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2018 Korean Clinical Imaging Guideline for Hemoptysis

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In 2014, the American College of Radiology (ACR) announced the guideline for the appropriate diagnostic approach and treatment of patients according to the severity of hemoptysis and risk for lung cancer. However, the application of the ACR guideline in Korea may not be appropriate, because many patients in Korea have active tuberculosis or pulmonary fibrosis due to previous tuberculosis. The Korean Society of Radiology and Korean Society of Thoracic Radiology have proposed a new guideline suitable for Korean practice. This new guideline was prepared through the consensus of a development committee, working party, and an advisory committee. The guideline proposal process was based on an evidence-based clinical imaging guideline proposed by the development committee. Clinical imaging guideline for adult patients with hemoptysis is as follows: Chest radiography is an initial imaging modality to evaluate hemoptysis. Contrast-enhanced chest CT is recommended in patients with two risk factors for lung cancer (> 40 years old and > 30 pack-year smoking history), moderate hemoptysis (> 30 mL/24 hours) or recurrent hemoptysis. Contrast-enhanced chest CT is also recommended in patients with massive hemoptysis (> 400 mL/24 hours) without cardiopulmonary compromise.

Keywords: Guideline; Hemoptysis; Thorax; Lung; Radiography; X-ray; Computed tomography; Evidence; Republic of Korea

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Background of the Development of the Korean Clinical Imaging Guideline

Hemoptysis is the expectoration of blood to mouth or nose that originates from the respiratory tract. Clinical management is different according to the severity of hemoptysis based on the volume of blood loss. However, retained blood in lung parenchyma can be underestimated due to the difficulty in determining the exact volume of blood loss in routine practice. Diagnostic imaging is salient for appropriate diagnosis and treatment because hemoptysis can originate from various diseases such as acute benign disease (including bronchietasis), or malignant tumor.

A clinical imaging guideline for hemoptysis has been previously published by the American College of Radiology (ACR) (1). ACR guideline states diagnostic and treatment strategies according to the severity of hemoptysis and risk factor for lung cancer. The ACR emphasized the importance of undiagnosed lung cancer in hemoptysis patients.

However, there is a need for a guideline specific to Korea, because it is an endemic tuberculosis area with a high rate of hemoptysis patients with pulmonary fibrosis caused by previous tuberculosis or hemoptysis from active tuberculosis. Consequently, the development committee, working party, and an advisory committee have developed a quideline suitable for Korean practice.

Adaptation Process of the Guidelines

The guideline development process was based on a guideline adaptation methodology process suggested by the development committee (2).

Key Questions

The key questions made by working group were reviewed and revised by the development committee and consensus group. The consensus group represented the suggested end user and was selected from The Korean Academy of Tuberculosis and Respiratory Disease. The finalized key question is in a structured form below.

1. What is the most appropriate imaging test to diagnose the cause of hemoptysis in an adult patient with hemoptysis?

Search Guidelines

The development committee searched for guidelines using international databases (Ovid-MEDLINE and Ovid-

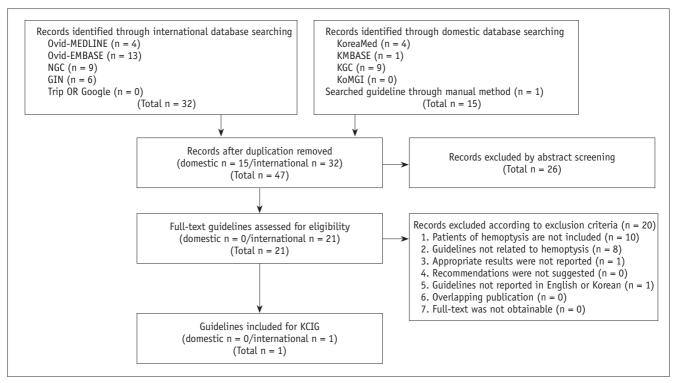


Fig. 1. Flow chart for literature selection. GIN = Guideline International Network, KCIG = Korean Clinical Imaging Guidelines, KGC = Korean Guideline Clearinghouse, KoMGI = Korean Medical Guidelines and Information, NGC = National Guideline Clearinghouse

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Table 1. AGREE II

Source of Recommendation	AGREE II Score	Proposal of Developmental Committee
ACR Appropriateness Criteria® hemoptysis	66	Recommended

Not recommended: AGREE score < 50. ACR = American College of Radiology, AGREE II = Appraisal of Guidelines for Research & Evaluation II

Table 2. Criteria for Evidence Level of Each Evidence Literature

Level	Content		
	Research satisfying all of criteria following three		
	1) Good reference standard		
1	2) Consecutive patients study		
1	3) Blind interpretation		
	Systematic review of level 1		
	Randomized controlled trial or cross-sectional cohort study that compares index test to comparators		
Research satisfying all of criteria following two			
	1) Good reference standard		
2	2) Consecutive patients study or blind interpretation		
	Systematic review of level 2		
	Observational studies that compares index test to comparators		
3	Without consistently applied reference standards		
,	Case-control study		
4	Poor or non-independent reference standard		
5	Expert opinion		

Adapted from Choi et al. Korean J Radiol 2017;18:208-216 (2)

Table 3. Grades of Korean Clinical Imaging Guideline Recommendation

Grading	Content	Meaning	
Α	Recommended	This intervention (examination) has enough evidence to support desired effect, and therefore, is recommended	
В	(Conditional) recommended	This intervention (examination) has intermediate to enough level of evidence to support desired effect Provide intervention (examination) selectively, or for specific individuals based on expert's judgment	
С	Not recommended	This intervention (examination) has enough evidence to support non-desired effect, and therefore, is not recommended (use of this examination is not recommended)	
I	No recommendation	This intervention (examination) does not have enough evidence to either support or reject effectiveness and needs further research This intervention (examination) has very low level of certainty for desired effect, and decision based on recommendation grading has no meaning	

Adapted from Choi et al. Korean J Radiol 2017;18:208-216 (2)

EMBASE), several international sites associated with guidelines (National Guideline Clearinghouse and Guideline International Network) and major domestic databases (KoreaMed, KMbase, Korean Medical Guidelines and Information, and Korean Guideline Clearinghouse).

In addition, the websites of major academic societies and institutions were also searched along with a manual database search. We performed literature searches from 2000 to July, 2015 for the keywords hemoptysis, computerized tomography, or bronchoscopy, using Medical Subject Heading (MeSH) terms or Embase subject headings (EMTREE). Guidelines or articles published after that period

were included by an up-to-dated search.

Selection of Searched Guidelines

A total of 47 guidelines were searched: 32 guidelines in an international database, 14 guidelines in a domestic database, and 1 guideline by a manual search. In the primary screening process, 21 records were selected after reviewing the title and abstract of an identified study or guideline. One guideline was selected after the full-texts of identified literature were reviewed in the secondary screening process that also noted the reason for the exclusion of any specific literature selected (Fig 1).



Quality Appraisal of the Guidelines

The final selected guidelines underwent quality appraisal using the Korean Appraisal of Guidelines for Research & Evaluation II (AGREE II) tool of the development committee. Final appraisal results were provided to the working group (Table 1). The working group selected one guideline for the key question in consideration of the quality of the selected quideline.

Grading the Level of Evidence and Drafting the Recommendation Document

Working group members reviewed evidence in the literature that supported the final selected guideline recommendations. Grading the level of evidence for each literature was performed to create an evidence table (Supplementary Table 1 in the online-only Data Supplement) according to the evidence level criteria of the Korean Clinical Imaging Guidelines (K-CIG) (Tables 2, 3). The draft version of the recommendation document was based on the prepared evidence table after a final discussion of the working group.

Agreement of the Recommendation Grades

The draft version of the recommendation document prepared by the working group was reviewed and discussed by the development committee. The final level of evidence and grade of recommendation was determined by the consensus of the development committee and working group, according to K-CIG criteria (2).

Finalizing the Recommendation Document

The Delphi method was used for formal consensus. The consensus group was composed of clinical imaging experts, end-users from the clinical imaging guideline-related academic societies (The Korean Academy of Tuberculosis and Respiratory Disease), and research methodology experts.

The results of the Delphi were as follows: degree of agreement on primary survey was 7.3–8.6 (standard deviation: 0.5–0.9) and degree of agreement on secondary survey 6.9–8.5 (standard deviation: 0.5–0.8).

External Review and Approval of Clinical Guideline

Clinical experts related to the area of the recommendations, such as members from The Korean Academy of Tuberculosis and Respiratory Disease reviewed the final recommendations. Feedback and modification was subsequently reflected in the documents.

Recommendations

Key Question

1. What is the most appropriate imaging test to diagnose the cause of hemoptysis in an adult patient with hemoptysis?

[Recommendation 1-1]

Chest radiography is suggested for initial evaluation in all adult patients with hemoptysis.

(recommendation grade A, evidence level II)

[Recommendation 1-2]

Contrast-enhanced chest CT scan is recommended to diagnose the cause of hemoptysis in all adult patients with hemoptysis and who have two risk factors for lung cancer (> 40 years old and > 30 pack-year smoking history).

(recommendation grade A, evidence level II)

[Recommendation 1-3]

Contrast-enhanced chest CT scan is recommended to diagnose the cause of hemoptysis in all adult patients with moderate hemoptysis (> 30 mL/24 hours) or recurrent hemoptysis.

(recommendation grade A, evidence level II)

[Recommendation 1-4]

Contrast-enhanced chest CT scan should also be considered in adult patients with massive hemoptysis (> 400 mL/24 hours) and preserved cardiopulmonary function. (recommendation grade A, evidence level III)

Evidence Summary

Hemoptysis is spitting blood from the respiratory tract due to various causes that range from blood tinged sputum to massive hemoptysis. Causes of hemoptysis include benign disease such as bronchiectasis, acute bronchitis, chronic bronchitis, tuberculosis, diffuse interstitial lung disease, and pulmonary venous malformation or malignant disease such as lung cancer and idiopathic. Therefore, appropriate screening test and complete medical examinations are required (3-5). This paper proposes a clinical imaging guideline for adult patients with hemoptysis.

The guideline recommends chest radiography as an initial evaluation for all adult hemoptysis patients. Chest radiography can help lateralize the bleeding site and screen for the presence or absence of abnormality in the pleura

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and lung parenchyma (3, 6, 7).

Clinical imaging guidelines differ according to the amount of hemoptysis: minor (< 30 mL/24 hours), moderate (30–400 mL/24 hours), or massive hemoptysis (> 400 mL/24 hours) (7-10).

Contrast-enhanced chest CT scan is recommended to exclude the possibility of lung cancer in patients with hemoptysis and normal chest radiographs, if the patient has two risk factors for malignancy (> 40 years old and > 30 pack-year smoking history). In adult patients with hemoptysis, the incidence of malignancy was reported as 10–35% (2, 3). In a report on the long-term outcome and incidence of lung cancer in patients with hemoptysis of unknown origin, unresectable lung cancer developed in 6% of patients within 3 years after the first presentation; all patients were smokers and > 40 years old (11). In another retrospective study of patients with hemoptysis and normal radiography, 9.6% of patients were diagnosed with lung cancer; all patients were current or ex-smokers (12).

The accurate assessment of the cause is more important than prompt treatment in patients with moderate hemoptysis (> 30 mL/24 hours) or recurrent hemoptysis. Previous reports indicate that the most common cause of hemoptysis in Korea was bronchiectasis, followed by active tuberculosis, and tuberculosis sequelae (5). Possible remaining causes were pneumonia, aspergilloma, lung cancer, others, or unknown (5). CT is more useful than bronchoscopy, because it is noninvasive and accurately assesses the cause and site of hemoptysis; in addition, it is also possible to diagnose bronchiectasis, tuberculosis and lung cancer with non-contrast-enhanced CT. However, contrast-enhanced CT is recommended to accurately assess the bleeding site, because non-bronchial collateral arteries could be the source of bleeding in tuberculosis patients despite the bronchial artery being the common source of bleeding (13-15). Inoperable massive hemoptysis was considered a major indication of bronchial artery

embolization; however, moderate hemoptysis or mild but chronic and worsening hemoptysis is also considered an indication of bronchial artery embolization (14). Contrastenhanced CT can guide bronchial artery embolization and reduce procedure time, because it adequately depicts the imaging anatomy of the bronchial artery and non-bronchial collateral arteries (intercostal artery, subclavian artery, axillary artery, and inferior phrenic artery), pulmonary artery and pulmonary vein (16).

A contrast-enhanced CT scan is recommended for patients with massive hemoptysis (> 400 mL/24 hours) who preserve cardiopulmonary function. Massive hemoptysis can be treated by bronchial artery embolization or surgery; in addition, CT prior to these treatments can be helpful to detect the causative vessel (7, 17).

Consideration for Recommendation

Benefit and Harm

Performing chest radiography as a screening test in patients with hemoptysis is useful in terms of low radiation exposure, lateralization of bleeding site, and as a screening test for lung disease.

Regardless of the amount of hemoptysis, chest CT is recommended in patients who have two risk factors for malignancy (> 40 years old and > 30 pack-year smoking history). Chest CT has the benefit to detect hidden lung cancer that cannot be detected on chest radiography; however, caution is advised due to radiation exposure.

CT is very useful for the simultaneous detection of the bleeding site and cause of bleeding in patients with moderate hemoptysis (> 30 mL/24 hours) or recurrent hemoptysis. In Korea, hemoptysis is commonly associated with inflammatory diseases that include active tuberculosis and tuberculosis sequelae. In these cases, chest radiography has several limitations in the evaluation of the bleeding site. CT is therefore very useful for the exact evaluation of

Table 4. Korean Relative Radiation Level

Symbol	RRL (mSv)	Example
0	0	Sonography, MRI
A	< 1	Chest radiography (PA view), mammography
	1–5	IVU, UGIS, low-dose chest CT, brain CT, brain CTA
$\triangle \triangle \triangle$	> 5–10	Routine chest CT, abdominal CT, coronary CT
	> 10	3-phase dynamic CT (abdomen)

CT = computed (computer) tomography, CTA = computed tomography angiography, IVU = intravenous urography, MRI = magnetic resonance imaging, RRL = relative radiation level, PA view = posteroanterior view, UGIS = upper gastrointestinal series. Adapted from Choi et al. *Korean J Radiol* 2017;18:208-216 (2).



bleeding size and causative disease.

Bronchial artery embolization is considered a treatment option in patients with massive hemoptysis, moderate hemoptysis, or recurrent hemoptysis. CT performed before embolization shows a detailed anatomy of both the bronchial artery and pulmonary artery, helps guide the procedure, and can reduce the procedure time. CT before embolization has significant benefits that outweigh radiation hazards.

Contrast-enhanced CT could have hazards related to the iodine contrast media. We should follow the consultation guide for contrast media that indicates the benefits of contrast media versus the harm from iodine before choosing to use contrast media.

Radiation Dose

Relative radiation level of chest radiography is less than 1 mSv and that of chest CT is 5–10 mSv (Table 4).

Supplementary Materials

The online-only Data Supplement is available with this article at https://doi.org/10.3348/kjr.2018.19.5.866.

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