

Female Fertility

Folate

Protects genes during rapid cell division which increases likelihood of a healthy embryo (via methylation of DNA); Deficiency raises homocysteine which damages reproductive cells.^{1,2,3,4}

Vitamin B₆ & B₁₂

Both are needed to convert toxic homocysteine to a benign form; Low homocysteine levels linked to a better chance of pregnancy.^{5,6,7,8}

Vitamin C

Increases serum progesterone levels; Induces ovulation in some women; Enhances effect of the fertility drug clomiphene.^{9,10,11,12}

Minerals

Several enzymes needed to protect a woman's reproductive organs (such as superoxide dismutase) are dependent on the trace elements **zinc, copper and magnesium.**

^{22,30,31,32}

Vitamin D

Higher levels linked to better success rates of IVF (in vitro fertilization); Influences production of the sex hormones estradiol and progesterone.^{13,14,15}

Antioxidant Status

Reproductive cells, including embryos, are very susceptible to damage from oxidative stress due to the rapid rate of growth; Low antioxidant status can cause infertility or miscarriage.^{19,22,28,29}

Vitamin E

Protects reproductive cells (follicles); May improve endometrial response (ability of fertilized egg to implant into uterine wall properly) during IVF.^{16,17,18,19}

Cysteine

N-acetyl cysteine can improve ovulation and pregnancy rates in women with infertility due to PCOS (polycystic ovary syndrome) that do not respond to fertility drugs; Improves viability of endometrial cells in vitro; Precursor to glutathione.^{25,26,27}

Glutathione

Protects eggs (fertilized or not) from damage by reactive oxygen species; Protective action of follicle stimulating hormone on embryonic development is due largely to glutathione synthesis.^{22,23,24}

Selenium

Deficiency implicated in miscarriage and infertility; In one trial, 100% of infertile women achieved pregnancy after supplementation.^{20,21}

REFERENCES

- ¹Laanpere M, Altmäe S, Stavreus-Evers A et al. Folate-mediated one-carbon metabolism and its effect on female fertility and pregnancy viability. *Nutr Rev* 2010;68:99-113.
- ²Forges T, Monnier-Barbarino P, Alberto J et al. Impact of folate and homocysteine metabolism on human reproductive health. *Hum Reprod Update* 2007;13:225-238.
- ³Ebisch I, Thomas C, Peters W et al. The importance of folate, zinc and antioxidants in the pathogenesis and prevention of subfertility. *Hum Reprod Update* 2007;13:163-174.
- ⁴Dawson D, Sawers A. Infertility and folate deficiency. *Case Reports. Br J Obstet Gynaecol* 1982;89:678-680.
- ⁵Ocal P, Ersoylu B, Cepni I et al. The association between homocysteine in the follicular fluid with embryo quality and pregnancy rate in assisted reproductive techniques. *J Assist Reprod Genet* 2012;29:299-304.
- ⁶Berker B, Kaya C, Aytac R et al. Homocysteine concentrations in follicular fluid are associated with poor oocyte and embryo qualities in polycystic ovary syndrome patients undergoing assisted reproduction. *Hum Reprod* 2009;24:2293-2302.
- ⁷Ebisch I, Peters W, Thomas C et al. Homocysteine, glutathione and related thiols affect fertility parameters in the (sub)fertile couple. *Hum Reprod* 2006;21:1725-1733.
- ⁸Jerzak M, Putowski L, Baranowski W. Homocysteine level in ovarian follicular fluid or serum as a predictor of successful fertilization. *Gynekol Pol* 2003;74:949-952.
- ⁹Henmi H, Endo T, Kitajima Y et al. Effects of ascorbic acid supplementation on serum progesterone levels in patients with a luteal phase defect. *Fertil Steril* 2003;80:459-461.
- ¹⁰Tannetta D, Sargent I, Linton E et al. Vitamins C and E Inhibit Apoptosis of Cultured Human Term Placenta Trophoblast. *Placenta* 2008;29:680-690.
- ¹¹Luck M, Jeyaseelan I, Scholes R. Ascorbic acid and fertility. *Biol Reprod* 1995;52:262-266.
- ¹²Igarashi M. Augmentative effect of ascorbic acid upon induction of human ovulation in clomiphene-ineffective anovulatory women. *Int J Fertility* 1977;22:168-173.
- ¹³Lerchbaum E, Obermayer-Pietsch B. Vitamin D and fertility: a systematic review. *Eur J Endocrinol* 2012;166:765-778.
- ¹⁴Anagnostis P, Karras S, Goulis D. Vitamin D in human reproduction: a narrative review. *Int J Clin Pract* 2013; Epub ahead of print.
- ¹⁵Ozkan S, Jindal S, Greenfield K et al. Replete vitamin D stores predict reproductive success following in vitro fertilization. *Fertil Steril* 2010;94:1314-1319.
- ¹⁶Cicek N, Eryilmaz O, Sarikaya E et al. Vitamin E effect on controlled ovarian stimulation of unexplained infertile women. *J Assist Reprod Genet* 2012;29:325-328.
- ¹⁷Nugent D, Newton H, Gallivan L et al. Protective effect of vitamin E on ischaemia-reperfusion injury in ovarian grafts. *J Reprod Fertil* 1998;114:341-346.
- ¹⁸Campos C, Ferriani R, Dos Reis R et al. Lipid peroxidation and vitamin E in serum and follicular fluid of infertile women with peritoneal endometriosis submitted to controlled ovarian hyperstimulation: a pilot study. *Fertil Steril* 2008;90:2080-2085.
- ¹⁹Tarin J, Ten J, Vendrell F et al. Effects of maternal ageing and dietary antioxidant supplementation on ovulation, fertilisation and embryo development in vitro in the mouse. *Reprod Nutr Dev* 1998;38:499-508.
- ²⁰Mistry H, Broughton Pipkin F, Redman C et al. Selenium in reproductive health. *Am J Obstet Gynecol* 2012;206:21-30.
- ²¹Howard J, Davies S, Hunnisett A. Red cell magnesium and glutathione peroxidase in infertile women – effects of oral supplementation with magnesium and selenium. *Magnes Res* 1994;7:49-57.
- ²²Fujii J, Iuchi Y, Okada F. Fundamental roles of reactive oxygen species and protective mechanisms in the female reproductive system. *Reprod Biol Endocrinol* 2005;3:43.
- ²³Tsai-Turton M, Luderer U. Opposing effects of glutathione depletion and follicle-stimulating hormone on reactive oxygen species and apoptosis in cultured preovulatory rat follicles. *Endocrinology* 2006;147:1224-1236.
- ²⁴Gardiner C, Salmen J, Brandt C. Glutathione is present in reproductive tract secretions and improves development of mouse embryos after chemically induced glutathione depletion. *Biol Reprod* 1998;59:431-436.
- ²⁵Badawy A, State O, Abdelgawad S. N-acetyl cysteine and clomiphene citrate for induction of ovulation in polycystic ovary syndrome: a cross-over trial. *Acta Obstet Gynecol Scand* 2007;86:218-222.
- ²⁶Rizk A, Bedaiwy M, Al-Inany H. N-acetyl cysteine is a novel adjuvant to clomiphene citrate in clomiphene citrate-resistance patients with polycystic ovary syndrome. *Fertil Steril* 2005;83:367-270.
- ²⁷Estany s, Palacio J, Barnadas R et al. Antioxidant activity of N-acetylcysteine, flavonoids and alpha-tocopherol on endometrial cells in culture. *J Reprod Immunol* 2007;75:1-10.
- ²⁸Agarwal A, Gupta S, Sharma R. Role of oxidative stress in female reproduction. *Reprod Biol Endocrinol* 2005;3:28.
- ²⁹Ruder E, Hartman T, Blumberg J et al. Oxidative stress and antioxidants: exposure and impact on female fertility. *Hum Reprod Update* 2008;14:345-357.
- ³⁰Pathak P, Kapil U. Role of trace elements zinc, copper and magnesium during pregnancy and its outcomes. *Indian J Pediatr* 2004;71:1003-1005.
- ³¹Noda Y, Ota K, Shirasawa T et al. Copper/zinc superoxide dismutase insufficiency impairs progesterone secretion and fertility in female mice. *Biol Reprod* 2012;86:1-8.
- ³²Cetin I et al. Role of micronutrients in the periconceptual period. *Hum Reprod Update* 2010;16:80-95.

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