

呼吸器・アレルギー内科学講座

○主な研究内容

- 1 特発性間質性肺炎の疫学、病態、治療に関する研究
- 2 肺胞上皮細胞の分化、再生に関する研究
- 3 肺サーファクタント蛋白質の機能に関する研究
- 4 肺がんの分子機構に関する研究
- 5 気管支喘息の病態に関する研究
- 6 膠原病肺疾患の診断、病態に関する研究
- 7 肺炎球菌感染症とそのワクチンに関する研究
- 8 インフルエンザ桿菌感染症に関する研究

○Pub Med 掲載論文 (2018 年)

1. Characterization of distal airway stem-like cells expressing N-terminally truncated p63 and thyroid transcription factor-1 in the human lung.

Tanaka Y, Yamaguchi M, Hirai S, Sumi T, Tada M, Saito A, Chiba H, Kojima T, Watanabe A, Takahashi H, Sakuma Y.

Exp Cell Res. 2018 Nov 15;372(2):141-149. doi: 10.1016/j.yexcr.2018.09.020. Epub 2018 Sep 27.

PMID: 30268759

2. Poorer Prognosis of Idiopathic Pleuroparenchymal Fibroelastosis Compared with Idiopathic Pulmonary Fibrosis in Advanced Stage.

Shioya M, Otsuka M, Yamada G, Umeda Y, Ikeda K, Nishikiori H, Kuronuma K, Chiba H, Takahashi H.

Can Respir J. 2018 Aug 13;2018:6043053. doi: 10.1155/2018/6043053. eCollection 2018.

PMID: 30186537 Free PMC Article

3. A new ENG mutation in a Japanese family with hereditary hemorrhagic telangiectasia and pulmonary arteriovenous malformations.

Yokoo K, Yamada G, Chiba H, Ishikawa A, Morisaki H, Saijo H, Kudoh S, Kitamura Y, Hirokawa N, Miyajima M, Watanabe A, Takahashi H.

Respir Med Case Rep. 2018 Jul 2;25:73-77. doi: 10.1016/j.rmcr.2018.07.001. eCollection 2018.

PMID: 30073140 Free PMC Article

4. Significance of molecular biomarkers in idiopathic pulmonary fibrosis: A mini review.

Chiba H, Otsuka M, Takahashi H.

Respir Investig. 2018 Sep;56(5):384-391. doi: 10.1016/j.resinv.2018.06.001. Epub 2018 Jul 17. Review.

PMID: 30030108

5. Response to pneumococcal vaccine in interstitial lung disease patients: Influence of systemic immunosuppressive treatment.

Kuronuma K, Honda H, Mikami T, Saito A, Ikeda K, Otsuka M, Chiba H, Yamada G, Sato T, Yokota SI, Takahashi H.
Vaccine. 2018 Aug 9;36(33):4968–4972. doi: 10.1016/j.vaccine.2018.06.062. Epub 2018 Jul 6.
PMID: 29983256

6. Impaired diversity of the lung microbiome predicts progression of idiopathic pulmonary fibrosis.

Takahashi Y, Saito A, Chiba H, Kuronuma K, Ikeda K, Kobayashi T, Ariki S, Takahashi M, Sasaki Y, Takahashi H.
Respir Res. 2018 Feb 27;19(1):34. doi: 10.1186/s12931-018-0736-9.
PMID: 29486761 Free Article

7. Multiclonal Expansion and High Prevalence of β -Lactamase-Negative Haemophilus influenzae with High-Level Ampicillin Resistance in Japan and Susceptibility to Quinolones.

Honda H, Sato T, Shinagawa M, Fukushima Y, Nakajima C, Suzuki Y, Shiraishi T, Kuronuma K, Takahashi S, Takahashi H, Yokota SI.
Antimicrob Agents Chemother. 2018 Aug 27;62(9). pii: e00851-18. doi: 10.1128/AAC.00851-18. Print 2018 Sep.
PMID: 29987153

8. Whole genome analysis of a multidrug-resistant Streptococcus pneumoniae isolate from a patient with invasive pneumococcal infection developing disseminated intravascular coagulation.

Ohkoshi Y, Sato T, Wada T, Fukushima Y, Murabayashi H, Takakuwa Y, Nishiyama K, Honda H, Shiraishi T, Kuronuma K, Takahashi H, Nakajima C, Suzuki Y, Yokota SI.
J Infect Chemother. 2018 Aug;24(8):674–681. doi: 10.1016/j.jiac.2018.01.012. Epub 2018 Feb 26.
PMID: 29496334

9. Occult ovarian clear-cell carcinoma diagnosed as primary adenocarcinoma of the lung: A case report of a diagnostic pitfall for clinicians and pathologists.

Kubo T, Hirohashi Y, Fujita H, Sugita S, Kikuchi Y, Shinkawa T, Nakatsugawa M, Tsujiwaki M, Sudo Y, Asai Y, Umeda Y, Takahashi H, Hasegawa T, Torigoe T.
Respir Med Case Rep. 2018 Oct 18;25:306–308. doi: 10.1016/j.rmcr.2018.10.013. eCollection 2018.
PMID: 30386720 Free PMC Article

○その他の論文 (2018年)

1. 両肺の自然気胸を反復したサルコイドーシスの1例(原著論文)
Author : 小玉 賢太郎(札幌医科大学 医学部呼吸器・アレルギー内科学講座), 角 俊行, 高橋 洋平, 吉川 匠, 多屋 哲也, 森 勇樹, 宮島 さつき, 高橋 守, 千葉 弘文, 山田 玄, 高橋 弘毅
Source : 日本サルコイドーシス/肉芽腫性疾患学会雑誌 (1883-1273) 38 巻 1-2 号 Page66-70(2018. 10)
2. 特発性肺線維症における肺マイクロバイオーム研究の展開(総説)
Author : 黒沼 幸治(札幌医科大学 医学部呼吸器・アレルギー内科学講座), 齋藤 充史, 池田 貴美之, 大塚 満雄, 千葉 弘文, 高橋 弘毅
Source : 日本肺サーファクタント・界面医学会雑誌 (1882-1812) 49 巻 Page1-7(2018. 10)
3. 間質性肺炎の新しい診断・治療概念(解説)
Author : 高橋 弘毅(札幌医科大学 呼吸器・アレルギー内科学講座)
Source : 日本内科学会雑誌 (0021-5384) 107 巻 9 号 Page1819-1825(2018. 09)
4. 【間質性肺炎・肺線維症のトピックス】 特発性肺線維症の重症度評価(解説/特集)
Author : 千葉 弘文(札幌医科大学 医学部呼吸器・アレルギー内科学講座), 高橋 弘毅
Source : 呼吸器内科 (1884-2887) 34 巻 3 号 Page211-216(2018. 09)
5. 【ぜひ知っておきたい水・電解質異常と輸液管理】 各種病態における輸液管理 呼吸不全における輸液・栄養療法(解説/特集)
Author : 錦織 博貴(札幌医科大学 医学部呼吸器・アレルギー内科学講座), 千葉 弘文, 高橋 弘毅
Source : 臨牀と研究 (0021-4965) 95 巻 8 号 Page857-860(2018. 08)
6. 【特発性間質性肺炎の診療の現状と将来展望】 特発性肺線維症の予後因子と重症度分類(解説/特集)
Author : 千葉 弘文(札幌医科大学 医学部呼吸器・アレルギー内科学講座), 錦織 博貴, 高橋 弘毅
Source : 呼吸臨床 (2433-3778) 2 巻 5 号 Page1-8(2018. 05)
7. セツキシマブと放射線療法の併用により間質性肺炎を発症した下咽頭癌の2例(原著論文/症例報告)
Author : 吉川 匠(札幌医科大学 医学部呼吸器・アレルギー内科学講座), 大塚 満雄, 池田 貴美之, 高橋 知之, 山田 玄, 高橋 弘毅
Source : 日本呼吸器学会誌 (2186-5876) 7 巻 4 号 Page234-239(2018. 07)
8. 薬剤安全使用のための薬剤性肺障害診断のコツ すべての薬剤において肺障害を起こす可能性があることを念頭に、早期診断することが重要(Q&A)
Author : 坂東 政司(自治医科大学 内科学講座呼吸器内科学部門), 高橋 弘毅
Source : 日本医事新報 (0385-9215) 4910 号 Page52-53(2018. 06)
9. 有癭性膿気胸に対して局所麻酔下胸腔鏡で自己血注入とフィブリンシートを用いて治療した肺扁平上皮癌の1例(原著論文/症例報告)
Author : 田中 悠祐(札幌医科大学 医学部呼吸器・アレルギー内科学講座), 高橋 守, 橋本 みどり, 本田 泰人, 山田 玄, 高橋 弘毅
Source : 肺癌 (0386-9628) 58 巻 1 号 Page41-45(2018. 02)
10. 経気管支生検を施行した線維形成型悪性胸膜中皮腫の1剖検例(原著論文/症例報告)

Author: 本田 宏幸(函館五稜郭病院 呼吸器内科), 森 裕二, 石川 立, 小野 貴広, 中田 尚志,
高橋 弘毅

Source: 肺癌 (0386-9628) 58 卷 1 号 Page29-34 (2018. 02)

11. 特発性肺線維症の投薬タイミングと治療薬の使いわけ 肺活量が 10%以上の経年減少を認めた症例には積極的に投与。潜在的膠原病(各種自己抗体陽性)を基礎疾患とする IPF にはピルフェニドンを推奨(Q&A)

Author: 高橋 弘毅(札幌医科大学 呼吸器内科), 永田 真

Source: 日本医事新報 (0385-9215) 4902 号 Page54 (2018. 04)

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- 1 特発性間質性肺炎の診断、病態、治療に関する研究
- 2 肺サーファクタント蛋白の機能に関する研究
- 3 肺癌の診断、予後因子、治療に関する研究
- 4 肺癌の浸潤転移に関する分子生物学的研究
- 5 サルコイドーシスの診断、病態に関する研究
- 6 気管支喘息の病態、治療に関する研究
- 7 環境と気管支喘息、過敏性肺炎に関する研究
- 8 膠原病の肺疾患の診断、病態、治療に関する研究
- 9 各種肺疾患の三次元的病態像の解析
- 10 臨床CT、MRI、超音波像の解析
- 11 肺気腫の診断、病態に関する研究
- 12 HRC Tを用いた末梢気道立体構築の研究

○Pub Med 掲載論文 (2017 年)

1. Treatment of pulmonary fibrosis with siRNA against a collagen-specific chaperone HSP47 in vitamin A-coupled liposomes.

Otsuka M, Shiratori M, Chiba H, Kuronuma K, Sato Y, Niitsu Y, Takahashi H.
Exp Lung Res. 2017 Oct 16:1-12. doi: 10.1080/01902148.2017.1354946. [Epub ahead of print]
PMID: 29035148

2. Surfactant protein A downregulates epidermal growth factor receptor by mechanisms different from those of surfactant protein D.

Hasegawa Y, Takahashi M, Ariki S, Saito A, Uehara Y, Takamiya R, Kuronuma K, Chiba H, Sakuma Y, Takahashi H, Kuroki Y.
J Biol Chem. 2017 Sep 27. pii: jbc.M117.800771. doi: 10.1074/jbc.M117.800771. [Epub ahead of print]
PMID: 28972165

3. Serum surfactant protein D predicts the outcome of patients with idiopathic pulmonary fibrosis treated with pirfenidone.

Ikeda K, Shiratori M, Chiba H, Nishikiori H, Yokoo K, Saito A, Hasegawa Y, Kuronuma K, Otsuka M, Yamada G, Takahashi H.
Respir Med. 2017 Oct;131:184-191. doi: 10.1016/j.rmed.2017.08.021. Epub 2017 Aug 24.
PMID: 28947028

4. Daily physical activity affects exercise capacity in patients with idiopathic pulmonary fibrosis.

Morino A, Takahashi H, Chiba H, Ishiai S.
J Phys Ther Sci. 2017 Aug;29(8):1323-1328. doi: 10.1589/jpts.29.1323. Epub 2017 Aug 10.

PMID: 28878456

5. Disease severity staging system for idiopathic pulmonary fibrosis in Japan.

Kondoh Y, Taniguchi H, Kataoka K, Furukawa T, Ando M, Murotani K, Mishima M, Inoue Y, Ogura T, Bando M, Hagiwara K, Suda T, Chiba H, Takahashi H, Sugiyama Y, Homma S. *Respirology*. 2017 Nov;22(8):1609–1614. doi: 10.1111/resp.13138. Epub 2017 Aug 8. PMID: 28787101

6. Surfactant protein D inhibits activation of non-small cell lung cancer-associated mutant EGFR and affects clinical outcomes of patients.

Umeda Y, Hasegawa Y, Otsuka M, Arika S, Takamiya R, Saito A, Uehara Y, Saijo H, Kuronuma K, Chiba H, Ohnishi H, Sakuma Y, Takahashi H, Kuroki Y, Takahashi M. *Oncogene*. 2017 Jul 24. doi: 10.1038/onc.2017.253. [Epub ahead of print] PMID: 28745320

7. Factors affecting dyspnea after the 6-minute walk test in idiopathic pulmonary fibrosis patients presenting with exercise-induced hypoxemia.

Morino A, Takahashi H, Chiba H, Ishiai S. *J Phys Ther Sci*. 2017 Aug;29(8):1458–1462. doi: 10.1589/jpts.29.1458. Epub 2017 Aug 10. PMID: 28878483

8. Volatile Organic Compounds in Exhaled Breath of Idiopathic Pulmonary Fibrosis for Discrimination from Healthy Subjects.

Yamada YI, Yamada G, Otsuka M, Nishikiori H, Ikeda K, Umeda Y, Ohnishi H, Kuronuma K, Chiba H, Baumbach JI, Takahashi H. *Lung*. 2017 Apr;195(2):247–254. doi: 10.1007/s00408-017-9979-3. Epub 2017 Mar 1. PMID: 28247041

9. Fibroblastic foci, covered with alveolar epithelia exhibiting epithelial-mesenchymal transition, destroy alveolar septa by disrupting blood flow in idiopathic pulmonary fibrosis.

Yamaguchi M, Hirai S, Tanaka Y, Sumi T, Miyajima M, Mishina T, Yamada G, Otsuka M, Hasegawa T, Kojima T, Niki T, Watanabe A, Takahashi H, Sakuma Y. *Lab Invest*. 2017 Mar;97(3):232–242. doi: 10.1038/labinvest.2016.135. Epub 2016 Dec 12. PMID: 27941755

10. Angiotensin-converting enzyme 2 is a potential therapeutic target for EGFR-mutant lung adenocarcinoma.

Yamaguchi M, Hirai S, Sumi T, Tanaka Y, Tada M, Nishii Y, Hasegawa T, Uchida H, Yamada G, Watanabe A, Takahashi H, Sakuma Y. *Biochem Biophys Res Commun*. 2017 Jun 3;487(3):613–618. doi: 10.1016/j.bbrc.2017.04.102. Epub 2017 Apr 20. PMID: 28433633

11. Surfactant protein A (SP-A) and SP-A-derived peptide attenuate chemotaxis of mast cells induced by human β -defensin 3.

Uehara Y, Takahashi M, Murata M, Saito A, Takamiya R, Hasegawa Y, Kuronuma K, Chiba H, Hashimoto J, Sawada N, Takahashi H, Kuroki Y, Ariki S.

Biochem Biophys Res Commun. 2017 Mar 25;485(1):107-112. doi: 10.1016/j.bbrc.2017.02.028. Epub 2017 Feb 8.

PMID:28188794

12. Congenital Pulmonary Vein Atresia in an Adult.

Kuronuma K, Kobayashi T, Takahashi Y, Takahashi H.

Intern Med. 2017;56(7):877-878. doi: 10.2169/internalmedicine.56.7590. Epub 2017 Apr 1.

No abstract available.

PMID:28381760

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- 11 肺気腫の診断、病態に関する研究
- 12 HRC Tを用いた末梢気道立体構築の研究

OPub Med 掲載論文 (2016 年)

1. ESICM LIVES 2016: part three : Milan, Italy. 1-5 October 2016.

Intensive Care Med Exp. 2016 Sep;4(Suppl 1):28. doi: 10.1186/s40635-016-0100-7. No abstract available.

Yatabe T, Tamura T, Yamashita K, Yokoyama M, Idei M, Nomura T, Yamamoto N, Sakai Y, Yoshida T, Matsuda Y, Yamaguchi Y, Takaki S, Yamaguchi O, Goto T, Longani N, Medar S, Abdel-Aal IR, El Adawy AS, Mohammed HM, Mohamed AN, Parry SM, E, Hadley JS, Millar M, Hall D, Hewitt H, Yasuda H, Sanui M, Komuro T, Kawano S, Andoh K, Yamamoto H, Noda E, Hatakeyama J, Saitou N, Okamoto H, Kobayashi A, Takei T, Matsukubo S; JSEPTIC (Japanese Society of Education for Physicians and Trainees in Intensive Care) Clinical Trial Group., Rotzel HB, Lázaro AS, Sadamoto Y, Katabami K, Wada T, Ono Y, Maekawa K, Hayakawa M, Sawamura A, Gando S, Ohta Y, Sakuma J, Lee C, Hatib F, Jian Z, Buddi S, Cannesson M, Fileković S, Turel M, Knafelj R, Gorjup V, Stanić R, Gradišek P, Cerović O, Mirković T, Noč M, Maekawa K, Hayakawa M, Ono Y, Kodate A, Sadamoto Y, Tominaga N, Mizugaki A, Murakami H, Yoshida T, Katabami K, Wada T, Sawamura A, FR, SimóD, Hayashi K, Hirayama T, Yumoto T, Tsukahara K, Iida A, Nosaka N, Sato K, Ugawa T, Nakao A, Ujike Y, Hirohata S, Mojoli F, Torriglia F, Giannantonio M, Orlando A, Bianzina S, Tavazzi G, Mongodi S, Suzuki S, Mihara Y, Hikasa Y, Okahara S, Morimatsu H; Okayama Research Investigation Organizing Network Bodingbauer M, Yamaguchi Y, Nomura T, Yoshitake E, Idei M, Yoshida T, Takaki S, Yamaguchi O, Kaneko Wee S, Ong C, Lau YH, Wong Y, Banderas-Bravo ME, Olea-Jiménez V, Yoshihiro S, Sakuraya M, Hayakawa M, Hirata A, Kawamura N, Tsutui T, Yoshida K, Hashimoto Y, Takei T, Fushimi K, Iwamoto M, Nakagawa S,

Intensive Care Med Exp. 2016 Sep;4(Suppl 1):28. doi: 10.1186/s40635-016-0100-7. No abstract available.

PMID: 27686459 Free PMC Article

2. The novel heart-specific RING finger protein 207 is involved in energy metabolism in cardiomyocytes.

Mizushima W, Takahashi H, Watanabe M, Kinugawa S, Matsushima S, Takada S, Yokota T, Furihata T, Matsumoto J, Tsuda M, Chiba I, Nagashima S, Yanagi S, Matsumoto M, Nakayama KI, Tsutsui H, Hatakeyama S.

J Mol Cell Cardiol. 2016 Sep 25;100:43-53. doi: 10.1016/j.yjmcc.2016.09.013.

PMID: 27677939

3. Internal Movement of the Prostate During Intensity Modulated Radiation Therapy Detected by Using Cone Beam Computed Tomography Considering the Prostate and Bladder Volumes and Rectal Diameter.

Oguma Y, Matsuura T, Ishikawa K, Tatebe H, Fukuda K, Nishikawa T, Takahashi H, Okajima K.

Int J Radiat Oncol Biol Phys. 2016 Oct 1;96(2S):E275. doi: 10.1016/j.ijrobp.2016.06.1314. No abstract available.

PMID: 27674242

4. Response to: 'Could abatacept directly target expanded plasmablasts in IgG4-related disease?' by Alegria et al.

Yamamoto M, Takahashi H, Takano K, Himi T, Nakase H.

Ann Rheum Dis. 2016 Nov;75(11):e74. doi: 10.1136/annrheumdis-2016-210403. No abstract available.

PMID: 27651399

5. Validation of the Japanese disease severity classification and the GAP model in Japanese patients with idiopathic pulmonary fibrosis.

Kondoh S, Chiba H, Nishikiori H, Umeda Y, Kuronuma K, Otsuka M, Yamada G, Ohnishi H, Mori M, Kondoh Y, Taniguchi H, Homma S, Takahashi H.

Respir Investig. 2016 Sep;54(5):327-33. doi: 10.1016/j.resinv.2016.02.009. Epub 2016 Mar 30.

PMID: 27566380

6. Antithrombin Supplementation and Mortality in Sepsis-Induced Disseminated Intravascular Coagulation: A Multicenter Retrospective Observational Study.

Hayakawa M, Kudo D, Saito S, Uchino S, Yamakawa K, Iizuka Y, Sanui M, Takimoto K, Mayumi T, Ono K, Azuhata T, Ito F, Yoshihiro S, Hayakawa K, Nakashima T, Ogura T, Noda E, Nakamura Y, Sekine R, Yoshikawa Y, Sekino M, Ueno K, Okuda Y, Watanabe M, Tampo A, Saito N, Kitai Y, Takahashi H, Kobayashi I, Kondo Y, Matsunaga W, Nachi S, Miike T, Takahashi H, Takauji S, Umakoshi K, Todaka T, Kodaira H, Andoh K, Kasai T, Iwashita Y, Arai H, Murata M, Yamane M, Shiga K, Hori N.

Shock. 2016 Aug 19. [Epub ahead of print]

PMID: 27548460

7. Surfactant protein D is a useful biomarker for monitoring acute lung injury in rats.

Murata M, Otsuka M, Ashida N, Yamada G, Kuronuma K, Chiba H, Takahashi H.

Exp Lung Res. 2016 Aug;42(6):314-21. doi: 10.1080/01902148.2016.1215570.

PMID: 27541374

8. Eclampsia and hypertension: difficult to predict based solely on blood pressure monitoring. Matsubara S, Suzuki H, Baba Y, Takahashi H, Minakami H. *Acta Obstet Gynecol Scand*. 2016 Aug 19. doi: 10.1111/aogs.13001. [Epub ahead of print] No abstract available.
PMID: 27538383

9. Benefit profile of anticoagulant therapy in sepsis: a nationwide multicentre registry in Japan. Yamakawa K, Umemura Y, Hayakawa M, Kudo D, Sanui M, Takahashi H, Yoshikawa Y, Hamasaki T, Fujimi S; Japan Septic Disseminated Intravascular Coagulation (J-Septic DIC) study group. *Crit Care*. 2016 Jul 29;20(1):229. doi: 10.1186/s13054-016-1415-1.
PMID: 27472991 Free PMC Article

10. IgG4-Related Disease in Organs Other than the Hepatobiliary-Pancreatic System. Yamamoto M, Takahashi H. *Semin Liver Dis*. 2016 Aug;36(3):274-82. doi: 10.1055/s-0036-1584317. Epub 2016 Jul 28.
PMID: 27466796

11. Lung adenocarcinoma may be a more susceptible subtype to a dendritic cell-based cancer vaccine than other subtypes of non-small cell lung cancers: a multicenter retrospective analysis. Takahashi H, Shimodaira S, Ogasawara M, Ota S, Kobayashi M, Abe H, Morita Y, Nagai K, Tsujitani S, Okamoto M, Suzuki Y, Nakanishi Y, Yonemitsu Y; DC Vaccine Study Group at the Japanese Society of Immunotherapy and Cell Therapy. *Cancer Immunol Immunother*. 2016 Sep;65(9):1099-111. doi: 10.1007/s00262-016-1872-z. Epub 2016 Jul 22.
PMID: 27448677

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○主な研究内容

- 1 特発性間質性肺炎の診断、病態、治療に関する研究
- 2 肺サーファクタント蛋白の機能に関する研究
- 3 肺癌の診断、予後因子、治療に関する研究
- 4 肺癌の浸潤転移に関する分子生物学的研究
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- 8 膠原病の肺疾患の診断、病態、治療に関する研究
- 9 各種肺疾患の三次元的病態像の解析
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- 11 肺気腫の診断、病態に関する研究
- 12 HRC Tを用いた末梢気道立体構築の研究

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