

Review

Narrow-band imaging (NBI) magnifying endoscopic classification of colorectal tumors proposed by the Japan NBI Expert Team

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Many clinical studies on narrow-band imaging (NBI) magnifying endoscopy classifications advocated so far in Japan (Sano, Hiroshima, Showa, and Jikei classifications) have reported the usefulness of NBI magnifying endoscopy for qualitative and quantitative diagnosis of colorectal lesions. However, discussions at professional meetings have raised issues such as: (i) the presence of multiple terms for the same or similar findings; (ii) the necessity of including surface patterns in magnifying endoscopic classifications; and (iii) differences in the NBI findings in elevated and superficial lesions. To resolve these problems, the Japan NBI Expert Team (JNET) was constituted with the aim of establishing a universal NBI magnifying endoscopic classification for colorectal

tumors (JNET classification) in 2011. Consensus was reached on this classification using the modified Delphi method, and this classification was proposed in June 2014. The JNET classification consists of four categories of vessel and surface pattern (i.e. Types 1, 2A, 2B, and 3). Types 1, 2A, 2B, and 3 are correlated with the histopathological findings of hyperplastic polyp/sessile serrated polyp (SSP), low-grade intramucosal neoplasia, high-grade intramucosal neoplasia/shallow submucosal invasive cancer, and deep submucosal invasive cancer, respectively.

Key words: classification, colorectal neoplasms, magnifying endoscopy, narrow band imaging, the Japan NBI expert team (JNET)

INTRODUCTION

Narrow-band imaging (NBI) was initially developed by Sano *et al.* under the supervision of Dr Shigeaki Yoshida at the National Cancer Center Hospital East in 1999.^{1,2} A prototype short-wavelength narrow-band red/green/blue

(RGB) filter was successfully created in 2001 (monochrome NBI),^{1,2} and the microvascular architecture of the gastrointestinal tract and tumor surface structure were successfully visualized in color using 415- and 540-nm short- and medium-wavelength filters in 2003.³ Subsequently, various improvements, such as noise reduction, light amount adjustment and color adjustment, were made, and OLYMPUS EVIS LUCERA SPECTRUM (Olympus Medical Systems Corp., Tokyo, Japan) was launched in the market as the final mass production model in 2006.

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Sano *et al.* reported the usefulness of observing the pit pattern and the usefulness of vascular classification using NBI in 2004,³ and Sano *et al.* reported the first classification of the capillary pattern using NBI magnifying colonoscopy in 2006.⁴ Thereafter, the Magnifying Endoscopy Study Groups of the Ministry of Health, Labour and Welfare (Kudo, Sano, and Inoue groups) discussed the usefulness of NBI classification based on the findings of NBI magnifying endoscopy, which led to the proposal of new classifications (Hiroshima, Showa, and Jikei classifications) based on the capillary pattern classification from several institutions.

SANO, HIROSHIMA, SHOWA, AND JIKEI CLASSIFICATIONS BASED ON THE FINDINGS OF NBI MAGNIFYING ENDOSCOPY

Sano classification

The Sano classification (Fig. 1) was the first published NBI magnifying endoscopic classification, in 2006. This classification is based on the vascular findings. Validation studies have reported its usefulness in qualitative and quantitative diagnosis, its learning curve etc.^{5–8}

Hiroshima classification

The Hiroshima classification (Fig. 2), published in 2008, is based on the surface pattern in addition to the vascular findings on NBI magnifying endoscopy.^{9,10}

Showa classification

The Showa classification (Fig. 3) was proposed in 2009 as a classification mainly based on the vascular findings on NBI magnifying endoscopy.¹¹

Jikei classification

The Jikei classification (Fig. 4) was also proposed in 2009 as a classification based mainly on the vascular findings on NBI magnifying endoscopy.¹²

NBI INTERNATIONAL COLORECTAL ENDOSCOPIC (NICE) CLASSIFICATION OF TUMORS

Tanaka *et al.* constituted the Colon Tumor NBI Interest Group (CTNIG) in 2009, with the aim of creating a simple NBI endoscopic classification of colorectal tumors that

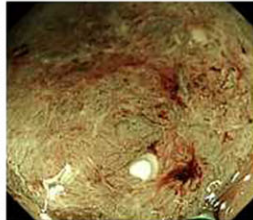
Capillary pattern	I	II	IIIA	IIIB
Schema				
Endoscopic findings				
Capillary characteristics	Meshed capillary vessels (-)	• Meshed capillary vessels (+) • Capillary vessel surrounds mucosal glands	Meshed capillary vessels characterized by: blind ending, branching and curtailed irregularly • Lack of uniformity • High density of capillary vessels	• Nearly avascular or loose micro capillary vessels

Figure 1 Sano classification (capillary pattern classification).

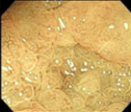
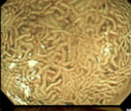
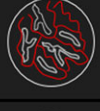
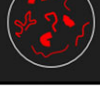
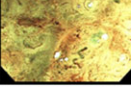
A type				Microvessel intensity are vague or invisible. None or isolated lacy vessels may be present coursing across the lesion. Brown or black dots, star or round shaped surrounded by white.
B type				Regular surface pattern is observed by the increased microvessel intensity around the pits and image enhance. Or regular meshed microvessel network pattern is observed.
C type	1			Irregular surface pattern is observed by the increased microvessel intensity around the pits and image enhance. Thickness and distribution of vessels are homogenous.
	2			More irregular surface pattern is observed by the increased microvessel intensity around the pits and image enhance. Thickness and distribution of vessels are heterogenous.
	3			Surface pattern is completely unclear. Thickness and distribution of vessels are heterogenous. Avascular area (AVA) and scattered microvessel fragments are observed.

Figure 2 Hiroshima classification.

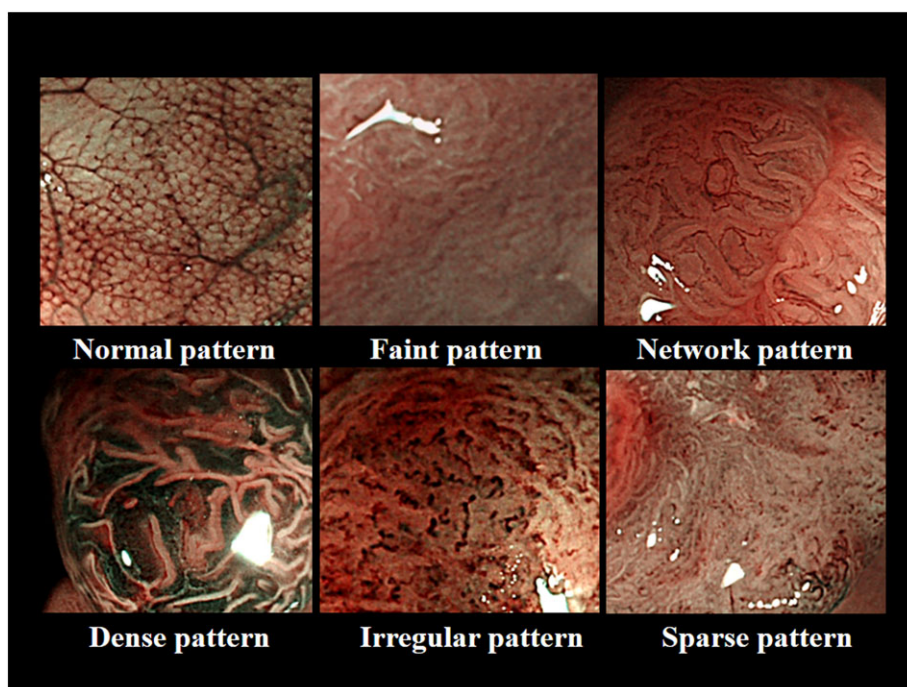


Figure 3 Showa classification.

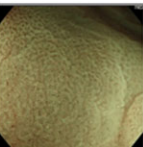
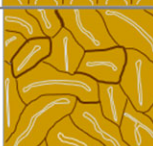
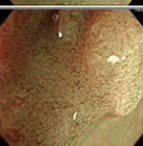
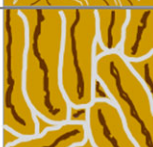
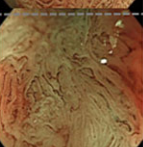
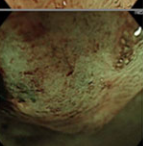
pattern 1			Invisible vessels
pattern 2			Slight caliber dilatation
pattern 3V			Regular arrangement of dilated vessels in the same places as the stroma of the ducts that are thought to have a villous structure
pattern 3I			Marked caliber dilatation Irregular arrangement of dilated vessels
pattern 4			Sparse distribution of vessels and untraceable vessel arrangement

Figure 4 Jikei classification.

can be used internationally. The CTNIG consist of the following six members: Chair, S Tanaka, Hiroshima University (Japan); Y Sano, Sano Hospital (Japan); DK Rex, Indiana University (USA); RM Soetikno, Stanford University (USA); T Ponchon, Edouard Herriot Hospital (France); and BP Saunders, St Mark's Hospital (UK). The CTNIG proposed the NBI International Colorectal Endoscopic (NICE) classification in 2009, which can also be used in assessment using non-magnifying endoscopy, and is based on the color, vessels, and surface pattern on endoscopy (Fig. 5). Naming of the NICE classification was proposed by Sano in London in February 2010 and was approved by the members of the CTNIG. International validation studies of the NICE classification have also proved useful in qualitative and quantitative diagnosis based on non-magnifying endoscopy.^{13,14}

CLINICAL PROBLEMS IN JAPAN AND SOME EXPERTS IN THE WEST

A number of validation studies of NBI magnifying endoscopic classifications proposed in Japan have reported the usefulness of NBI magnifying endoscopy in the qualitative and quantitative diagnosis of colorectal

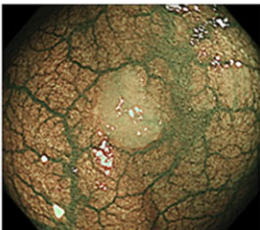
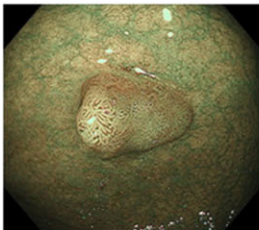
lesions. However, clinical studies have raised the following concerns.

- 1 Existence of multiple terms for the same or similar magnifying findings.
- 2 Necessity of including the surface patterns in magnifying endoscopic classifications.
- 3 Differences in the NBI magnifying findings in elevated and superficial lesions.

To resolve the aforementioned problems, the necessity of developing a universal classification is discussed.

UNIVERSAL NBI MAGNIFYING ENDOSCOPIC CLASSIFICATION OF COLORECTAL TUMORS: JAPAN NBI EXPERT TEAM (JNET) CLASSIFICATION

To establish a universal NBI magnifying endoscopic classification of colorectal tumors, the Japan NBI Expert Team (JNET), consisting of 38 members, mainly specialists in colonoscopy from throughout Japan, was formed within the Research Group of the National Cancer Center Research and Development Fund (Yutaka Saito Group) in 2011.

	Type 1	Type 2	Type 3
Color	Same or lighter than background	Browner relative to background (verify color arises from vessels)	Brown to dark brown relative to background; sometimes patchy whiter areas
Vessels	None, or isolated lacy vessels may be present coursing across the lesion	Brown vessels surrounding white structures**	Has area(s) of disrupted or missing vessels
Surface pattern	Dark or white spots of uniform size, or homogeneous absence of pattern	Oval, tubular or branched white structures** surrounded by brown vessels	Amorphous or absent surface pattern
Most likely pathology	Hyperplastic & sessile serrated polyp (SSP) ***	Adenoma****	Deep submucosal invasive cancer
Endoscopic image			

* Can be applied using colonoscopes with/ without optical (zoom) magnification

** These structures (regular or irregular) may represent the pits and the epithelium of the crypt opening.

*** In the WHO classification, sessile serrated polyp and sessile serrated adenoma are synonymous.

**** Type 2 consists of Vienna classification types 3, 4 and superficial 5 (all adenomas with either low or high grade dysplasia, or with superficial submucosal carcinoma). The presence of high grade dysplasia or superficial submucosal carcinoma may be suggested by an irregular vessel or surface pattern, and is often associated with atypical morphology (e.g., depressed area).

Figure 5 Narrow-band imaging (NBI) International Colorectal Endoscopic (NICE) classification.

First, it was necessary to establish common evaluation criteria, and a working group mainly consisting of young but experienced researchers (Shoichi Saito, Yoshiki Wada, Hiroaki Ikematsu, Hiroshi Kawano and Shiro Oka, Chairs; Takahisa Matsuda and Yutaka Saito) from six institutions, including the institution to which the member who first proposed establishment of a classification belonged, was organized to hold discussions. Consequently, normal/hyperplastic lesions/sessile serrated polyp were classified as Type 1, low-grade intramucosal neoplasia including intramucosal cancer with low-grade structural atypia as Type 2A, high-grade intramucosal neoplasia/shallow submucosal invasive cancer as Type 2B, and deep submucosal invasive cancers as Type 3, and a magnifying NBI scale including the vascular and surface patterns was created (Fig 6).¹⁵

To scientifically evaluate the NBI scale and determine the NBI findings and diagnostic criteria used in the universal classification, a web image interpretation study

(UMIN000010292: Multicenter study for developing universal NBI magnifying endoscopic classification of colorectal tumors in Japan, principal investigator: Yasushi Sano) was conducted by 25 specialists in colonoscopy belonging to the JNET in 2013. Twenty-five specialists in colonoscopy evaluated NBI findings and histology with 100 NBI still images on the web. Univariate and multivariate analyses and analysis of the diagnosability from five candidate NBI findings such as: (i) loose vessel areas; (ii) interruption of thick vessels; (iii) scattered vessels; (iv) thick, linearized/meandering atypical vessels in the tumor; and (v) amorphous areas of surface patterns, identified findings (i), (ii) and (v) for the diagnosis of Type 3. In addition, from among five candidate findings such as: (i) variable caliber of vessels; (ii) thick vessels; (iii) irregular distribution of vessels; (iv) vessel meandering; and (v) irregular or obscure surface pattern, findings (i), (iii) and (v) were selected for the diagnosis of Type 2B.

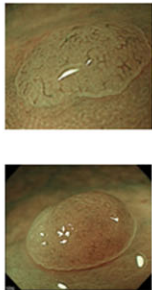
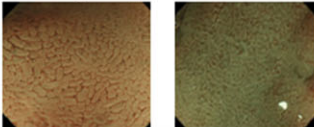
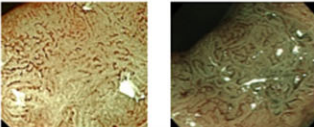
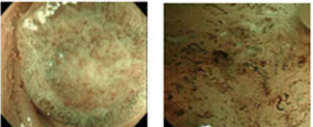
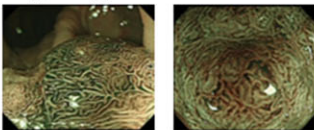
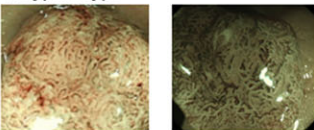
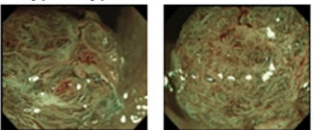
Normal/ Hyperplasia Type 1	Low grade adenoma Type 2A	High grade adenoma Type 2B	Deep submucosal invasive carcinoma Type 3
	Non-Polypoid type 	Non-Polypoid type 	Non-Polypoid type 
	Polypoid type 	Polypoid type 	Polypoid type 
None, or isolated lacy vessels may be present coursing across the lesion	Regular	Has area(s) with moderately distorted vessels	Has area(s) with markedly distorted or missing vessels
•Vessels are invisible •If vessels are visible, the vessel caliber in the lesion is the same as that in the surrounding normal mucosa •Lacy vessels coursing across the lesion	•Distribution of dark brown microvessels •Uniform and regular, relatively well-ordered reticular pattern (*Note that microvessels are often distributed in a punctate pattern and the well-ordered reticular pattern is not commonly observed in depressed lesions.)	•Varied caliber/caliber change •Thick vessels/vessel dilation •Uneven and irregular distribution of vessels •Vessel meandering *Approximately ≥ 1.5 times thicker than in adenomas	•Avascular areas or loose vascular areas •Disrupted thick vessels

Figure 6 Magnifying narrow-band imaging (NBI) scale. (a) Vascular pattern. (b) Appendix. (c) Surface pattern.

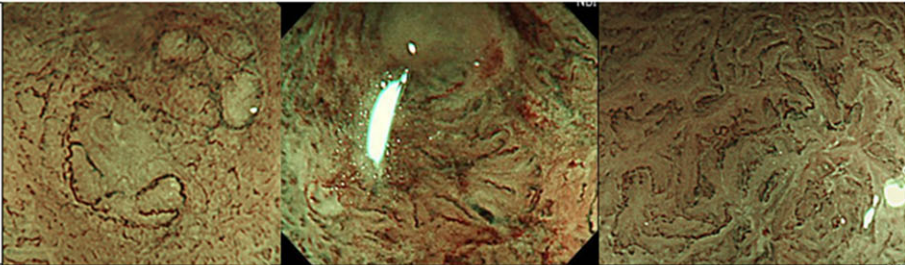
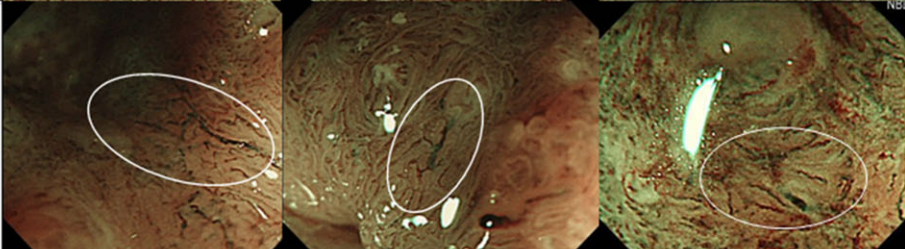
Scattered vessels Coarsely distributed, irregular vessels Probably corresponding to desmoplastic reaction	
Thick, linearized/meandering atypical vessels in the tumor Vessels similar to string sign described by Terai et al, but those found at the periphery are not taken into account	

Figure 6b (Continued)

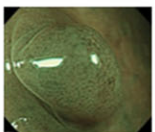
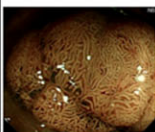
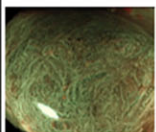
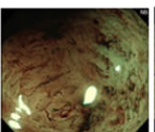
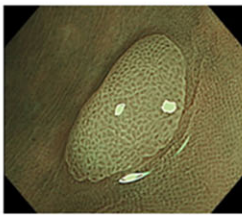
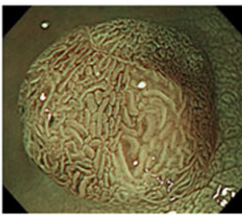
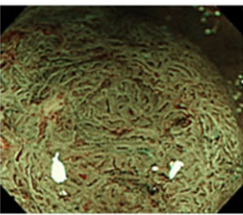
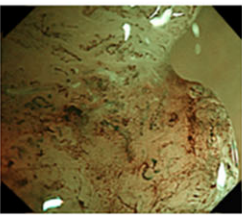
Normal/ Hyperplasia Type 1	Low grade adenoma Type 2A	High grade adenoma Type 2B	Deep submucosal invasive carcinoma Type 3
			
Dark or white spots of uniform size, or homogeneous absence of pattern	Regular	Irregular	Amorphous
•Regular dark or white spots •Uniformly obscure structure	•Tubular or dendritic or papillary •Regular surface pattern •Corresponding to type III or IV pit pattern	•Visible surface pattern with irregularity •Corresponding to the type Vi pit pattern	•Invisible surface pattern •Corresponding to the type Vn pit pattern

Figure 6c (Continued)

	Type 1	Type 2A	Type 2B	Type 3
Vessel pattern	• Invisible* ¹	• Regular caliber • Regular distribution (meshed/spiral pattern)* ²	• Variable caliber • Irregular distribution	• Loose vessel areas • Interruption of thick vessels
Surface pattern	• Regular dark or white spots • Similar to surrounding normal mucosa	• Regular (tubular/branched/papillary)	• Irregular or obscure	• Amorphous areas
Most likely histology	Hyperplastic polyp/ Sessile serrated polyp	Low grade intramucosal neoplasia	High grade intramucosal neoplasia/ Shallow submucosal invasive cancer* ³	Deep submucosal invasive cancer
Endoscopic image				

*1. If visible, the caliber in the lesion is similar to surrounding normal mucosa.

*2. Micro-vessels are often distributed in a punctate pattern and well-ordered reticular or spiral vessels may not be observed in depressed lesions.

*3. Deep submucosal invasive cancer may be included.

Figure 7 The Japan NBI Expert Team (JNET) classification.

The Yutaka Saito Group meeting on 6 June 2014 reached a consensus on the universal NBI magnifying endoscopic classification of colorectal tumors based on scientific grounds using a modified Delphi method, followed by its proposal by the JNET (Fig. 7).

The JNET classification unifying previous classifications will give us common diagnostic criteria to promote academic progress of NBI. Now we need to standardize the interpretation of each NBI finding through various discussions and validate the JNET classification itself.

FUTURE PERSPECTIVES

Fifteen years have passed since the introduction of NBI (14 December 1999), and a consensus was reached on the original Japanese universal NBI magnifying endoscopic classification of colorectal tumors (JNET classification). At present, validation studies for the JNET classification are proposed to be conducted by the Research Group of the National Cancer Center Research and Development Fund (Yutaka Saito Group) and a study group of the Japan Gastroenterological Endoscopy Society (a study group for the standardization and dissemination of the magnifying colonoscopic diagnostic criteria, representative organizer: Takahisa Matsuda). Modifications may be made through these studies, but it should be kept in mind that the classification will be continuously updated by many new findings and innovations in endoscopy. Moreover, it is an urgent task to disseminate this classification in many clinical settings in Japan and to scientifically prove that this is useful in qualitative and quantitative diagnosis, which can determine the therapeutic strategy for colorectal tumors.

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CONFLICTS OF INTEREST

AUTHORS DECLARE NO conflicts of interest for this article.

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