EXAMINING THE EVIDENCE CLINICAL IMPLICATIONS OF KEY TRIALS

Ryerson AB, Eheman C, Burton J, et al. Symptoms, diagnoses, and time to key diagnostic procedures among older US women with ovarian cancer. Obstet Gynecol. 2007;109: 1053–1061.

Q. Which symptoms are most likely to arouse suspicion of ovarian cancer?

Gynecologic symptoms such as abdominal and pelvic swelling, abnormal bleeding, and genital organ pain were most likely to prompt evaluation for ovarian cancer in this review of Surveillance, Epidemiology, and End Results (SEER) and Medicare data for 3,250 women 65 years and older. Gastrointestinal (GI) symptoms such as nausea and vomiting, constipation, diarrhea, and other digestive disorders were associated with diagnosis at a later stage, whereas gynecologic symptoms were associated with earlier disease.

EXPERT COMMENTARY

Barbara A. Goff, MD, Professor and Director of Gynecologic Oncology, and Adjunct Professor, Department of Medicine, Division of Surgery, University of Washington, Seattle.

A substantial body of research over the past decade has demonstrated that, even in the earliest stages, ovarian cancer pro-

duces symptoms. This study by Ryerson and colleagues is a significant contribution to that body of evidence and underscores the importance of including ovarian cancer in the differential diagnosis when a patient reports gynecologic, GI, or abdominal symptoms.

Early recognition is critical

VEDIO

Because we lack an effective screening test for ovarian cancer, early recognition and prompt evaluation of symptoms may be critical to improve prognosis. The cure rate for early-stage disease is 70% to 90%, compared with 10% to 30% for advanced stages.

This study is consistent with other research demonstrating that more than 80% of women with early-stage ovarian cancer have symptoms months before they are given a diagnosis. Compared with women who have GI symptoms, those who have gynecologic symptoms

Experts agree: Ovarian cancer is not silent

he so-called silent killer isn't, apparently. That's the word from the Gynecologic Cancer Foundation, American Cancer Society, and Society of Gynecologic Oncologists, which jointly issued an evidence-based consensus statement on ovarian cancer symptoms in June. Often referred to as the "silent killer" because of the common belief that there are no warning signs or symptoms, ovarian cancer is the deadliest reproductive cancer.

Barbara A. Goff, MD, who comments here on the study by Ryerson and colleagues, helped formulate the consensus statement, which cites the following symptoms that occur in women with ovarian cancer:

- Bloating
- Pelvic or abdominal pain

- Difficulty eating or feeling full quickly
- Urinary symptoms (urgency or frequency).

A woman who has symptoms almost daily for more than a few weeks should see her gynecologist, and the gynecologist should include ovarian cancer in the differential diagnosis. Ovarian cancer should be suspected if symptoms persist more than 2 to 3 weeks and occur 12 or more times a month.

"This agreement on common symptoms of ovarian cancer hopefully will lead to earlier diagnosis when a cure is more likely," said Dr. Goff. "We know that when women are diagnosed in stage I of the disease, it is 90% curable."

For more on the detection of ovarian cancer, see Dr. Barbieri's editorial on page 8.

CONTINUED

Prenatal 90+DHA

Rx Prenatal Vitamin Tablet and 250 mg DHA Capsule

DESCRIPTION: Citracal® Prenatal 90+DHA is a prescription prenatal/postnatal multivitamin/mineral tablet and a capsule of an essential fatty acid. The prenatal vitamin is a scored, white, oval multivitamin/mineral tablet. The tablet is embossed "CITRACAL" on one side and "PN 90" on the other. The essential fatty acid DHA capsule is clear and contains an amber to light/ dark orange semi-solid mixture.

Each prenatal tablet contains:

Vitamin A (as Palmitate and beta-carotene) .	2700	IU
Vitamin C (Ascorbic acid)	120	mg
Calcium (Calcium citrate)	200	mg
Iron (Carbonyl iron)	90	mg
Vitamin D3 (Cholecalciferol)	400	IU
Vitamin E (dl-alpha tocopheryl acetate)	30	IU
Thiamin (Vitamin B1)	3	mg
Riboflavin (Vitamin B2)	3.4	mg
Niacinamide (Vitamin B3)	20	mg
Vitamin B6 (Pyridoxine HCI)	20	mg
Folic Acid	1	mg
Vitamin B12 (Cyanocobalamin)	12	mcg
Iodine (Potassium iodide)	150	mcg
Zinc (Zinc oxide)	25	mg
Copper (Cupric oxide)	2	mg
Docusate sodium	50	mg

Each DHA gelatin capsule contains:

INDICATIONS: Citracal® Prenatal 90+DHA is a multivitamin/ mineral prescription drug indicated for use in improving the nutritional status of women prior to conception, throughout pregnancy, and in the postnatal period for both lactating and nonlactating mothers. Citracal® Prenatal 90+DHA can also be beneficial in improving the nutritional status of women prior to conception.

CONTRAINDICATIONS: This product is contraindicated in patients with a known hypersensitivity to any of the ingredients.

WARNING: Accidental overdose of iron-containing products is a leading cause of fatal poisoning in children under 6. KEEP THIS PRODUCT OUT OF THE REACH OF CHILDREN. In case of accidental overdose, call a doctor or poison control center immediately.

WARNING: Ingestion of more than 3 grams of omega-3 fatty acids per day has been shown to have potential antithrombotic effects, including an increased bleeding time and INR. Administration of omega-3 fatty acids should be avoided in patients on anticoagulants and in those known to have an inherited or acquired bleeding diathesis.

WARNING: Folic acid alone is improper therapy in the treatment of pernicious anemia and other megaloblastic anemias where vitamin B_{12} is deficient.

PRECAUTIONS: Folic acid in doses above 0.1 mg daily may obscure pernicious anemia in that hematologic remission can occur while neurological manifestations progress.

ADVERSE REACTIONS: Allergic sensitization has been reported following both oral and parenteral administration of folic acid.

CAUTION: Exercise caution to ensure that the prescribed dosage of DHA does not exceed 1 gram (1000 mg) per day.

DOSAGE AND ADMINISTRATION: One tablet and one capsule daily or as directed by a physician.

Store at controlled room temperature.

NOTICE: Contact with moisture can discolor or erode the tablet.

HOW SUPPLIED: Six child-resistant blister packs of 5 tablets and 5 capsules each - NDC 0178-4200-30

U.S. Patent 6,818,228

MISSION PHARMACAL COMPANY San Antonio, TX 78230 1355

PRE-07202

Rev 010060

Trademark of Martek Biosciences Corporation U.S. Patent No. 5,407,957

U.S. Patent No. 5,492,938





underwent abdominal or pelvic imaging and surgical evaluation sooner after reporting symptoms.

Large database and inclusion of ICD-9 data are strengths of this study

This study utilized population data from 13 geographic regions of the country. More important, it evaluated the ICD-9 diagnosis that prompted a visit to a physician or a procedure before ovarian cancer was diagnosed. Because women were unaware that they would subsequently be given a diagnosis of ovarian cancer, the design eliminates recall bias that has been at issue in earlier studies.

ICD-9 data are also a weakness

The fact that ICD-9 diagnostic data for billing purposes were used as a substitute for patient symptoms is a weakness of this study. Physicians frequently report only one ICD-9 diagnosis per encounter, so this study may significantly underreport the symptoms experienced by women with ovarian cancer.

The fact that this study included only women older than 65 years is another weakness, as is the fact that only 3,250 of 8,357 women were ultimately included in the final analysis—raising concern of selection bias.

Bottom line: Keep symptoms in mind

Both patients and physicians alike should be aware that gynecologic, GI, and urinary symptoms may be indicators of ovarian cancer and may warrant further diagnostic evaluation. Until there is a screening test for ovarian cancer, heightened awareness is best.

Did you see these recent articles on ovarian cancer?

Check our Web archives for full-text access

I Update on Gynecologic Oncology

by Karen Lu, MD, and Michael R. Milam, MD, MPH University of Texas M.D. Anderson Cancer Center July 2007

Does menopausal HRT increase the risk of ovarian cancer?

Commentary by Andrew M. Kaunitz, MD University of Florida College of Medicine Examining the Evidence, July 2007

www.obgmanagement.com