## 哪些人需要內視鏡檢查及定期追蹤?

## Who need endoscopic screening and surveillance

Hsiu-Chi Cheng, MD, PhD

Department of Internal Medicine, Institute of Clinical Medicine and Molecular Medicine,

National Cheng Kung University Hospital, College of Medicine, National Cheng Kung University;

Department of Internal Medicine, Tainan Hospital, Ministry of Health and Welfare,

## Tainan, Taiwan

The purpose of endoscopic screening or surveillance is to detect precancerous lesions and/or early gastric cancer because the prognosis is highly correlated with the stage of cancer at diagnosis.

Patients who are at risk of gastric cancer are suggested to receive endoscopy screening and surveillance.

The etiologies of gastric cancer include *Helicobacter pylori* (*H. pylori*) infection, autoimmune gastritis, hereditary gene defects, and others. *H. pylori* is a class I carcinogen and *H. pylori*-related gastritis results in atrophic gastritis and intestinal metaplasia, which are precancerous conditions of gastric cancer. Approximately one third (33% [95% CI 26%–41%]) and one fourth (25% [95% CI 19%–30%]) of the population in the world have chronic atrophic gastritis and intestinal metaplasia, respectively. Moreover, the half of them has extensive chronic atrophic gastritis (16% [95% CI 12%–20%]) and intestinal metaplasia (13% [95% CI 9.0%–17%]) [1]. The incidence rates of gastric

cancer in patients with atrophic gastritis and intestinal metaplasia are 2.25 (95% CI, 1.67-2.90) and 7.58 (95% CI, 4.10–1.91) per 1000 person-years in East Asia, respectively [2]. Intestinal metaplasia almost occurs in underlying atrophic mucosa [3]; thus, the characteristics or biomarkers of atrophic gastritis could be used to screen subjects who need endoscopic screening. Male, H. pylori infection, old age ≥ 40 years, and residents in countries with high gastric cancer incidences are risk factors to have chronic atrophic gastritis [1]. Autoimmune gastritis induces corpus atrophic gastritis, pernicious anemia and/or iron deficiency anemia. Subjects with autoimmune gastritis are at risk of not only gastric adenocarcinoma (the odds ratio [OR] 2.18 [95% CI 1.94-2.45]) but also gastric carcinoid tumors (OR 11.43 [95% CI 8.90-14.69]) [4]. Thus, patients with autoimmune gastritis are suggested to receive endoscopic screening and surveillance to detect both gastric adenocarcinoma and type 1 gastric neuroendocrine tumor. Family history of gastric cancer, including first- and second-degree relative, increases the risk of gastric cancer (the relative risk 2.35 [95% CI 1.96-2.81]) [5]; thus, subjects with family history of gastric cancer are suggested to commence endoscopy screening at the age of 10 years earlier than that of affected relatives while gastric cancer is diagnosed [6]. In addition, there are various suggestions for gastric cancer screening for those with hereditary genetic mutations, i.e., familial gastric cancer syndromes [7].

The severity of atrophic gastritis and intestinal metaplasia could be evaluated by histological analysis by topographic biopsy based on operative link for gastritis assessment (OLGA) and/or operative link on gastric intestinal metaplasia assessment (OLGIM), endoscopic mucosal visualization

according to Kimura-Takemoto classification or endoscopic grading of gastric intestinal metaplasia (EGGIM), or serological tests by serum pepsinogen (PG) I and I/II ratio. OLGA or OLGIM stages III–IV predict higher risk of gastric cancer than stages 0–II (OR 2.64 [95% CI 1.84–3.79] for OLGA and 3.99 [95% CI 3.05–5.21] for OLGIM) [8]. Kimura-Takemoto severe-type or open-type of atrophy has increased risk of gastric neoplasm (the pooled risk ratio 3.89 [95% CI 2.92–5.17] for severe-type and 8.02 [95% CI 2.39–26.88] for open-type) [9]. EGGIM by using narrow-band imaging could predict OLGIM stages III–IV accurately (the area under the receiver operating characteristic curve 0.96 [95% CI 0.93–0.98])[10]. PG I  $\leq$  70 ng/mL and PG I/II ratio  $\leq$  3 could predict subjects to develop non-cardiac gastric cancer (OR 11.1 [95% CI 4.3–28.8]) [11]. Moreover, PG I  $\leq$  45 ng/mL and PG I/II ratio  $\leq$  6 could predict subjects with OLGA/OLGIM stages III-IV or gastric cancer (sensitivity 0.60 [95% CI 0.36–0.80] and specificity 0.71 [95% CI 0.65–0.76])[12].

In addition to down-staging gastric cancer, the balance between efficacy and costs of endoscopic surveillance is important. A cost-effectiveness analysis reported the optimal intervals of surveillance for subjects with different gastric cancer risk. The interval is annual for OR 5.46–21.5 and 2-yearly for OR 2.4–5.46 [13]. According to the OR of gastric cancer, the optimal interval of endoscopic surveillance for subjects with OLGA/OGLIM stages III–IV in Taiwan may be 2 to 3 years.

## Reference

1. Marques-Silva L, Areia M, Elvas L, Dinis-Ribeiro M. Prevalence of gastric precancerous conditions:

- a systematic review and meta-analysis. Eur J Gastroenterol Hepatol. 2014; 26:378-387.
- Akbari M, Tabrizi R, Kardeh S, Lankarani KB. Gastric cancer in patients
  with gastric atrophy and intestinal metaplasia: A systematic review and meta-analysis. PLoS One
  2019; 14:e0219865.
- Shah SC, Piazuelo MB, Kuipers EJ, Li D. AGA clinical practice update on the diagnosis and management of atrophic gastritis: expert review. Gastroenterology 2021; 161:1325-1332.e7.
- 4. Murphy G, Dawsey SM, Engels EA, et al. Cancer risk after pernicious anemia in the US elderly population. Clin Gastroenterol Hepatol 2015; 13:2282-2289.e1-4.
- 5. Yaghoobi M, McNabb-Baltar J, Bijarchi R, Hunt RH. What is the quantitative risk of gastric cancer in the first-degree relatives of patients? A meta-analysis. World J Gastroenterol. 2017; 23:2435-2442.
- 6. Luu MN, Quach DT, Hiyama T. Screening and surveillance for gastric cancer: Does family history play an important role in shaping our strategy? Asia Pac J Clin Oncol 2022; 18:353-362.
- 7. Setia N, Clark JW, Duda DG, et al. Familial Gastric Cancers. Oncologist 2015; 20:1365-1377.
- 8. Yue H, Shan L, Bin L. The significance of OLGA and OLGIM staging systems in the risk assessment of gastric cancer: a systematic review and meta-analysis. Gastric Cancer 2018; 21:579-587.
- 9. Xiao S, Fan Y, Yin Z, Zhou L. Endoscopic grading of gastric atrophy on risk assessment of gastric neoplasia: A systematic review and meta-analysis. J Gastroenterol Hepatol. 2021; 36:55-63.
- 10. Esposito G, Pimentel-Nunes P, Angeletti S, et al. Endoscopic grading of gastric intestinal metaplasia (EGGIM): a multicenter validation study. Endoscopy 2019; 51:515-521.
- 11. In H, Sarkar S, Ward J, et al. Serum pepsinogen as a biomarker for gastric cancer in the United States: A nested case-control study using the PLCO cancer screening trial data. Cancer Epidemiol Biomarkers Prev 2022; 31:1426-1432.
- 12. Chiang TH, Maeda M, Yamada H, et al. Risk stratification for gastric cancer after Helicobacter pylori eradication: A population-based study on Matsu Islands. J Gastroenterol Hepatol 2021; 36:671-679.
- 13. Zhou HJ, Dan YY, Naidoo N, Li SC, Yeoh KG. A cost-effectiveness analysis evaluating endoscopic surveillance for gastric cancer for populations with low to intermediate risk. PLoS One. 2013; 8:e83959.