.46	0.46	0.47	0.96
<i>APOE</i>	rs7412	T	0.05	0.04	0.06	0.04	0.24
<i>ARNTL</i>	rs2278749	A	0.22	0.25	0.18	0.22	0.02
<i>ARNTL</i>	rs6486121	T*	0.49	0.49	0.49	0.47	0.59
<i>ARNTL</i>	rs7950226	G*	0.47	0.45	0.50	0.49	0.08
<i>ARNTL</i>	rs11022775	T	0.11	0.10	0.12	0.10	0.53
<i>CELSR2-PSRC1-SORT1</i>	rs646776	G	0.19	0.18	0.18	0.22	0.31
<i>CELSR2-PSRC1-SORT1</i>	rs599839	T	0.21	0.20	0.21	0.25	0.34
<i>CETP</i>	rs1800775	C	0.47	0.45	0.48	0.46	0.68
<i>CLOCK</i>	rs3749474	T	0.45	0.47	0.43	0.43	0.18
<i>CLOCK</i>	rs4580704	G	0.31	0.31	0.33	0.32	0.58
<i>CLOCK</i>	rs1464490	C	0.45	0.47	0.43	0.43	0.18
<i>CLOCK</i>	rs6843722	C	0.43	0.44	0.41	0.41	0.30
<i>CLOCK</i>	rs6850524	C	0.36	0.35	0.39	0.38	0.34

<i>CLOCK</i>	rs4864548	A	0.45	0.47	0.42	0.43	0.15
<i>CLOCK</i>	rs1801260	C	0.21	0.21	0.22	0.23	0.73
<i>CRY1</i>	rs3809236	T	0.12	0.12	0.11	0.13	0.73
<i>CRY1</i>	rs12315175	C	0.19	0.21	0.18	0.16	0.22
<i>CRY1</i>	rs2287161	C	0.45	0.44	0.47	0.42	0.26
<i>CRY2</i>	rs2292912	C	0.34	0.33	0.36	0.30	0.25
<i>DHCR7</i>	rs1630498	G	0.24	0.25	0.23	0.25	0.81
<i>DHCR7</i>	rs2002064	C	0.25	0.25	0.24	0.25	0.85
<i>FOXO1</i>	rs10507486	T	0.21	0.21	0.20	0.23	0.54
<i>FOXO1</i>	rs2297627	C	0.43	0.44	0.43	0.38	0.30
<i>HMGCR</i>	rs2303152	T	0.07	0.07	0.07	0.12	0.03
<i>IGF2</i>	rs74050124	A	0.01	0.01	0.01	0.01	0.86
<i>LCAT</i>	rs1109166	G	0.18	0.18	0.17	0.19	0.73
<i>LDLR</i>	rs6511720	T	0.09	0.09	0.09	0.11	0.59
<i>LEPR</i>	rs1137101	A*	0.48	0.47	0.47	0.44	0.06
<i>LIPC</i>	rs6083	A*	0.49	0.50	0.45	0.44	0.02
<i>LIPC</i>	rs1800588	T	0.40	0.40	0.42	0.36	0.26
<i>LIPG</i>	rs2156552	A	0.12	0.11	0.13	0.12	0.33
<i>LPL</i>	rs328	G	0.09	0.09	0.09	0.07	0.68
<i>MTNR1B</i>	rs10830963	G	0.31	0.32	0.30	0.30	0.59
<i>NCAN</i>	rs16996148	T	0.08	0.07	0.09	0.09	0.18
<i>NPAS2</i>	rs2305160	T	0.27	0.26	0.27	0.30	0.49
<i>NPAS2</i>	rs11541353	A	0.11	0.11	0.11	0.11	0.98
<i>PCSK9</i>	rs11591147	T	0.007	0.005	0.008	0.01	0.67
<i>PER1</i>	rs2585405	C	0.19	0.20	0.19	0.15	0.32
<i>PER1</i>	rs3027178	C	0.39	0.38	0.42	0.35	0.20
<i>PER2</i>	rs2304672	C	0.05	0.05	0.05	0.07	0.29

<i>PER2</i>	rs56013859	C	0.14	0.14	0.13	0.16	0.62
<i>PER2</i>	rs7602358	G	0.18	0.17	0.19	0.21	0.22
<i>PER3</i>	rs228669	A	0.21	0.21	0.23	0.11	0.003
<i>PER3</i>	rs2640908	T	0.22	0.22	0.24	0.20	0.57
<i>PNPLA3</i>	rs738409	G	0.37	0.38	0.33	0.40	0.08
<i>POMC</i>	rs28932472	C	0.21	0.22	0.20	0.20	0.48
<i>PPARA</i>	rs1800206	G	0.04	0.05	0.04	0.03	0.40
<i>PPARG</i>	rs1801282	G	0.10	0.11	0.09	0.08	0.35
<i>PPARG</i>	rs7638903	A	0.10	0.11	0.09	0.09	0.29
<i>PPARGC1A</i>	rs8192678	A	0.31	0.32	0.31	0.34	0.75
<i>PPARGC1A</i>	rs12640088	C	0.10	0.10	0.10	0.09	0.78
<i>PPARGC1B</i>	rs251464	C	0.36	0.36	0.36	0.40	0.52
<i>SIRT1</i>	rs12413112	A	0.16	0.16	0.16	0.16	0.99
<i>SIRT1</i>	rs3758391	T*	0.50	0.49	0.49	0.49	0.89
<i>SIRT1</i>	rs2273773	C	0.14	0.14	0.14	0.12	0.67
<i>TM6SF2</i>	rs58542926	T	0.06	0.05	0.07	0.06	0.39
<i>TRIB1</i>	rs17321515	G	0.44	0.43	0.47	0.43	0.23
<i>VDR</i>	rs17383291	G	0.43	0.42	0.43	0.43	0.91
<i>VDR</i>	rs2228570	A	0.44	0.45	0.44	0.34	0.02

\*Minor allele for rs6486121 in White-Not Hispanic is C; Minor allele for rs7950226 in White-Not Hispanic is A; Minor allele for rs11605924 in Hispanic is A; Minor allele for rs3758391 in Hispanic is C; Minor allele for rs10997860 in White-Not Hispanic is T; Minor allele for rs1137101 in Asian is G; Minor allele for rs6083 in Asian is G.

MAF, minor allele frequency as calculated from the study population.

P-value represents the chi-square test for differences in MAF by race.

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