厚生労働科学研究費補助金

循環器疾患・糖尿病等生活習慣病対策総合研究事業

バイオマーカーを用いた加熱式たばこによる受動喫煙の健康影響を評価するための研究

令和５（２０２３）年度　総合研究報告書

研究代表者　大森久光

令和６年（２０２４年）３月

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令和 5 年度　厚生労働科学研究費
（循環器疾患・糖尿病等生活習慣病対策総合研究事業）
総括研究報告書

バイオマーカーを用いた加熱式たばこによる受動喫煙の健康影響を評価するための研究

研究代表者　大森　久光　熊本大学

研究要旨

本研究は，加熱式たばこ喫煙者・受動喫煙者の健康影響を評価することを目的とした。本研究では、先行研究での受動喫煙の評価方法に加えて、曝露指標（尿中ニコチン代謝物)、臨床指標（炎症、呼吸機能）および肺への影響指標を分析し、加熱式たばこの使用によって影響の生じるバイオマーカーの抽出および選定を目指すことに特色がある。加熱式たばこによる受動喫煙が人の健康に及ぼす影響をバイオマーカーから予測し、評価することを最終目的とする。

本研究は、「**項目 1：加熱式たばこ使用による受動喫煙曝露の実態調査（長期間の臨床指標への影響の評価）」および「項目 2：呼吸器疾患につながりうる影響指標の探索**」で構成されている。

加熱式たばこ使用による受動喫煙曝露の実態を把握する（長期の臨床指標への影響を評価する）ために、熊本大学および協力機関の倫理委員会で承認後、1）令和 5 年 10 月 20 日より、健康診断（人間ドック）受診者を対象として、リクルートを開始した。

令和 6 年 1 月末までに 2,414 名の研究参加に同意した者の質問票の回収および余剰尿、余剰血清を採取した。回収した質問項目をエクセルに順次入力を行い、人間ドックデータを突合したデータベースを作成し、長期的な縦断研究の体制を構築した。**現時点、加熱式たばこによる受動喫煙者と考えられる 62 名が登録されている。**

また、微量血清サンプルからの miRNA プロファイリングの実験条件の設定のための検討を行った。微量血清サンプルからの miRNA プロファイリングの実験条件の設定のため、異なる被検者 12 名から取得した血清サンプル(200ml～1ml)から miRNA 抽出を行った。その結果、健診の残余検体で得られる検体量から本解析に必要な miRNA を分解することなく十分量取得できることが確認された。さらに、取得した miRNA サンプルから次世代シーケンサー解析用のライブラリーを作成し、そのサイズ分布及び定量解析から解析に適したライブラリーを作成できることを確認した。次世代シーケンサーによる 12 サンプルの解析から、各血清サンプル中の miRNA プロファイルの取得に成功した。

令和 6 年度以降、呼吸器疾患につながり得る指標であるフィブリン-3、SP-D 解析に加えて、次世代シーケンシング （NGS：Next Generation Sequencing） による miRNA の

発現プロファイル解析による病態モニタリングを行う。現在リクルートを進めている血清サンプルから、加熱式使用群、紙巻喫煙群、非喫煙群について、次世代シーケンサーによる miRNA プロファイリングを行う。取得した結果についてクラスター分析を行い、各群を分類しうるマーカーmiRNA を見出す。臨床指標（特に呼吸機能）の経年変化および炎症指標（高感度 C 反応性蛋白、白血球）との関連について分析する。肺胞破壊や COPD の病態マーカーである Fibulin-3 及び SP-D の解析を実施し、その結果及び健康診断の結果と miRNA を指標とした解析結果との相関性を解析することで、呼吸器疾患につながり得るバイオマーカーを見出し、その評価方法を開発する。

開発した方法を用いて、順次、加熱式たばこの使用および受動喫煙の評価を行う。さらに、国内で主に使用されている複数のメーカーの加熱式たばこについての解析を行う。

本研究成果の発信により、国民の加熱式たばこへの意識の変革および受動喫煙に対する認知の向上を促し、「改正健康増進法」で店内を喫煙可能としている施設において屋内禁煙化の選択を促すことにつながることが期待される。最終的には、本研究の成果が、受動喫煙による疾病および喫煙関連疾患の予防に貢献することが期待される。

研究分担者	所属機関名	職名
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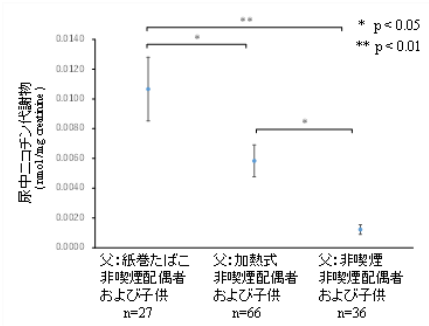
A. 研究目的

加熱式たばこによる受動喫煙の健康影響について明らかにすることが目的である。

これまで、紙巻たばこについては受動喫煙による健康影響が明らかになっているが、加熱式たばこについては主流煙にニコチンや発がん性物質が含まれていることは明らかであるものの、現時点では受動喫煙による長期的な健康影響を予測することは難しい状況である。 このような状況を踏まえ、健康増進法の改正において、加熱式たばこによる受動喫煙が人の健康に影響を及ぼす調査研究を一層推進し、可能な限り早期に結論を得るよう附帯決議がなされた。

申請者らは、日本医療研究開発機構（AMED）（2018–19 年）および厚生労働科学研究費補助金（2019–22 年）の助成を受け、父親が加熱式たばこを使用している非喫煙家族（配偶者と子供）の尿中ニコチン代謝物は、非喫煙・非使用家族に対して有意に高値を示し、受動喫煙による曝露の実態を明らかにした。（Onoue A, Omori H. et al. *Int J Environ Res Pub Health.* 2022, 19, 6275.）



本研究では、曝露指標に加えて、影響指標を用いた評価方法の開発を行い、加熱式たばこによる受動喫煙が人の健康に及ぼす影響をバイオマーカーから予測し、評価することを最終目的として、以下の４点について検討する。

① 呼吸機能、炎症、尿中のニコチン代謝物など、加熱式たばこの使用によって影響の生じるバイオマーカーの抽出・選定
② 疾患につながり得るバイオマーカーを用いた、加熱式たばこの喫煙及び受動喫煙の評価方法の開発
③ 開発した方法に基づいて実施する加熱式たばこの喫煙及び受動喫煙の評価
④ 国内で主に使用されている複数のメーカーの加熱式たばこについての解析

令和 5～6 年度に各バイオマーカーからの加熱式たばこによる受動喫煙の健康影響の予測の解析を行い、令和 7 年までにその評価の達成を目指す。

B. 研究方法

項目１：加熱式たばこ使用による受動喫煙曝露の実態調査（長期間の臨床指標への影響の評価）（全班員）

【研究対象者】

1-1：人間ドック受診者を対象として、加熱式たばこ使用者の呼吸機能の横断・縦断研究（前向き・後ろ向きコホート研究）の体制を構築し、曝露の実態調査を実施する。

1-2：紙巻たばこ、加熱式たばこ、紙巻および加熱式たばこ併用者およびその家族（配偶者、子供）を対象とした曝露の実態調査を実施する。

1-3：加熱式たばこ使用者およびその家族（AMED、厚労科研リクルート者）の追跡調査（前向きコホート研究）

【質問票による受動喫煙の分析】

喫煙状況（**加熱式たばこの種類**、紙巻たばこ喫煙歴等）、受動喫煙状況（**家族の目の前での使用、最終曝露日時**等）、自覚症状（咳、痰、**喘息症状**等）、身体状況（治療中疾患等）、**新規疾患の罹患状況**、職場の喫煙環境、加熱式たばこに関する知識等、先行研究の項目をさらに充実。

1－2、1－3 では、負担軽減費：1 人 2 千円。

【目標サンプル数の設定に関して】

先行研究がない状況であり、明確にサンプルサイズを説明することは困難な状況にあると考えられたため、1－2、1－3 のサンプルサイズは、先行研究と同様に、Cohen の計算法（Cohen, J. **1988**）により算出した。

Cohen, J. *Statistical power analysis for the behavioral sciences*, 2nd Edition.; Hillsdale, NJ: Lawrence Erlbaum Associates. 1988.

Cohen の計算法によると、3 群比較が主

の解析方法であり、有意水準を両側で 5%、効果量は 0.4 で、目指す検出力を 0.8 の場合、**それぞれ 21 以上が必要**であり、**目標症例数は、それぞれの群で 30 人以上とする。**

項目 2：呼吸器疾患につながりうる影響指標の探索
（首藤班員、河野研究協力者、大坪班員、橋本弘司研究協力者、黒澤班員、緒方班員、大森代表者）

【曝露指標の分析】

・尿中ニコチン代謝物: コチニン、3-ハイドロキシコチニン（3-OHCot）
依存性化学物質。過去 2-3 日間のニコチン曝露量を反映。

・尿中たばこ特異的ニトロソアミン類: NNAL: 4-(methylnitrosamino)-1- (3-pyridyl)-1-butanone
たばこ製品由来の発がん性物質。中期的な曝露マーカー（半減期 15-45 日）

・ liquid chromatography [LC] / mass spectrometry [MS] / [MS] (LC-MS/MS)を用いて測定する。

【呼吸器疾患につながり得る影響指標の分析】

・fibulin-3：生体を構成する殆ど全ての細胞が分泌する**細胞外小胞**が、細胞機能の制御や疾患の進展に大きく寄与することが報告され、現在、血液や尿等の体液を利用した疾患診断（**リキッドバイオプシー**）として、世界的に注目されている。慢性閉塞性肺疾患（COPD: Chronic Obstructive Pulmonary Disease）患者血清のプロテオミクス解析により、**エラスチン繊維の変性•破壊を反映する細胞外小胞タンパク質**として同定された**肺胞破壊（肺気腫）関連バイオマーカー**である。（Koba T, *ERJ open research*, 2020）

・**SP-D**：肺の II 型肺胞上皮細胞より産生•分泌される**肺特異的糖タンパク質**である。SP-D 遺伝子の SNPs(Rs721917, Met11Thr）が血清中の SD-P レベルの低下を招く、COPD、肺気腫と相関、血清 SP-D が喫煙者で高値、COPD 患者で高値などが報告されており、**COPD の病態バイオマーカー**である。

・**次世代シーケンシング （NGS：Next Generation Sequencing）** による miRNA の発現プロファイル解析による病態モニタリング

本研究は、横断研究の指針である 「The Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) statement」に従って実施した。Vandenvbroucke, J.P.et al. STROBE Initiative. Epidemiology 2007, 18, 805–835

本研究は、熊本大学倫理委員会の承認（倫理第 2150 号、令和 5 年 8 月 18 日承認）を得て実施した。

C. 結果

項目 1：加熱式たばこ使用による受動喫煙曝露の実態調査（長期間の臨床指標への影響の評価）（全班員）

加熱式たばこ使用による受動喫煙曝露の実態を把握する（長期の臨床指標への影響

を評価する）ために、熊本大学および協力機関の倫理委員会で承認後、１）令和 5 年 10 月 20 日より、健康診断（人間ドック）受診者を対象として、リクルートを開始した。

令和 6 年 1 月末までに 2,414 名の研究参加に同意した者の質問票の回収および余剰尿、余剰血清を採取した。

回収した質問項目をエクセルに順次入力を行い、人間ドックデータを突合したデータベースを作成し、長期的な縦断研究の体制を構築した。

現時点、加熱式たばこによる受動喫煙者と考えられる 62 名が登録されている。

項目２：呼吸器疾患につながりうる影響指標の探索
（首藤班員、河野研究協力者、大坪班員、橋本弘司研究協力者、黒澤班員、緒方班員、大森代表者）

【曝露指標の分析】

liquid chromatography [LC] / mass spectrometry [MS] / [MS] (LC-MS/MS)の手法の確立を行った。現在、余剰尿を分析中である。

【呼吸器疾患につながり得る影響指標の分析】

微量血清サンプルからの miRNA プロファイリングの実験条件の設定のための検討を行った。

微量血清サンプルからの miRNA プロファイリングの実験条件の設定のため、異なる被検者 12 名から取得した血清サンプル(200μl～1ml)から miRNA 抽出を行った。

その結果、**健診の残余検体で得られる検体量から本解析に必要な miRNA を分解することなく十分量取得できることが確認された。**

さらに、取得した miRNA サンプルから**次世代シーケンサー解析用のライブラリーを作成し、そのサイズ分布及び定量解析から解析に適したライブラリーを作成できることを確認した。**

次世代シーケンサーによる 12 サンプルの解析から、**各血清サンプル中の miRNA プロファイルの取得に成功した。**

D．考察

項目１：加熱式たばこ使用による受動喫煙曝露の実態調査（長期間の臨床指標への影響の評価）（全班員）

令和 6 年 1 月末までに 2,414 名の研究参加に同意した者の質問票の回収および余剰尿、余剰血清を採取した。回収した質問項目をエクセルに順次入力を行い、人間ドックデータを突合したデータベースを作成し、長期的な縦断研究の体制を構築した。**現時点、加熱式たばこによる受動喫煙者と考えられる 62 名を登録した。**

臨床指標（特に呼吸機能）の経年変化および炎症指標（高感度 C 反応性蛋白、白血球）との関連について分析を行っている。

令和 6 年度もリクルートを継続し対象者の増加を図る。加熱式たばこ使用による受動喫煙曝露の実態調査（項目１）を継続する。

また、1-2. 紙巻たばこ、加熱式たばこ、紙巻および加熱式たばこ併用者およびその家族（配偶者、子供）を対象とした曝露の実態調査、および 1-3. 我々の先行研究に参加し

た加熱式たばこ使用者およびその家族（AMED、厚労科研リクルート者）の追跡調査（前向きコホート研究）を実施する。

質問票により、特に喘息症状の有無、新規の発症疾患の有無を聴取する。

項目２：呼吸器疾患につながりうる影響指標の探索
（首藤班員、河野研究協力者、大坪班員、橋本弘司研究協力者、黒澤班員、緒方班員、大森代表者）

健診の残余検体で得られる検体量から本解析に必要な miRNA を分解することなく十分量取得できることが確認された。

さらに、取得した miRNA サンプルから次世代シーケンサー解析用のライブラリーを作成し、そのサイズ分布及び定量解析から解析に適したライブラリーを作成できることを確認した。

次世代シーケンサーによる 12 サンプルの解析から、各血清サンプル中の miRNA プロファイルの取得に成功した。

加熱式使用群、紙巻喫煙群、非喫煙群（受動喫煙の有無で分類）に分け、曝露指標、影響指標の測定を行う。呼吸器疾患につながり得る指標である**フィブリン-3、SP-D 解析**に加えて、**次世代シーケンシング（NGS：Next Generation Sequencing）による miRNA の発現プロファイル解析による病態モニタリングを行う。**

リクルートした血清サンプルから、加熱式使用群、紙巻喫煙群、非喫煙群について、次世代シーケンサーによる miRNA プロファイリングを行う。

取得した結果について**クラスター分析を行い、各群を分類しうるマーカーmiRNA を見出す。**

Ｅ．結語

健診の残余検体で得られる検体量から本解析に必要な miRNA を分解することなく十分量取得できることが確認された。

さらに、取得した miRNA サンプルから**次世代シーケンサー解析用のライブラリーを作成し、そのサイズ分布及び定量解析から解析に適したライブラリーを作成できることを確認した。**

次世代シーケンサーによる解析から、**各血清サンプル中の miRNA プロファイルの取得に成功した。**

現在リクルートを進めている血清サンプルから、加熱式使用群、紙巻喫煙群、非喫煙群について、次世代シーケンサーによる miRNA プロファイリングを行う。取得した結果についてクラスター分析を行い、各群を分類しうるマーカーmiRNA を見出す。

臨床指標（特に呼吸機能）の経年変化および炎症指標（高感度 C 反応性蛋白、白血球）との関連について分析する。

肺胞破壊や COPD の病態マーカーである Fibulin-3 及び SP-D の解析を実施し、その結果及び健康診断の結果と miRNA を指標とした解析結果との相関性を解析することで、呼吸器疾患につながり得るバイオマーカーを見出し、その**評価方法を開発**する。

開発した方法を用いて、順次、加熱式たばこの使用および受動喫煙の評価を行う（目的③）**。さらに、国内で主に使用されている複数のメーカーの加熱式たばこについての解析を行う**

F．健康危険情報

なし.

G．研究発表

1．論文発表（本研究に関連するもの）

なし.

2．学会発表

なし.

H．知的財産権の出願・登録状況

この研究において、知的財産権に該当するものはなかった。

資料１　血清 miRNA プロファイル（その他：参照）

令和 5 年度　厚生労働科学研究費

（循環器疾患・糖尿病等生活習慣病対策総合研究事業）

加熱式たばこ使用による受動喫煙曝露の実態調査（長期間の臨床指標への影響の評価）

分担研究報告書

研究代表者	大森久光	熊本大学	研究協力者	尾上あゆみ	熊本大学
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研究分担者	首藤　剛	熊本大学			

研究要旨

本研究は，加熱式たばこ喫煙者・受動喫煙者の健康影響を評価することを目的とした。本研究では、先行研究での受動喫煙の評価方法に加えて、曝露指標（尿中ニコチン代謝物)、臨床指標（炎症、呼吸機能）および肺への影響指標を分析し、加熱式たばこの使用によって影響の生じるバイオマーカーの抽出および選定を目指すことに特色がある。加熱式たばこによる受動喫煙が人の健康に及ぼす影響をバイオマーカーから予測し、評価することを最終目的とする。

本研究は、「**項目 1：加熱式たばこ使用による受動喫煙曝露の実態調査（長期間の臨床指標への影響の評価）」および「項目 2：呼吸器疾患につながりうる影響指標の探索」**で構成されている。本分担研究は項目 1 を担当する。

本分担研究では、加熱式たばこ使用による受動喫煙曝露の実態を把握する（長期の臨床指標への影響を評価する）ために、熊本大学および協力機関の倫理委員会で承認後、１）令和 5 年 10 月 20 日より、健康診断（人間ドック）受診者を対象として、リクルートを開始した。令和 6 年 1 月末までに 2,414 名の研究参加に同意した者の質問票の回収および余剰尿、余剰血清を採取した。回収した質問項目をエクセルに順次入力を行い、人間ドックデータを突合したデータベースを作成し、長期的な縦断研究の体制を構築した。現時点、加熱式たばこによる受動喫煙者と考えられる 62 名が登録されている。

臨床指標（特に呼吸機能）の経年変化および炎症指標（高感度 C 反応性蛋白、白血球）との関連について分析を行っている。令和 6 年度もリクルートを継続し対象者の増加を図る。加熱式たばこ使用による受動喫煙曝露の実態調査を継続する。

A. 研究目的

加熱式たばこによる受動喫煙の健康影響について明らかにすることが目的である。これまで、紙巻たばこについては受動喫煙による健康影響が明らかになっているが、加熱式たばこについては主流煙にニコチンや発がん性物質が含まれていることは明らかであるものの、現時点では受動喫煙によ

る長期的な健康影響を予測することは難しい状況である。 このような状況を踏まえ、健康増進法の改正において、加熱式たばこによる受動喫煙が人の健康に影響を及ぼす調査研究を一層推進し、可能な限り早期に結論を得るよう附帯決議がなされた。

申請者らは、日本医療研究開発機構(AMED)(2018-19 年)および厚生労働科学研究費補助金 (2019-22 年) の助成を受け、父親が加熱式たばこを使用している非喫煙家族 (配偶者と子供)の尿中ニコチン代謝物は、非喫煙・非使用家族に対して有意に高値を示し、受動喫煙による曝露の実態を明らかにした。(Onoue A, Omori H. et al. *Int J Environ Res Pub Health.* 2022, 19, 6275.)

本研究では、曝露指標に加えて、影響指標を用いた評価方法の開発を行い、加熱式たばこによる受動喫煙が人の健康に及ぼす影響をバイオマーカーから予測し、評価することを最終目的として、以下の 4 点について検討する。

① 呼吸機能、炎症、尿中のニコチン代謝物など、加熱式たばこの使用によって影響の生じるバイオマーカーの抽出・選定
② 疾患につながり得るバイオマーカーを用いた、加熱式たばこの喫煙及び受動喫煙の評価方法の開発
③ 開発した方法に基づいて実施する加熱式たばこの喫煙及び受動喫煙の評価
④ 国内で主に使用されている複数のメーカーの加熱式たばこについての解析

令和 5～6 年度に各バイオマーカーからの加熱式たばこによる受動喫煙の健康影響の予測の解析を行い、令和 7 年までにその評価の達成を目指す。

B. 研究方法

C. 項目 1 ：加熱式たばこ使用による受動喫煙曝露の実態調査（長期間の臨床指標への影響の評価）（全班員）

【研究対象者】

1-1：人間ドック受診者を対象として、加熱式たばこ使用者の呼吸機能の横断・縦断研究(前向き・後ろ向きコホート研究)の体制を構築し、曝露の実態調査を実施する。

1-2：紙巻たばこ、加熱式たばこ、紙巻および加熱式たばこ併用者およびその家族（配偶者、子供)を対象とした曝露の実態調査を実施する。

1-3：加熱式たばこ使用者およびその家族（AMED、厚労科研リクルート者）の追　跡調査（前向きコホート研究）

【質問票による受動喫煙の分析】

喫煙状況(**加熱式たばこの種類**、紙巻たばこ喫煙歴等)、受動喫煙状況(**家族の目の前での使用、最終曝露日時**等)、自覚症状(咳、痰、**喘息症状**等)、身体状況(治療中疾患等)、**新規疾患の罹患状況**、職場の喫煙環境、加熱式たばこに関する知識等、先行研究の項目をさらに充実。

1－2、1－3 では、負担軽減費：1 人 2 千円。

【目標サンプル数の設定に関して】

先行研究がない状況であり、明確にサンプルサイズを説明することは困難な状況にあると考えられたため、1－2、1－3 のサン

プルサイズは、先行研究と同様に、Cohen の計算法（Cohen, J. 1988）により算出した。

Cohen, J. *Statistical power analysis for the behavioral sciences*, 2nd Edition.; Hillsdale, NJ: Lawrence Erlbaum Associates. 1988.

Cohen の計算法によると、3群比較が主の解析方法であり、有意水準を両側で5%、効果量は0.4で、目指す検出力を0.8の場合、**それぞれ21以上が必要**であり、**目標症例数は、それぞれの群で30人以上とする。**

D．結果

項目１：加熱式たばこ使用による受動喫煙曝露の実態調査（長期間の臨床指標への影響の評価）（全班員）

加熱式たばこ使用による受動喫煙曝露の実態を把握する（長期の臨床指標への影響を評価する）ために、熊本大学および協力機関の倫理委員会で承認後、１）令和5年10月20日より、健康診断（人間ドック）受診者を対象として、リクルートを開始した。

令和6年1月末までに2,414名の研究参加に同意した者の質問票の回収および余剰尿、余剰血清を採取した。

回収した質問項目をエクセルに順次入力を行い、人間ドックデータを突合したデータベースを作成し、長期的な縦断研究の体制を構築した。

現時点、加熱式たばこによる受動喫煙者と考えられる62名が登録されている。

D．考察

項目１：加熱式たばこ使用による受動喫煙曝露の実態調査（長期間の臨床指標への影響の評価）（全班員）

令和6年1月末までに2,414名の研究参加に同意した者の質問票の回収および余剰尿、余剰血清を採取した。回収した質問項目をエクセルに順次入力を行い、人間ドックデータを突合したデータベースを作成し、長期的な縦断研究の体制を構築した。**現時点、加熱式たばこによる受動喫煙者と考えられる62名を登録した。**

臨床指標（特に呼吸機能）の経年変化および炎症指標（高感度C反応性蛋白、白血球）との関連について分析を行っている。

令和6年度もリクルートを継続し対象者の増加を図る。加熱式たばこ使用による受動喫煙曝露の実態調査（項目１）を継続する。

また、1-2.紙巻たばこ、加熱式たばこ、紙巻および加熱式たばこ併用者およびその家族（配偶者、子供）を対象とした曝露の実態調査、および1-3.我々の先行研究に参加した加熱式たばこ使用者およびその家族（AMED、厚労科研リクルート者）の追跡調査（前向きコホート研究）を実施する。

質問票により、特に喘息症状の有無、新規の発症疾患の有無を聴取する。

E．結語

F．健康危険情報

なし.

G．研究発表

１．論文発表（本研究に関連するもの）

なし．

２．学会発表

なし．

H．知的財産権の出願・登録状況

この研究において、知的財産権に該当するものはなかった。

令和 5 年度　厚生労働科学研究費
（循環器疾患・糖尿病等生活習慣病対策総合研究事業）

呼吸器疾患につながりうる影響指標の探索

分担研究報告書

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研究分担者　緒方裕光　　女子栄養大学
研究代表者　大森久光　　熊本大学
研究協力者　橋本弘司　　熊本大学
研究協力者　河野圭亮　　熊本大学

研究要旨

本研究は，加熱式たばこ喫煙者・受動喫煙者の健康影響を評価することを目的とした。本研究では、先行研究での受動喫煙の評価方法に加えて、曝露指標（尿中ニコチン代謝物)、臨床指標（炎症、呼吸機能）および肺への影響指標を分析し、加熱式たばこの使用によって影響の生じるバイオマーカーの抽出および選定を目指すことに特色がある。加熱式たばこによる受動喫煙が人の健康に及ぼす影響をバイオマーカーから予測し、評価することを最終目的とする。

本研究は、「**項目 1：加熱式たばこ使用による受動喫煙曝露の実態調査（長期間の臨床指標への影響の評価）」および「項目 2：呼吸器疾患につながりうる影響指標の探索」**で構成されている。本分担研究は項目 2 を担当する。

本分担研究では、微量血清サンプルからの miRNA プロファイリングの実験条件の設定のための検討を行った。微量血清サンプルからの miRNA プロファイリングの実験条件の設定のため、異なる被検者 12 名から取得した血清サンプル(200ml～1ml)から miRNA 抽出を行った。その結果、健診の残余検体で得られる検体量から本解析に必要な miRNA を分解することなく十分量取得できることが確認された。さらに、取得した miRNA サンプルから次世代シーケンサー解析用のライブラリーを作成し、そのサイズ分布及び定量解析から解析に適したライブラリーを作成できることを確認した。次世代シーケンサーによる 12 サンプルの解析から、各血清サンプル中の miRNA プロファイルの取得に成功した。

令和 6 年度以降、呼吸器疾患につながり得る指標であるフィブリン-3、SP-D 解析に加えて、次世代シーケンシング （NGS：Next Generation Sequencing） による miRNA の発現プロファイル解析による病態モニタリングを行う。現在リクルートを進めている血清サンプルから、加熱式使用群、紙巻喫煙群、非喫煙群について、次世代シーケンサーによる miRNA プロファイリングを行う。取得した結果についてクラスター分析を行い、各群を分類しうるマーカーmiRNA を見出す。臨床指標（特に呼吸機能）の経年変化および炎症指標（高感度 C 反応性蛋白、白血球）との関連について分析する。肺胞破壊や COPD

の病態マーカーである Fibulin-3 及び SP-D の解析を実施し、その結果及び健康診断の結果と miRNA を指標とした解析結果との相関性を解析することで、呼吸器疾患につながり得るバイオマーカーを見出し、その評価方法を開発する。

開発した方法を用いて、順次、加熱式たばこの使用および受動喫煙の評価を行う。さらに、国内で主に使用されている複数のメーカーの加熱式たばこについての解析を行う。

A. 研究目的

加熱式たばこによる受動喫煙の健康影響について明らかにすることが目的である。これまで、紙巻たばこについては受動喫煙による健康影響が明らかになっているが、加熱式たばこについては主流煙にニコチンや発がん性物質が含まれていることは明らかであるものの、現時点では受動喫煙による長期的な健康影響を予測することは難しい状況である。 このような状況を踏まえ、健康増進法の改正において、加熱式たばこによる受動喫煙が人の健康に影響を及ぼす調査研究を一層推進し、可能な限り早期に結論を得るよう附帯決議がなされた。

申請者らは、日本医療研究開発機構(AMED)(2018–19 年)および厚生労働科学研究費補助金 (2019–22 年) の助成を受け、父親が加熱式たばこを使用している非喫煙家族 (配偶者と子供)の尿中ニコチン代謝物は、非喫煙・非使用家族に対して有意に高値を示し、受動喫煙による曝露の実態を明らかにした。(Onoue A, Omori H. et al. *Int J Environ Res Pub Health.* 2022, 19, 6275.)

本研究では、曝露指標に加えて、影響指標を用いた評価方法の開発を行い、加熱式たばこによる受動喫煙が人の健康に及ぼす影響をバイオマーカーから予測し、評価することを最終目的として、以下の 4 点について検討する。

① 呼吸機能、炎症、尿中のニコチン代謝物など、加熱式たばこの使用によって影響の生じるバイオマーカーの抽出・選定
② 疾患につながり得るバイオマーカーを用いた、加熱式たばこの喫煙及び受動喫煙の評価方法の開発
③ 開発した方法に基づいて実施する加熱式たばこの喫煙及び受動喫煙の評価
④ 国内で主に使用されている複数のメーカーの加熱式たばこについての解析

令和 5～6 年度に各バイオマーカーからの加熱式たばこによる受動喫煙の健康影響の予測の解析を行い、令和 7 年までにその評価の達成を目指す。

B. 研究方法

項目 2：呼吸器疾患につながりうる影響指標の探索
（首藤班員、河野研究協力者、大坪班員、橋本弘司研究協力者、黒澤班員、緒方班員、大森代表者）

【曝露指標の分析】

・尿中ニコチン代謝物: コチニン、3-ハイドロキシコチニン（3-OHCot）

依存性化学物質。過去 2-3 日間のニコチ

ン曝露量を反映。

・**尿中たばこ特異的ニトロソアミン類: NNAL: 4-(methylnitrosamino)-1- (3-pyridyl)-1-butanone**

たばこ製品由来の発がん性物質。中期的な曝露マーカー（半減期 15-45 日）

・ liquid chromatography [LC] / mass spectrometry [MS] / [MS] (LC-MS/MS)を用いて測定する。

【呼吸器疾患につながり得る影響指標の分析】

・fibulin-3：生体を構成する殆ど全ての細胞が分泌する**細胞外小胞**が、細胞機能の制御や疾患の進展に大きく寄与することが報告され、現在、血液や尿等の体液を利用した疾患診断（**リキッドバイオプシー**）として、世界的に注目されている。慢性閉塞性肺疾患（COPD: Chronic Obstructive Pulmonary Disease）患者血清のプロテオミクス解析により、**エラスチン繊維の変性•破壊を反映する細胞外小胞タンパク質**として同定された**肺胞破壊（肺気腫）関連バイオマーカー**である。（Koba T, *ERJ open research*, 2020）

・**SP-D**：肺の II 型肺胞上皮細胞より産生•分泌される**肺特異的糖タンパク質**である。SP-D 遺伝子の SNPs(Rs721917, Met11Thr）が血清中の SD-P レベルの低下を招く、COPD、肺気腫と相関、血清 SP-D が喫煙者で高値、COPD 患者で高値などが報告されており、**COPD の病態バイオマーカー**である。

・**次世代シーケンシング （NGS：Next Generation Sequencing）** による miRNA の発現プロファイル解析による病態モニタリング

C. 結果

項目２：呼吸器疾患につながりうる影響指標の探索

（首藤班員、河野研究協力者、大坪班員、橋本弘司研究協力者、黒澤班員、緒方班員、大森代表者）

【曝露指標の分析】

liquid chromatography [LC] / mass spectrometry [MS] / [MS] (LC-MS/MS)の手法の確立を行った。現在、余剰尿を分析中である。

【呼吸器疾患につながり得る影響指標の分析】

微量血清サンプルからの miRNA プロファイリングの実験条件の設定のための検討を行った。

微量血清サンプルからの miRNA プロファイリングの実験条件の設定のため、異なる被検者 12 名から取得した血清サンプル(200μl～1ml)から miRNA 抽出を行った。

その結果、**健診の残余検体で得られる検体量から本解析に必要な miRNA を分解することなく十分量取得できることが確認された。**

さらに、取得した miRNA サンプルから**次世代シーケンサー解析用のライブラリーを作成し、そのサイズ分布及び定量解析から解析に適したライブラリーを作成できることを確認した。**

次世代シーケンサーによる 12 サンプルの解析から、**各血清サンプル中の miRNA プ**

ロファイルの取得に成功した。

呼吸器疾患につながり得る指標であるフィブリン-3、SP-D 解析に加えて、次世代シーケンシング （NGS：Next Generation Sequencing） による miRNA の発現プロファイル解析による病態モニタリングを行う。

現在リクルートを進めている血清サンプルから、加熱式使用群、紙巻喫煙群、非喫煙群について、次世代シーケンサーによる miRNA プロファイリングを行う。取得した結果についてクラスター分析を行い、各群を分類しうるマーカーmiRNA を見出す。

臨床指標（特に呼吸機能）の経年変化および炎症指標（高感度 C 反応性蛋白、白血球）との関連について分析する。

肺胞破壊や COPD の病態マーカーである Fibulin-3 及び SP-D の解析を実施し、その結果及び健康診断の結果と miRNA を指標とした解析結果との相関性を解析することで、呼吸器疾患につながり得るバイオマーカーを見出し、その**評価方法を開発**する。

開発した方法を用いて、順次、加熱式たばこの使用および受動喫煙の評価を行う（目的③）**。さらに、国内で主に使用されている複数のメーカーの加熱式たばこについての解析を行う**（目的④）**。**（全班員）

研究成果は順次、学会発表、論文を作成し公表する。

E．結語

健診の残余検体で得られる検体量から本解析に必要な miRNA を分解することなく十分量取得できることが確認された。

さらに、取得した miRNA サンプルから次世代シーケンサー解析用のライブラリーを作成し、そのサイズ分布及び定量解析から解析に適したライブラリーを作成できることを確認した。

次世代シーケンサーによる解析から、各血清サンプル中の miRNA プロファイルの取得に成功した。

現在リクルートを進めている血清サンプルから、加熱式使用群、紙巻喫煙群、非喫煙群について、次世代シーケンサーによる miRNA プロファイリングを行う。取得した結果についてクラスター分析を行い、各群を分類しうるマーカーmiRNA を見出す。

臨床指標（特に呼吸機能）の経年変化および炎症指標（高感度 C 反応性蛋白、白血球）との関連について分析する。

肺胞破壊や COPD の病態マーカーである Fibulin-3 及び SP-D の解析を実施し、その結果及び健康診断の結果と miRNA を指標とした解析結果との相関性を解析することで、呼吸器疾患につながり得るバイオマーカーを見出し、その**評価方法を開発**する。

開発した方法を用いて、順次、加熱式たばこの使用および受動喫煙の評価を行う（目的③）**。さらに、国内で主に使用されている複数のメーカーの加熱式たばこについての解析を行う**

F．健康危険情報

なし.

G．研究発表

1．論文発表（本研究に関連するもの）

なし.

2．学会発表

なし.

H．知的財産権の出願・登録状況

この研究において、知的財産権に該当するものはなかった。

資料１　血清 miRNA プロファイル（その他：参照）

その他　資料1　血清miRNAプロファイル

Sample#

miRNA levels
high to low

1	
ClustersPF	1035655
hsa-miR-486-5p	125687
hsa-miR-16-5p	119247
hsa-let-7b-5p	19574
hsa-miR-92a-3p	12513
hsa-miR-451a	11415
hsa-miR-22-3p	10666
hsa-miR-25-3p	6452
hsa-let-7i-5p	5482
hsa-miR-320a	2150
hsa-miR-26b-5p	1975
hsa-miR-26a-5p	1842
hsa-miR-423-5p	1840
hsa-miR-181a-5p	1819
hsa-miR-192-5p	1551
hsa-miR-186-5p	1349
hsa-miR-191-5p	1060
hsa-miR-21-5p	1024
hsa-miR-144-3p	769
hsa-miR-93-5p	767
hsa-miR-142-5p	766
hsa-let-7f-5p	661
hsa-miR-146a-5p	621
hsa-miR-30d-5p	602
hsa-miR-126-5p	591
hsa-let-7a-5p	577
hsa-let-7g-5p	519
hsa-miR-151a-5p	479
hsa-miR-182-5p	348
hsa-miR-126-3p	332
hsa-miR-486-3p	322
hsa-miR-20a-5p	309
hsa-miR-17-5p	304
hsa-miR-103a-3p	266
hsa-miR-148a-3p	264
hsa-miR-4732-5p	261
hsa-miR-24-3p	240
hsa-miR-140-3p	235
hsa-miR-185-5p	223
hsa-miR-101-3p	200
hsa-miR-144-5p	184
hsa-miR-146b-5p	166
hsa-miR-423-3p	137
hsa-miR-4732-3p	132
hsa-miR-106b-5p	127
hsa-miR-7-5p	114
hsa-miR-143-3p	113
hsa-miR-30b-5p	111
hsa-miR-151a-3p	103
hsa-miR-122-5p	97
hsa-miR-98-5p	95
hsa-miR-378a-3p	88
hsa-miR-15a-5p	86
hsa-miR-532-5p	85
hsa-miR-28-3p	83
hsa-miR-15b-5p	78
hsa-let-7d-5p	77
hsa-miR-30e-5p	75
hsa-miR-223-3p	67
hsa-miR-660-5p	66
hsa-miR-629-5p	62
hsa-miR-20b-5p	58
hsa-miR-106a-5p	56
hsa-miR-484	54
hsa-miR-221-3p	43
hsa-miR-425-5p	43
hsa-miR-181c-5p	42
hsa-miR-194-5p	42
hsa-miR-454-3p	40
hsa-miR-301a-3p	37
hsa-miR-27b-3p	35
hsa-miR-148b-3p	35
hsa-miR-1180-3p	34
hsa-miR-128-3p	33
hsa-miR-150-5p	30
hsa-let-7c-5p	29
hsa-miR-1255b-5p	29
hsa-miR-183-5p	27
hsa-miR-409-3p	26
hsa-miR-652-3p	25
hsa-miR-155-5p	23
hsa-miR-21-3p	22
hsa-miR-181b-5p	22
hsa-miR-30c-5p	21
hsa-miR-340-5p	20
hsa-miR-18a-5p	16
hsa-miR-27a-3p	16
hsa-miR-363-3p	16
hsa-miR-548k	16
hsa-let-7d-3p	15
hsa-miR-132-3p	15
hsa-miR-432-5p	15
hsa-miR-374a-5p	14
hsa-miR-1307-5p	14
hsa-miR-30e-3p	13
hsa-miR-941	13
hsa-miR-342-3p	12
hsa-miR-196b-5p	12
hsa-miR-148a-5p	11
hsa-miR-222-3p	10
hsa-miR-152-3p	10
hsa-miR-410-3p	10
hsa-miR-3200-3p	10
hsa-miR-411-5p	9
hsa-miR-744-5p	9
hsa-miR-19b-3p	8
hsa-miR-769-5p	8
hsa-miR-2110	8
hsa-miR-3688-3p	8
hsa-miR-382-5p	7
hsa-miR-342-5p	7
hsa-miR-210-3p	6
hsa-miR-29c-3p	6
hsa-miR-424-3p	6
hsa-miR-625-5p	6
hsa-miR-23a-3p	5
hsa-miR-28-5p	5
hsa-miR-100-5p	5
hsa-miR-199a-5p	5
hsa-miR-199a-3p	5
hsa-miR-361-5p	5
hsa-miR-381-3p	5
hsa-miR-485-3p	5
hsa-miR-493-5p	5
hsa-miR-550a-3p	5
hsa-miR-301b-3p	5
hsa-miR-3200-5p	5
hsa-miR-5010-5p	5
hsa-miR-6734-5p	5
hsa-miR-7706	5
hsa-miR-130a-3p	4
hsa-miR-134-5p	4
hsa-miR-324-3p	4
hsa-miR-590-3p	4
hsa-miR-421	4
hsa-miR-320b	4
hsa-miR-942-5p	4
hsa-miR-1270	4
hsa-miR-6513-3p	4
hsa-let-7e-5p	3
hsa-miR-19a-3p	3
hsa-miR-25-5p	3
hsa-miR-30a-3p	3
hsa-miR-181a-3p	3
hsa-miR-142-3p	3
hsa-miR-190a-5p	3
hsa-miR-296-5p	3
hsa-miR-130b-3p	3
hsa-miR-375	3
hsa-miR-377-5p	3
hsa-miR-328-3p	3
hsa-miR-425-3p	3
hsa-miR-431-5p	3
hsa-miR-92b-3p	3
hsa-miR-576-5p	3
hsa-miR-625-3p	3
hsa-miR-627-5p	3
hsa-miR-654-5p	3
hsa-miR-374b-5p	3
hsa-miR-1294	3
hsa-miR-548ae-5p	3
hsa-let-7a-3p	2
hsa-miR-17-3p	2
hsa-miR-92a-1-5p	2
hsa-miR-93-3p	2
hsa-miR-96-5p	2
hsa-miR-16-2-3p	2
hsa-miR-1-3p	2
hsa-miR-140-5p	2
hsa-miR-127-3p	2
hsa-miR-106b-3p	2
hsa-miR-200a-3p	2
hsa-miR-363-5p	2
hsa-miR-369-3p	2
hsa-miR-374a-3p	2
hsa-miR-379-5p	2
hsa-miR-339-5p	2
hsa-miR-345-5p	2
hsa-miR-433-3p	2
hsa-miR-146b-3p	2
hsa-miR-493-3p	2
hsa-miR-501-3p	2
hsa-miR-505-3p	2
hsa-miR-576-3p	2
hsa-miR-939-5p	2
hsa-miR-548j-5p	2
hsa-miR-1287-5p	2
hsa-miR-1255a	2
hsa-miR-3187-3p	2
hsa-miR-3615	2
hsa-miR-548ad-5p	2
hsa-miR-4446-3p	2
hsa-miR-4504	2
hsa-miR-4646-5p	2
hsa-miR-4677-3p	2
hsa-miR-4728-3p	2
hsa-miR-4746-5p	2
hsa-miR-6741-3p	2
hsa-miR-6805-5p	2
hsa-miR-18a-3p	1
hsa-miR-29a-3p	1
hsa-miR-101-5p	1
hsa-miR-196a-5p	1
hsa-miR-139-3p	1
hsa-miR-181a-2-3p	1
hsa-miR-199b-3p	1
hsa-miR-210-5p	1
hsa-miR-125b-5p	1
hsa-miR-132-5p	1
hsa-miR-191-3p	1
hsa-miR-125a-5p	1
hsa-miR-136-3p	1
hsa-miR-195-5p	1
hsa-miR-370-3p	1
hsa-miR-323a-3p	1

2	
ClustersPF	2045928
hsa-miR-16-5p	183011
hsa-miR-486-5p	164111
hsa-let-7b-5p	37153
hsa-miR-451a	17500
hsa-miR-22-3p	16573
hsa-miR-92a-3p	11168
hsa-miR-26a-5p	9843
hsa-miR-25-3p	9457
hsa-let-7f-5p	8301
hsa-let-7a-5p	7831
hsa-miR-146a-5p	7129
hsa-miR-320a	6098
hsa-let-7i-5p	5805
hsa-miR-21-5p	3438
hsa-miR-192-5p	3431
hsa-miR-423-5p	3171
hsa-miR-26b-5p	2956
hsa-miR-181a-5p	2918
hsa-miR-191-5p	2848
hsa-miR-122-5p	2694
hsa-miR-126-5p	2230
hsa-miR-142-5p	1659
hsa-miR-24-3p	1468
hsa-miR-186-5p	1164
hsa-miR-93-5p	1149
hsa-miR-148a-3p	1096
hsa-miR-151a-5p	1079
hsa-miR-126-3p	950
hsa-let-7g-5p	913
hsa-miR-30d-5p	902
hsa-miR-144-3p	866
hsa-miR-143-3p	765
hsa-miR-20a-5p	724
hsa-let-7d-5p	601
hsa-miR-4732-3p	598
hsa-miR-486-3p	574
hsa-miR-151a-3p	535
hsa-miR-146b-5p	522
hsa-miR-17-5p	504
hsa-miR-101-3p	500
hsa-miR-140-3p	470
hsa-miR-423-3p	432
hsa-miR-28-3p	397
hsa-miR-98-5p	376
hsa-miR-484	376
hsa-miR-103a-3p	374
hsa-miR-27b-3p	342
hsa-miR-144-5p	328
hsa-miR-182-5p	312
hsa-miR-30b-5p	308
hsa-miR-185-5p	295
hsa-miR-4732-5p	289
hsa-miR-106b-5p	268
hsa-let-7e-5p	255
hsa-miR-1-3p	217
hsa-miR-378a-3p	213
hsa-miR-221-3p	181
hsa-miR-409-3p	168
hsa-let-7c-5p	167
hsa-miR-27a-3p	163
hsa-miR-128-3p	148
hsa-miR-425-5p	148
hsa-miR-340-5p	146
hsa-miR-181c-5p	143
hsa-miR-223-3p	141
hsa-miR-150-5p	138
hsa-miR-660-5p	135
hsa-miR-30e-5p	116
hsa-miR-629-5p	116
hsa-let-7d-3p	113
hsa-miR-15b-5p	113
hsa-miR-20b-5p	95
hsa-miR-155-5p	83
hsa-miR-15a-5p	81
hsa-miR-432-5p	80
hsa-miR-941	80
hsa-miR-7-5p	78
hsa-miR-30e-3p	77
hsa-miR-148b-3p	70
hsa-miR-532-5p	68
hsa-miR-375	66
hsa-miR-30a-5p	65
hsa-miR-301a-3p	65
hsa-miR-148a-5p	63
hsa-miR-769-5p	63
hsa-miR-28-5p	61
hsa-miR-125a-5p	61
hsa-miR-744-5p	61
hsa-miR-23a-3p	59
hsa-miR-1180-3p	54
hsa-miR-181b-5p	51
hsa-miR-411-5p	42
hsa-miR-197-3p	39
hsa-miR-194-5p	39
hsa-miR-1307-5p	39
hsa-miR-652-3p	37
hsa-miR-21-3p	35
hsa-miR-152-3p	34
hsa-miR-106a-5p	33
hsa-miR-199a-3p	33
hsa-miR-1255b-5p	30
hsa-miR-134-5p	29
hsa-miR-485-3p	29
hsa-miR-92b-3p	28
hsa-miR-130a-3p	27
hsa-miR-132-3p	27
hsa-miR-1270	27
hsa-miR-328-3p	26
hsa-miR-625-3p	26
hsa-miR-548k	25
hsa-miR-342-3p	24
hsa-miR-222-3p	23
hsa-miR-454-3p	23
hsa-miR-382-5p	22
hsa-miR-425-3p	22
hsa-miR-199b-3p	21
hsa-miR-4446-3p	21
hsa-miR-4677-3p	21
hsa-miR-361-5p	20
hsa-miR-381-3p	20
hsa-miR-335-3p	20
hsa-miR-410-3p	19
hsa-miR-5010-5p	19
hsa-miR-139-3p	18
hsa-miR-3158-3p	18
hsa-miR-433-3p	17
hsa-miR-181d-5p	17
hsa-miR-3615	17
hsa-miR-18a-5p	16
hsa-miR-25-5p	16
hsa-miR-183-5p	16
hsa-miR-493-5p	16
hsa-miR-487b-3p	16
hsa-miR-181a-2-3p	15
hsa-miR-205-5p	15
hsa-miR-483-5p	15
hsa-miR-30c-5p	14
hsa-miR-320b	14
hsa-miR-19b-3p	13
hsa-miR-374a-3p	13
hsa-miR-576-5p	13
hsa-miR-3200-3p	13
hsa-miR-136-3p	12
hsa-miR-1908-5p	12
hsa-miR-29a-3p	11
hsa-miR-181a-3p	11
hsa-miR-215-5p	11
hsa-miR-99b-5p	11
hsa-miR-493-3p	11
hsa-miR-301b-3p	11
hsa-miR-1287-5p	11
hsa-miR-6741-3p	10
hsa-miR-30a-3p	9
hsa-miR-29c-3p	9
hsa-miR-93-3p	8
hsa-miR-363-3p	8
hsa-miR-374a-5p	8
hsa-miR-196b-5p	8
hsa-miR-421	8
hsa-miR-374b-5p	8
hsa-miR-199a-5p	7
hsa-miR-378a-5p	7
hsa-miR-342-5p	7
hsa-miR-339-5p	7
hsa-miR-485-5p	7
hsa-miR-766-5p	7
hsa-miR-1285-3p	7
hsa-miR-140-5p	6
hsa-miR-206	6
hsa-miR-379-5p	6
hsa-miR-339-3p	6
hsa-miR-574-3p	6
hsa-miR-766-3p	6
hsa-miR-2110	6
hsa-let-7a-3p	5
hsa-miR-100-3p	5
hsa-miR-107	5
hsa-miR-9-5p	5
hsa-miR-499a-5p	5
hsa-miR-509-3p	5
hsa-miR-550a-3p	5
hsa-miR-590-3p	5
hsa-miR-654-5p	5
hsa-miR-885-5p	5
hsa-miR-139-5p	4
hsa-miR-210-3p	4
hsa-miR-15b-3p	4
hsa-miR-195-5p	4
hsa-miR-130b-3p	4
hsa-miR-337-3p	4
hsa-miR-323a-3p	4
hsa-miR-338-3p	4
hsa-miR-345-5p	4
hsa-miR-146b-3p	4
hsa-miR-576-3p	4
hsa-miR-628-3p	4
hsa-miR-454-5p	4
hsa-miR-889-3p	4
hsa-miR-1228-5p	4
hsa-miR-3200-5p	4
hsa-miR-3605-5p	4
hsa-miR-3617-5p	4
hsa-miR-4781-3p	4
hsa-miR-6805-5p	4
hsa-miR-4433b-5p	4
hsa-let-7g-3p	3
hsa-miR-150-3p	3
hsa-miR-335-5p	3
hsa-miR-424-3p	3
hsa-miR-449a	3
hsa-miR-500a-3p	3
hsa-miR-501-3p	3
hsa-miR-532-3p	3
hsa-miR-551b-3p	3

3	
ClustersPF	1614891
hsa-miR-486-5p	160644
hsa-miR-16-5p	121530
hsa-let-7b-5p	21824
hsa-miR-22-3p	20863
hsa-miR-146a-5p	19258
hsa-miR-92a-3p	11598
hsa-miR-451a	9444
hsa-miR-26a-5p	9421
hsa-miR-25-3p	8533
hsa-let-7a-5p	6015
hsa-let-7f-5p	5785
hsa-miR-320a	5265
hsa-miR-191-5p	4306
hsa-let-7i-5p	4305
hsa-miR-21-5p	4282
hsa-miR-24-3p	4212
hsa-miR-423-5p	4105
hsa-miR-126-5p	3558
hsa-miR-192-5p	3355
hsa-miR-26b-5p	2904
hsa-miR-122-5p	2802
hsa-miR-181a-5p	2485
hsa-miR-148a-3p	2460
hsa-miR-151a-5p	1787
hsa-miR-143-3p	1780
hsa-miR-186-5p	1743
hsa-miR-142-5p	1621
hsa-miR-126-3p	1322
hsa-let-7g-5p	1085
hsa-miR-146b-5p	1065
hsa-miR-151a-3p	1062
hsa-miR-93-5p	976
hsa-miR-30d-5p	872
hsa-miR-423-3p	799
hsa-miR-27b-3p	744
hsa-miR-409-3p	729
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hsa-miR-126-5p	2488
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5	
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hsa-let-7i-5p	5985
hsa-miR-192-5p	3157
hsa-miR-26a-5p	2971
hsa-miR-26b-5p	2854
hsa-miR-423-5p	2405
hsa-miR-320a	1591
hsa-miR-21-5p	1559
hsa-miR-191-5p	1536
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hsa-let-7f-5p	1353
hsa-miR-181a-5p	1350
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hsa-miR-144-3p	1033
hsa-let-7a-5p	891
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hsa-let-7g-5p	732
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6	
ClustersPF	1893622
hsa-miR-486-5p	81789
hsa-miR-16-5p	51691
hsa-let-7b-5p	17172
hsa-miR-92a-3p	13001
hsa-miR-26a-5p	11536
hsa-miR-146a-5p	11434
hsa-miR-22-3p	11016
hsa-let-7f-5p	7018
hsa-let-7a-5p	6478
hsa-miR-451a	5554
hsa-let-7i-5p	4288
hsa-miR-126-5p	3746
hsa-miR-25-3p	3510
hsa-miR-191-5p	3305
hsa-miR-143-3p	2695
hsa-miR-192-5p	2642
hsa-miR-21-5p	2497
hsa-miR-320a	2140
hsa-miR-181a-5p	1863
hsa-miR-151a-5p	1826
hsa-miR-26b-5p	1525
hsa-miR-122-5p	1522
hsa-miR-423-5p	1481
hsa-miR-24-3p	1344
hsa-miR-142-5p	1285
hsa-miR-126-3p	1130
hsa-miR-423-3p	1098
hsa-let-7g-5p	961
hsa-miR-148a-3p	836
hsa-miR-146b-5p	803
hsa-miR-27b-3p	729
hsa-miR-151a-3p	687
hsa-let-7d-5p	647
hsa-miR-28-3p	557
hsa-miR-150-5p	429
hsa-miR-98-5p	402
hsa-let-7e-5p	382
hsa-miR-27a-3p	322
hsa-miR-144-3p	322
hsa-miR-30b-5p	284
hsa-miR-486-3p	284
hsa-miR-30d-5p	280
hsa-miR-140-3p	273
hsa-miR-93-5p	260
hsa-miR-103a-3p	258
hsa-miR-409-3p	256
hsa-miR-375	253
hsa-miR-101-3p	243
hsa-miR-182-5p	231
hsa-miR-186-5p	226
hsa-miR-221-3p	222
hsa-miR-484	194
hsa-miR-128-3p	178
hsa-miR-4732-3p	169
hsa-let-7c-5p	159
hsa-miR-378a-3p	154
hsa-miR-21-3p	146
hsa-miR-28-5p	146
hsa-miR-340-5p	145
hsa-let-7d-3p	138
hsa-miR-20a-5p	131
hsa-miR-17-5p	129
hsa-miR-106b-5p	124
hsa-miR-181c-5p	119
hsa-miR-342-3p	106
hsa-miR-148b-3p	104
hsa-miR-199b-3p	93
hsa-miR-30e-3p	92
hsa-miR-744-5p	90
hsa-miR-223-3p	87
hsa-miR-144-5p	87
hsa-miR-185-5p	80
hsa-miR-411-5p	80
hsa-miR-155-5p	79
hsa-miR-652-3p	78
hsa-miR-4732-5p	75
hsa-miR-199a-3p	72
hsa-miR-197-3p	69
hsa-miR-152-3p	65
hsa-miR-30a-5p	60
hsa-miR-301a-3p	55
hsa-miR-432-5p	55
hsa-miR-30e-5p	54
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hsa-miR-410-3p	54
hsa-miR-148a-5p	53
hsa-miR-125a-5p	52
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hsa-miR-132-3p	42
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hsa-miR-421	34
hsa-miR-4446-3p	34
hsa-miR-485-3p	31
hsa-miR-1255b-5p	31
hsa-miR-941	30
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hsa-miR-181b-5p	27
hsa-miR-15b-5p	27
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hsa-miR-99b-5p	20
hsa-miR-766-3p	20
hsa-miR-20b-5p	19
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hsa-miR-654-5p	18
hsa-miR-493-5p	17
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hsa-miR-877-5p	4
hsa-miR-1287-5p	4
hsa-miR-548k	4
hsa-miR-5193	4
hsa-miR-1273h-3p	4
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hsa-miR-320b	3
hsa-miR-885-5p	3
hsa-miR-1256	3
hsa-miR-2110	3
hsa-miR-2115-3p	3
hsa-miR-3909	3
hsa-miR-3928-3p	3
hsa-miR-4665-5p	3

7	
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hsa-miR-16-5p	119875
hsa-let-7b-5p	25698
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hsa-miR-451a	15344
hsa-miR-22-3p	11267
hsa-miR-25-3p	6623
hsa-let-7i-5p	5317
hsa-miR-26a-5p	4923
hsa-miR-146a-5p	3847
hsa-miR-26b-5p	3113
hsa-miR-192-5p	2959
hsa-let-7f-5p	2654
hsa-miR-423-5p	2325
hsa-miR-21-5p	2276
hsa-miR-126-5p	1802
hsa-let-7a-5p	1750
hsa-miR-181a-5p	1731
hsa-miR-191-5p	1728
hsa-miR-143-3p	1703
hsa-miR-320a	1428
hsa-miR-142-5p	1263
hsa-miR-144-3p	927
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hsa-miR-24-3p	747
hsa-miR-148a-3p	679
hsa-let-7g-5p	656
hsa-miR-122-5p	644
hsa-miR-126-3p	575
hsa-miR-30d-5p	548
hsa-miR-423-3p	512
hsa-miR-93-5p	511
hsa-miR-186-5p	509
hsa-miR-4732-3p	443
hsa-miR-486-3p	437
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hsa-let-7c-5p	170
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hsa-miR-106a-5p	26
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hsa-miR-125a-5p	12
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hsa-miR-181a-2-3p	10
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hsa-miR-200a-3p	9
hsa-miR-374a-5p	9
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hsa-miR-493-3p	9
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hsa-miR-130b-3p	6
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hsa-miR-454-5p	6
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hsa-miR-374b-5p	5
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hsa-miR-296-5p	4
hsa-miR-342-5p	4
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hsa-miR-6721-5p	3
hsa-miR-6805-5p	3
hsa-miR-18a-5p	2
hsa-miR-22-5p	2
hsa-miR-103a-2-5p	2
hsa-miR-196a-5p	2
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hsa-miR-195-3p	2
hsa-miR-378a-5p	2
hsa-miR-449a	2
hsa-miR-485-5p	2

8	
ClustersPF	2001247
hsa-miR-486-5p	84523
hsa-miR-16-5p	59549
hsa-let-7b-5p	16864
hsa-miR-22-3p	13734
hsa-miR-146a-5p	10465
hsa-miR-26a-5p	8821
hsa-let-7a-5p	8358
hsa-let-7f-5p	7077
hsa-miR-92a-3p	7034
hsa-miR-451a	4960
hsa-miR-25-3p	4888
hsa-miR-320a	4209
hsa-miR-21-5p	4080
hsa-let-7i-5p	3792
hsa-miR-191-5p	3644
hsa-miR-122-5p	3358
hsa-miR-181a-5p	2955
hsa-miR-126-5p	2915
hsa-miR-423-5p	2659
hsa-miR-192-5p	2514
hsa-miR-24-3p	2455
hsa-miR-142-5p	2048
hsa-miR-148a-3p	2026
hsa-miR-143-3p	1805
hsa-miR-26b-5p	1734
hsa-miR-126-3p	1191
hsa-miR-151a-5p	1172
hsa-miR-186-5p	1004
hsa-miR-146b-5p	878
hsa-let-7g-5p	658
hsa-miR-28-3p	639
hsa-miR-30d-5p	613
hsa-miR-150-5p	605
hsa-miR-27b-3p	567
hsa-miR-93-5p	536
hsa-let-7d-5p	533
hsa-miR-423-3p	453
hsa-miR-151a-3p	431
hsa-miR-223-3p	389
hsa-miR-140-3p	325
hsa-miR-409-3p	309
hsa-miR-98-5p	292
hsa-miR-27a-3p	283
hsa-miR-378a-3p	281
hsa-miR-221-3p	270
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hsa-miR-181c-5p	230
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hsa-miR-128-3p	221
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hsa-miR-340-5p	195
hsa-miR-20a-5p	194
hsa-miR-17-5p	193
hsa-miR-342-3p	182
hsa-miR-4732-3p	176
hsa-miR-197-3p	171
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hsa-let-7d-3p	168
hsa-miR-103a-3p	165
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hsa-miR-30e-5p	127
hsa-miR-106b-5p	118
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hsa-miR-185-5p	111
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hsa-miR-625-3p	8
hsa-miR-629-3p	8
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hsa-miR-205-5p	7
hsa-miR-374a-5p	7
hsa-miR-196b-5p	7
hsa-miR-548k	7
hsa-miR-1256	7
hsa-miR-19b-3p	6
hsa-miR-106a-5p	6
hsa-miR-142-3p	6
hsa-miR-376c-3p	6
hsa-miR-496	6
hsa-miR-654-3p	6
hsa-miR-656-3p	6
hsa-miR-4746-5p	6
hsa-miR-1273h-3p	6
hsa-let-7a-3p	5
hsa-miR-30a-3p	5
hsa-miR-183-5p	5
hsa-miR-195-5p	5
hsa-miR-130b-3p	5
hsa-miR-330-3p	5
hsa-miR-483-5p	5
hsa-miR-590-3p	5
hsa-miR-664a-3p	5
hsa-miR-2115-3p	5
hsa-miR-3615	5
hsa-miR-4707-3p	5
hsa-miR-210-3p	4
hsa-miR-193a-5p	4
hsa-miR-99b-5p	4
hsa-miR-373-3p	4
hsa-miR-339-5p	4
hsa-miR-339-3p	4
hsa-miR-335-5p	4
hsa-miR-584-5p	4
hsa-miR-889-3p	4
hsa-miR-190b	4
hsa-miR-548e-5p	4
hsa-miR-2110	4

9	
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hsa-miR-16-5p	87679
hsa-let-7b-5p	15631
hsa-miR-92a-3p	11700
hsa-miR-22-3p	9078
hsa-miR-451a	7300
hsa-miR-25-3p	5593
hsa-miR-146a-5p	4869
hsa-miR-26a-5p	4684
hsa-let-7f-5p	2846
hsa-let-7a-5p	2823
hsa-miR-320a	2258
hsa-miR-192-5p	2140
hsa-let-7i-5p	2121
hsa-miR-21-5p	1835
hsa-miR-423-5p	1827
hsa-miR-191-5p	1679
hsa-miR-126-5p	1524
hsa-miR-181a-5p	1289
hsa-miR-26b-5p	1059
hsa-miR-143-3p	1008
hsa-miR-148a-3p	776
hsa-miR-24-3p	738
hsa-miR-151a-5p	725
hsa-miR-122-5p	527
hsa-miR-186-5p	527
hsa-miR-93-5p	486
hsa-miR-126-3p	483
hsa-miR-142-5p	477
hsa-miR-151a-3p	431
hsa-miR-486-3p	390
hsa-miR-423-3p	372
hsa-let-7g-5p	371
hsa-miR-30d-5p	356
hsa-miR-4732-3p	329
hsa-miR-27b-3p	278
hsa-miR-28-3p	255
hsa-miR-140-3p	242
hsa-miR-146b-5p	237
hsa-miR-409-3p	221
hsa-miR-20a-5p	207
hsa-miR-17-5p	198
hsa-let-7d-5p	195
hsa-miR-4732-5p	184
hsa-miR-182-5p	158
hsa-miR-101-3p	153
hsa-miR-144-3p	148
hsa-miR-1-3p	135
hsa-miR-185-5p	128
hsa-miR-484	124
hsa-miR-432-5p	123
hsa-miR-221-3p	118
hsa-miR-27a-3p	114
hsa-miR-106b-5p	109
hsa-miR-378a-3p	108
hsa-let-7c-5p	105
hsa-miR-103a-3p	98
hsa-let-7e-5p	91
hsa-miR-144-5p	82
hsa-miR-98-5p	79
hsa-miR-629-5p	78
hsa-miR-148b-3p	72
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hsa-miR-150-5p	69
hsa-miR-660-5p	69
hsa-miR-30b-5p	67
hsa-miR-375	59
hsa-miR-181c-5p	57
hsa-miR-128-3p	57
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hsa-miR-576-3p	2
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hsa-miR-659-5p	2
hsa-miR-190b	2

10	
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hsa-miR-16-5p	78322
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hsa-miR-451a	7236
hsa-miR-22-3p	7173
hsa-miR-25-3p	3986
hsa-miR-26a-5p	2942
hsa-miR-26b-5p	2473
hsa-miR-146a-5p	2451
hsa-miR-192-5p	2246
hsa-let-7i-5p	2213
hsa-miR-320a	1966
hsa-miR-191-5p	1531
hsa-miR-126-5p	1504
hsa-miR-21-5p	1494
hsa-miR-423-5p	1400
hsa-miR-181a-5p	1262
hsa-miR-122-5p	1102
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hsa-let-7a-5p	819
hsa-miR-148a-3p	665
hsa-miR-143-3p	650
hsa-miR-24-3p	648
hsa-miR-142-5p	582
hsa-miR-186-5p	525
hsa-miR-151a-5p	480
hsa-let-7g-5p	450
hsa-miR-423-3p	399
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hsa-miR-486-3p	238
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hsa-miR-484	163
hsa-miR-106b-5p	107
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hsa-miR-103a-3p	75
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hsa-miR-324-3p	6
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hsa-miR-6837-3p	6
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hsa-miR-3120-3p	2
hsa-miR-3200-5p	2
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hsa-miR-628-3p	1
hsa-miR-636	1
hsa-miR-885-5p	1
hsa-miR-939-5p	1
hsa-miR-3127-5p	1
hsa-miR-3617-5p	1
hsa-miR-4646-5p	1

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hsa-miR-16-5p	58825
hsa-miR-22-3p	10229
hsa-let-7b-5p	8808
hsa-miR-92a-3p	8163
hsa-miR-146a-5p	5988
hsa-miR-122-5p	5936
hsa-miR-26a-5p	5526
hsa-let-7a-5p	4565
hsa-let-7f-5p	4045
hsa-miR-451a	3507
hsa-miR-25-3p	3008
hsa-miR-320a	2293
hsa-miR-192-5p	2214
hsa-miR-126-5p	1748
hsa-let-7i-5p	1718
hsa-miR-21-5p	1643
hsa-miR-186-5p	1367
hsa-miR-423-5p	1330
hsa-miR-191-5p	1252
hsa-miR-181a-5p	1000
hsa-miR-148a-3p	908
hsa-miR-126-3p	856
hsa-miR-24-3p	836
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hsa-miR-143-3p	777
hsa-miR-151a-5p	606
hsa-miR-142-5p	438
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hsa-let-7d-5p	330
hsa-let-7g-5p	323
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hsa-miR-486-3p	147
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hsa-miR-185-5p	103
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hsa-miR-150-5p	97
hsa-miR-378a-3p	96
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その他　資料1　血清miRNAプロファイル

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その他　資料1　血清miRNAプロファイル

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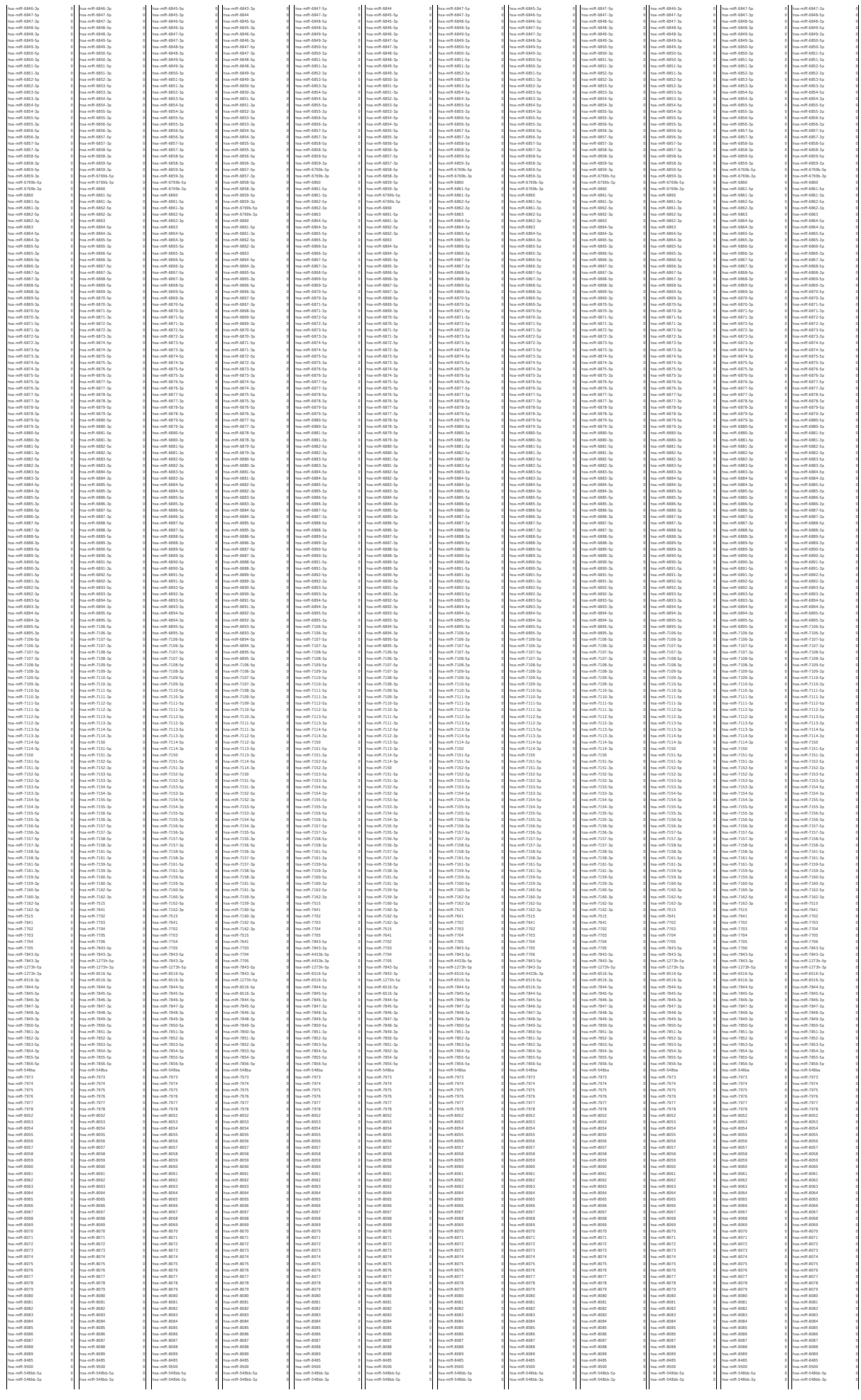
その他　資料1　血清miRNAプロファイル

その他　資料1　血清miRNAプロファイル



研究成果の刊行に関する一覧表

書籍

著者氏名	論文タイトル名	書籍全体の編集者名	書　籍　名	出版社名	出版地	出版年	ページ
該当なし							

雑誌

発表者氏名	論文タイトル名	発表誌名	巻号	ページ	出版年
該当なし					

2024年　4月　26日

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~~（国立保健医療科学院長）~~

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所属研究機関長　職　名　　学長

氏　名　小川　久雄

次の職員の（令和）５年度厚生労働科学研究費の調査研究における、倫理審査状況及び利益相反等の管理については以下のとおりです。

１．研究事業名　循環器疾患・糖尿病等生活習慣病対策総合研究事業

２．研究課題名　バイオマーカーを用いた加熱式たばこによる受動喫煙の健康影響を評価するための研究

３．研究者名　（所属部署・職名）　大学院生命科学研究部・教授

（氏名・フリガナ）　大森　久光・オオモリ　ヒサミツ

４．倫理審査の状況

	該当性の有無		左記で該当がある場合のみ記入（※1）		
	有	無	審査済み	審査した機関	未審査（※2）
人を対象とする生命科学・医学系研究に関する倫理指針（※3）	■	□	■	熊本大学	□
遺伝子治療等臨床研究に関する指針	□	■	□		□
厚生労働省の所管する実施機関における動物実験等の実施に関する基本指針	□	■	□		□
その他、該当する倫理指針があれば記入すること（指針の名称：　　）	□	■	□		□

（※1）当該研究者が当該研究を実施するに当たり遵守すべき倫理指針に関する倫理委員会の審査が済んでいる場合は、「審査済み」にチェックし一部若しくは全部の審査が完了していない場合は、「未審査」にチェックすること。

その他（特記事項）

（※2）未審査に場合は、その理由を記載すること。
（※3）廃止前の「疫学研究に関する倫理指針」、「臨床研究に関する倫理指針」、「ヒトゲノム・遺伝子解析研究に関する倫理指針」、「人を対象とする医学系研究に関する倫理指針」に準拠する場合は、当該項目に記入すること。

５．厚生労働分野の研究活動における不正行為への対応について

研究倫理教育の受講状況	受講 ■　　未受講 □

６．利益相反の管理

当研究機関におけるＣＯＩの管理に関する規定の策定	有 ■　無 □（無の場合はその理由：　　）
当研究機関におけるＣＯＩ委員会設置の有無	有 ■　無 □（無の場合は委託先機関：　　）
当研究に係るＣＯＩについての報告・審査の有無	有 ■　無 □（無の場合はその理由：　　）
当研究に係るＣＯＩについての指導・管理の有無	有 □　無 ■（有の場合はその内容：　　）

（留意事項）　・該当する□にチェックを入れること。
・分担研究者の所属する機関の長も作成すること。

令和６年３月１日

厚生労働大臣 殿

機関名　国立大学法人東北大学

所属研究機関長　職　名　　総長

氏　名　大野　英男

次の職員の令和５年度厚生労働科学研究費の調査研究における、倫理審査状況及び利益相反等の管理については以下のとおりです。

１．研究事業名　循環器疾患・糖尿病等生活習慣病対策総合研究事業

２．研究課題名　バイオマーカーを用いた加熱式たばこによる受動喫煙の健康影響を評価するための研究

３．研究者名　事業支援機構・教授

黒澤　一・クロサワ　ハジメ

４．倫理審査の状況

	該当性の有無	左記で該当がある場合のみ記入（※1）		
	有　無	審査済み	審査した機関	未審査（※2）
人を対象とする生命科学・医学系研究に関する倫理指針（※3）	□　■	□		□
遺伝子治療等臨床研究に関する指針	□　■	□		□
厚生労働省の所管する実施機関における動物実験等の実施に関する基本指針	□　■	□		□
その他、該当する倫理指針があれば記入すること（指針の名称：　　）	□　■	□		□

（※1）当該研究者が当該研究を実施するに当たり遵守すべき倫理指針に関する倫理委員会の審査が済んでいる場合は、「審査済み」にチェックし一部若しくは全部の審査が完了していない場合は、「未審査」にチェックすること。

その他（特記事項）

（※2）未審査に場合は、その理由を記載すること。

（※3）廃止前の「疫学研究に関する倫理指針」、「臨床研究に関する倫理指針」、「ヒトゲノム・遺伝子解析研究に関する倫理指針」、「人を対象とする医学系研究に関する倫理指針」に準拠する場合は、当該項目に記入すること。

５．厚生労働分野の研究活動における不正行為への対応について

研究倫理教育の受講状況	受講 ■　　未受講 □

６．利益相反の管理

当研究機関におけるＣＯＩの管理に関する規定の策定	有 ■　無 □（無の場合はその理由：　　）
当研究機関におけるＣＯＩ委員会設置の有無	有 ■　無 □（無の場合は委託先機関：　　）
当研究に係るＣＯＩについての報告・審査の有無	有 ■　無 □（無の場合はその理由：　　）
当研究に係るＣＯＩについての指導・管理の有無	有 ■　無 □（有の場合はその内容：研究実施の際の留意点を示した。　）

（留意事項）　・該当する□にチェックを入れること。
・分担研究者の所属する機関の長も作成すること。

令和6年　　3月　　31日

厚生労働大臣
~~（国立医薬品食品衛生研究所長）~~ 殿
~~（国立保健医療科学院長）~~

機関名　女子栄養大学

所属研究機関長　職　名　学長

氏　名　　香川　明夫

次の職員の（令和）５年度厚生労働科学研究費の調査研究における、倫理審査状況及び利益相反等の管理については以下のとおりです。

１．研究事業名　循環器疾患・糖尿病等生活習慣病対策総合研究事業

２．研究課題名　バイオマーカーを用いた加熱式たばこによる受動喫煙の健康影響を評価するための研究

３．研究者名　（所属部署・職名）　女子栄養大学大学院・教授

（氏名・フリガナ）　緒方　裕光・オガタ　ヒロミツ

４．倫理審査の状況

	該当性の有無	左記で該当がある場合のみ記入（※1）		
	有　無	審査済み	審査した機関	未審査（※2）
人を対象とする生命科学・医学系研究に関する倫理指針（※3）	■　□	■	熊本大学	□
遺伝子治療等臨床研究に関する指針	□　■	□		□
厚生労働省の所管する実施機関における動物実験等の実施に関する基本指針	□　■	□		□
その他、該当する倫理指針があれば記入すること（指針の名称：　　）	□　■	□		□

（※1）当該研究者が当該研究を実施するに当たり遵守すべき倫理指針に関する倫理委員会の審査が済んでいる場合は、「審査済み」にチェックし一部若しくは全部の審査が完了していない場合は、「未審査」にチェックすること。

その他（特記事項）

（※2）未審査に場合は、その理由を記載すること。
（※3）廃止前の「疫学研究に関する倫理指針」、「臨床研究に関する倫理指針」、「ヒトゲノム・遺伝子解析研究に関する倫理指針」、「人を対象とする医学系研究に関する倫理指針」に準拠する場合は、当該項目に記入すること。

５．厚生労働分野の研究活動における不正行為への対応について

研究倫理教育の受講状況	受講 ■　　未受講 □

６．利益相反の管理

当研究機関におけるＣＯＩの管理に関する規定の策定	有 ■　無 □（無の場合はその理由：　　）
当研究機関におけるＣＯＩ委員会設置の有無	有 ■　無 □（無の場合は委託先機関：　　）
当研究に係るＣＯＩについての報告・審査の有無	有 ■　無 □（無の場合はその理由：　　）
当研究に係るＣＯＩについての指導・管理の有無	有 □　無 ■（有の場合はその内容：　　）

（留意事項）　・該当する□にチェックを入れること。
・分担研究者の所属する機関の長も作成すること。

2024年　4月　26日

厚生労働大臣
~~（国立医薬品食品衛生研究所長）~~ 殿
~~（国立保健医療科学院長）~~

機関名　　国立大学法人熊本大学

所属研究機関長　職　名　　学長

氏　名　小川　久雄

次の職員の（令和）５年度厚生労働科学研究費の調査研究における、倫理審査状況及び利益相反等の管理については以下のとおりです。

１．研究事業名　循環器疾患・糖尿病等生活習慣病対策総合研究事業

２．研究課題名　バイオマーカーを用いた加熱式たばこによる受動喫煙の健康影響を評価するための研究

３．研究者名　（所属部署・職名）　大学院生命科学研究部・教授

（氏名・フリガナ）　大坪　和明・オオツボ　カズアキ

４．倫理審査の状況

	該当性の有無		左記で該当がある場合のみ記入（※1）		
	有	無	審査済み	審査した機関	未審査（※2）
人を対象とする生命科学・医学系研究に関する倫理指針（※3）	■	□	■	熊本大学	□
遺伝子治療等臨床研究に関する指針	□	■	□		□
厚生労働省の所管する実施機関における動物実験等の実施に関する基本指針	□	■	□		□
その他、該当する倫理指針があれば記入すること（指針の名称：　　）	□	■	□		□

（※1）当該研究者が当該研究を実施するに当たり遵守すべき倫理指針に関する倫理委員会の審査が済んでいる場合は、「審査済み」にチェックし一部若しくは全部の審査が完了していない場合は、「未審査」にチェックすること。

その他（特記事項）

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（※3）廃止前の「疫学研究に関する倫理指針」、「臨床研究に関する倫理指針」、「ヒトゲノム・遺伝子解析研究に関する倫理指針」、「人を対象とする医学系研究に関する倫理指針」に準拠する場合は、当該項目に記入すること。

５．厚生労働分野の研究活動における不正行為への対応について

研究倫理教育の受講状況	受講 ■　　未受講 □

６．利益相反の管理

当研究機関におけるＣＯＩの管理に関する規定の策定	有 ■　無 □（無の場合はその理由：　　）
当研究機関におけるＣＯＩ委員会設置の有無	有 ■　無 □（無の場合は委託先機関：　　）
当研究に係るＣＯＩについての報告・審査の有無	有 ■　無 □（無の場合はその理由：　　）
当研究に係るＣＯＩについての指導・管理の有無	有 □　無 ■（有の場合はその内容：　　）

（留意事項）　・該当する□にチェックを入れること。
・分担研究者の所属する機関の長も作成すること。

2024年　4月　26日

厚生労働大臣
~~（国立医薬品食品衛生研究所長）~~ 殿
~~（国立保健医療科学院長）~~

機関名　　国立大学法人熊本大学

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氏　名　小川　久雄

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２．研究課題名　バイオマーカーを用いた加熱式たばこによる受動喫煙の健康影響を評価するための研究

３．研究者名　（所属部署・職名）　大学院生命科学研究部・准教授

（氏名・フリガナ）　首藤　剛・シュトウ　ツヨシ

４．倫理審査の状況

	該当性の有無		左記で該当がある場合のみ記入（※1）		
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遺伝子治療等臨床研究に関する指針	□	■	□		□
厚生労働省の所管する実施機関における動物実験等の実施に関する基本指針	□	■	□		□
その他、該当する倫理指針があれば記入すること（指針の名称：　　）	□	■	□		□

（※1）当該研究者が当該研究を実施するに当たり遵守すべき倫理指針に関する倫理委員会の審査が済んでいる場合は、「審査済み」にチェックし一部若しくは全部の審査が完了していない場合は、「未審査」にチェックすること。

その他（特記事項）

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５．厚生労働分野の研究活動における不正行為への対応について

研究倫理教育の受講状況	受講 ■　　未受講 □

６．利益相反の管理

当研究機関におけるＣＯＩの管理に関する規定の策定	有 ■　無 □（無の場合はその理由：　　）
当研究機関におけるＣＯＩ委員会設置の有無	有 ■　無 □（無の場合は委託先機関：　　）
当研究に係るＣＯＩについての報告・審査の有無	有 ■　無 □（無の場合はその理由：　　）
当研究に係るＣＯＩについての指導・管理の有無	有 □　無 ■（有の場合はその内容：　　）

（留意事項）　・該当する□にチェックを入れること。
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