



Birchula™

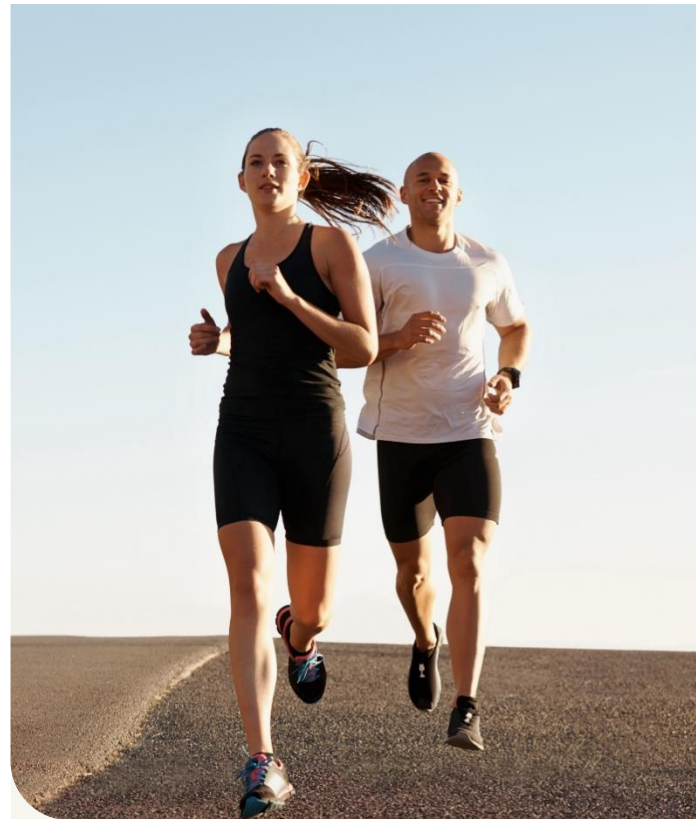
Birch bark extract for cardio metabolic disorders

Birchula™ birch bark extract contains highly effective triterpenes, Betulin, Betulinic Acid, Betulin Caffeates, and lupeol. These compounds are scientifically proven to reduce inflammation, improve cardiometabolic health, reduce cholesterol, improve glucose uptake and control blood sugar.

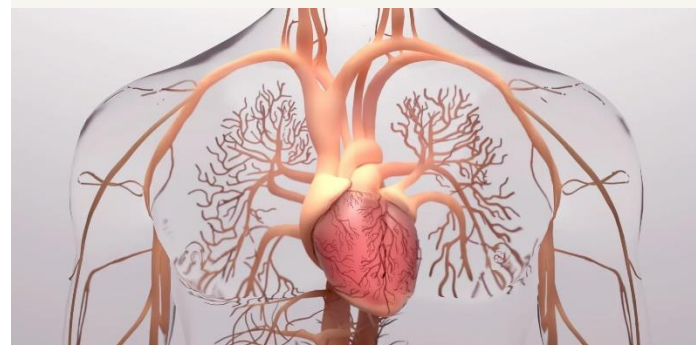
Scientifically proven benefits of birch bark extract

- Reduces inflammation B, BA, L
- Triggers the release of GLP-1 BA
- Controls carbohydrate metabolism BBE
- Anti-oxidant B, BA, L
- Controls blood sugars B, BA, L
- Improves glucose uptake B
- Reduces cholesterol B, L
- Lowers cardiac lipid levels B

(B) Betulin, (BA) betulinic acid, (L) lupeol, (BBE) birch bark extract



- Birchula™ Birch Bark Extract comes from the outer bark of the North American birch tree (*Betula Papyrifera*)
- Upcycled from the waste bark of the paper and timber industries, not peeled from live trees.
- Obtained from sustainably harvested and managed forests certified through the Sustainable Forestry Initiative (SFI)
- Proprietary technology for safe cGMP processing
- Identity preserved - Can trace the supply chain back to when the tree was harvested



The Actives Factory

1313 Fairgrounds Road #150 Two Harbors, MN 55616
bgarhofer@theactivesfactory.com | 877-723-8852 | theactivesfactory.com

The Actives Factory is a manufacturer of natural extracts and high purity compounds from the North American White Birch for nutraceutical, cosmeceutical, and pharmaceutical use.

There is a distressing worldwide rise in the incidence of Type 2 diabetes and associated. There is compelling evidence that the triterpenes betulin, betulinic acid, and lupeol found in birch bark have beneficial effects on carbohydrate and lipid metabolism, as well as insulin function, dyslipidemia, heart disease, and obesity.

Research and methods of action

Positive impact of birch bark triterpenes on carbohydrate metabolism

- Betulinic Acid inhibits PTP1B improving sensitivity to leptin, an anti-obesity hormone.⁽¹⁾
- Betulinic acid promotes energy expenditure by triggering the release of GLP-1 (10,11)
- Betulin stimulates glucose uptake at about 20% of the effect of Insulin. (12)
- Betulinic Acid impedes the breakdown of complex carbohydrates into simple sugars (12,13)

Birch bark triterpenes affect lipid metabolism and modification

- Betulinic Acid restricts fatty acid uptake from food²
- Lupeol inhibits the fatty acid synthase enzyme³
- Betulinic Acid promotes lipid breakdown by inhibiting cAMP-dependent phosphodiesterase⁴
- Betulin and Betulinic Acid inhibits sterol regulatory element binding proteins (SREBP) improving hyperlipidemia, insulin resistance and reducing atherosclerotic plaque.^{5,6}
- Betulinic acid promotes the efflux of lipid from these foam cells reducing the risk of atherosclerosis⁷
- Birch bark triterpenes inhibit the oxidation of “bad” low-density lipoprotein to form oxLDL⁸, which contributes to atherosclerotic plaque formation⁹

1 Choi YJ, Park SY, Kim JY, et al. Combined treatment of betulinic acid, a PTP1B inhibitor, with Orthosiphon stamineus extract decreases body weight in high-fat-fed mice. *J Med Food*. 2013;16(1):2-8.

2 Jang DS, Lee GY, Kim J, et al. A new pancreatic lipase inhibitor isolated from the roots of *Actinidia arguta*. *Arch Pharm Res*. 2008;31(5):666-670.

3 Ardiansyah, Yamaguchi E, Shirakawa H, et al. Lupeol supplementation improves blood pressure and lipid metabolism parameters in stroke-prone spontaneously hypertensive rats. *Biosci Biotechnol Biochem*. 2012;76(1):183-185.

4 Kim J, Lee YS, Kim CS, Kim JS. Betulinic acid has an inhibitory effect on pancreatic lipase and induces adipocyte lipolysis. *Phytother Res*. 2012;26(7):1103-1106.

5 Tang JJ, Li JG, Qi W, et al. Inhibition of SREBP by a small molecule, betulin, improves hyperlipidemia and insulin resistance and reduces atherosclerotic plaques. *Cell Metab*. 2011;13(1):44-56.

6 Quan HY, Kim do Y, Kim SJ, Jo HK, Kim GW, Chung SH. Betulinic acid alleviates non-alcoholic fatty liver by inhibiting SREBP1 activity via the AMPK-mTOR-SREBP signaling pathway. *Biochem Pharmacol*. 2013;85(9):1330-1340.

7 Zhao GJ, Tang SL, Lv YC, et al. Antagonism of betulinic acid on LPS-mediated inhibition of ABCA1 and cholesterol efflux through inhibiting nuclear factor-kappaB signaling pathway and miR-33 expression. *PLoS One*. 2013;8(9):e74782.

8 Andrikopoulos NK, Kaliora AC, Assimopoulou AN, Papapeorgiou VP. Biological activity of some naturally occurring resins, gums and pigments against in vitro LDL oxidation. *Phytother Res*. 2003;17(5):501-507.

9 Pirillo A, Norata GD, Catapano AL. LOX-1, OxLDL, and Atherosclerosis. *Mediators Inflamm*. 2013;2013.

10 Genet C, Strehle A, Schmidt C, et al. Structure-activity relationship study of betulinic acid, a novel and selective TGR5 agonist, and its synthetic derivatives: potential impact in diabetes. *J Med Chem*. 2010;53(1):178-190.

11 Inoue T, Wang JH, Higashiyama M, et al. Dipeptidyl peptidase IV inhibition potentiates amino acid- and bile acid-induced bicarbonate secretion in rat duodenum. *Am J Physiol Gastrointest Liver Physiol*. 2012;303(7):G810-816.

12 Deutschlander MS, Lall N, Van de Venter M, Hussein AA. Hypoglycemic evaluation of a new triterpene and other compounds isolated from *Euclea undulata* Thunb. var. *myrtina* (Ebenaceae) root bark. *J Ethnopharmacol*. 2011;133(3):1091-1095.

13 Kumar S, Kumar V, Prakash O. Enzymes inhibition and antidiabetic effect of isolated constituents from *Dillenia indica*. *Biomed Res Int*. 2013;2013:382063.