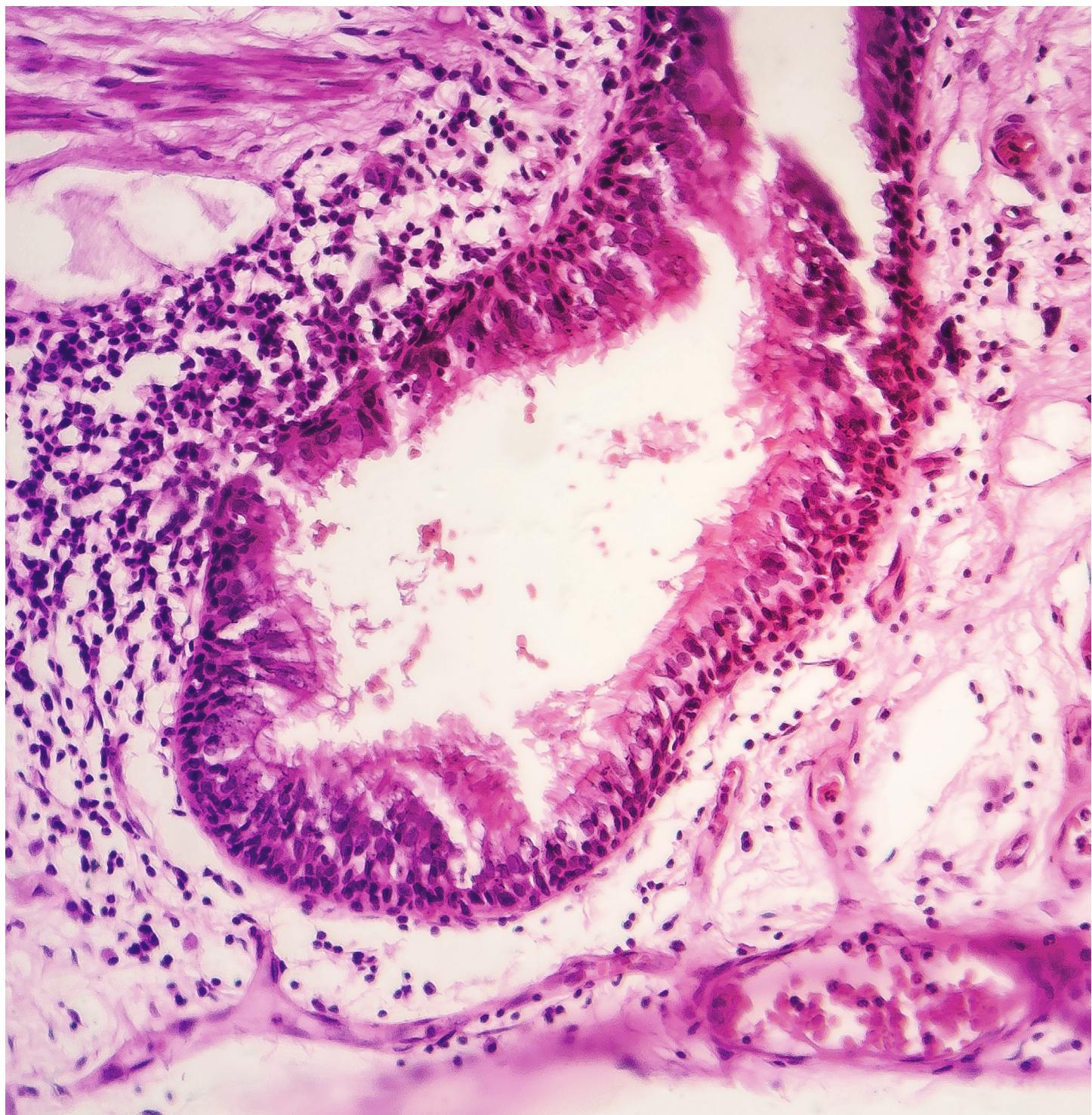


Tools for respiratory research

PromoCell®

Solutions for world-class
respiratory research

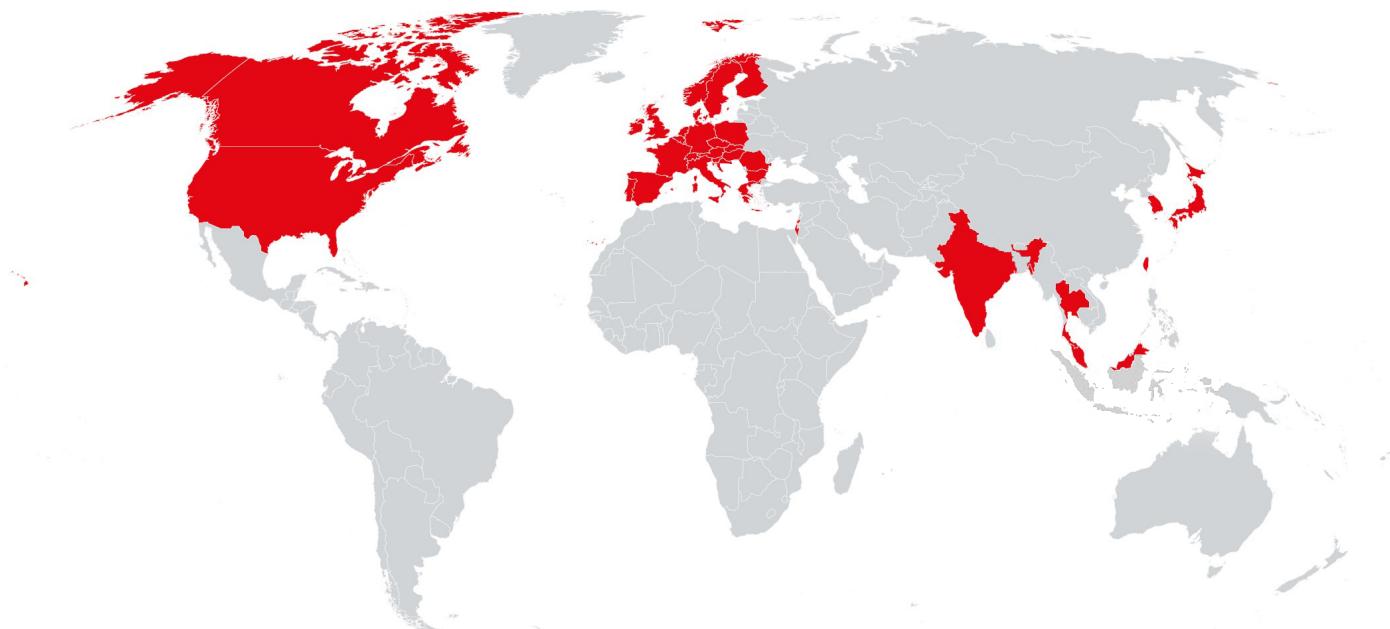


Who we are

At PromoCell, we help scientists do better research by offering a world-class portfolio of human primary cells, stem cells, blood cells, and optimized cell culture media.

With over 35 years of expertise, we are recognized as a trusted partner for scientists, providing tools and support for groundbreaking research, from discovery to clinical applications. At every stage of your research journey, we offer expert support from researchers who understand your work.

Each year, over 600 peer-reviewed publications feature PromoCell products. We operate in over 36 countries around the world, helping scientists with their research needs.



Austria	Croatia	France	India	Japan	Netherlands	Romania	South Korea	Taiwan
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Learn more about PromoCell on our [website](#), or connect with us on [LinkedIn](#), [YouTube](#) or [Facebook](#).

Our commitment to quality

Ethics and quality are at the heart of our business. We own the entire tissue collection and manufacturing process, which means we're able to provide fast and direct ethical regulatory support. All our products comply with European biomedical conventions, ensuring that human rights and donor privacy are always protected. Our ISO certifications demonstrate our commitment to quality, and our EXCiPACT™ GMP certification enables us to produce our cell culture media and reagents according to GMP standards.



- Our Quality Management System is certified according the EXCiPACT™ GMP standard



- PromoCell operates according to ISO 9001:2015 in order to consistently provide products and services that meet customer requirements as well as applicable statutory and regulatory requirements.

Our commitment to fighting respiratory diseases

Respiratory system

The respiratory tract is a complex network of organs and tissues that are vital for our survival. The respiratory network consists of the mouth, nose, sinuses, pharynx (throat), trachea, bronchial tubes, and lungs. The lungs have a highly organized hierarchical structure; they branch into lung lobes, bronchioles, and alveoli. The alveoli are in contact with capillaries, forming a complex alveolar capillary network.

Respiratory tissues are composed of various specialized cell types, including epithelial cells, fibroblasts, smooth muscle cells, and endothelial cells. These cells work

together to ensure that every organ, tissue, and cell of the body is supplied with oxygen through breathing. Respiratory organs and tissues help the body extract oxygen from the air and remove carbon dioxide and other waste gases that can become toxic to the body.

Air enters the body through the respiratory system; therefore, the airways and lungs are constantly exposed to potentially harmful bacteria, viruses, fungi, and pollutants carried through the air. These inhaled irritants can cause infections and damage the lungs, leading to various respiratory diseases.

Our primary cells from the respiratory tract

Respiratory diseases are a leading cause of death and disability worldwide. Respiratory research is key to reducing the burden of lung infections, lung cancer, and asthma, which are the most common respiratory diseases.

Primary cells are excellent models for studying respiratory diseases. Using our cell culture models, you can better understand

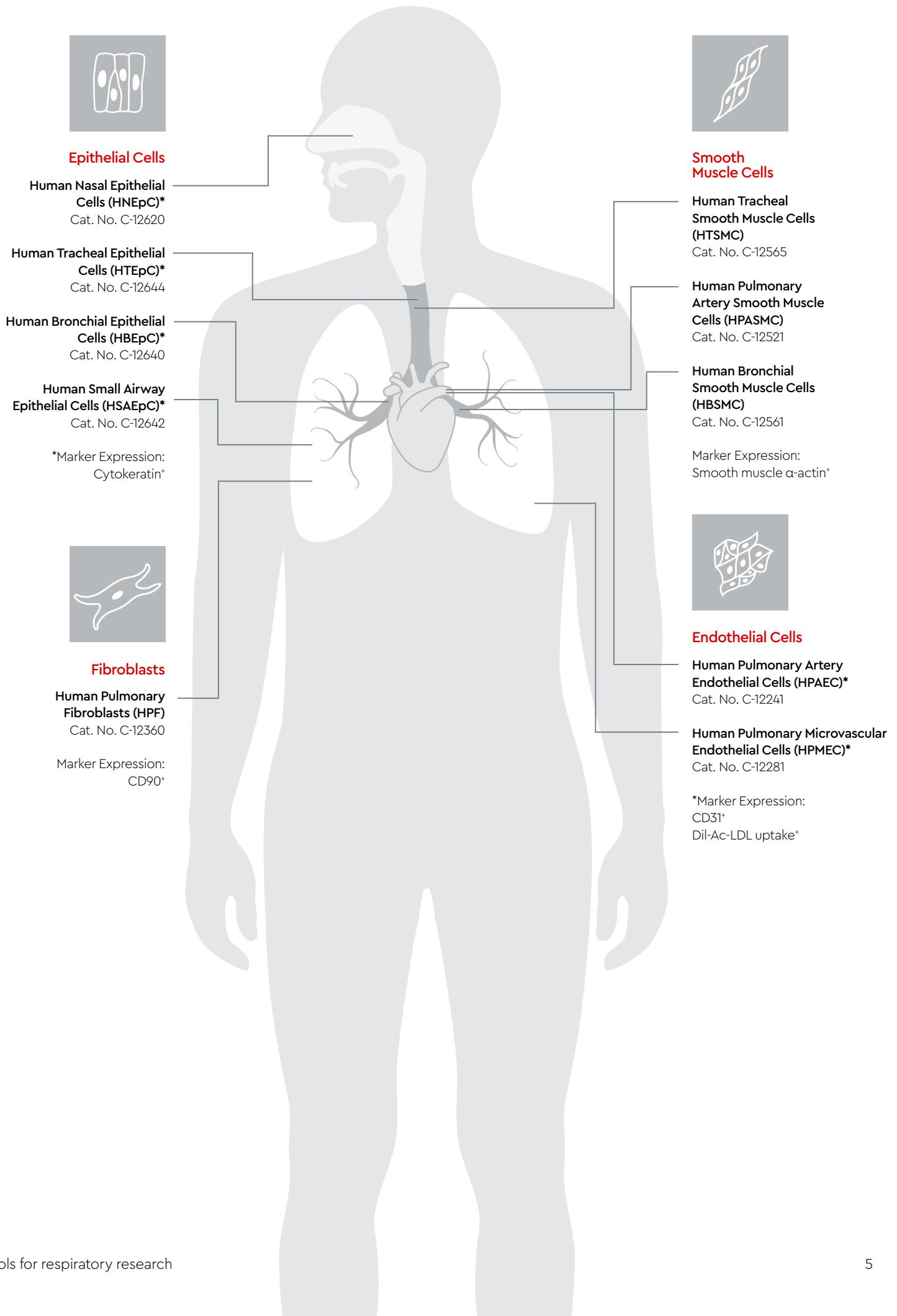
the cellular and molecular mechanisms that cause lung disease, identify novel therapeutic targets, and test the efficacy and safety of new therapies for respiratory diseases.

We offer an extensive portfolio of tools for basic and translational respiratory research, including various primary cell culture models.

Our respiratory models include:

- Nasal, tracheal, and bronchial airway epithelial cells
- Pulmonary smooth muscle cells
- Pulmonary fibroblasts
- Lung endothelial cells





In vitro airway models

In vitro airway models are becoming increasingly popular in respiratory research owing to their numerous advantages. Airways consist of a complex network of specialized cells, which are challenging to study using

in vivo models. In contrast, 3D cultures of primary cells from donors with specific disease states provide excellent, reproducible, and physiologically relevant models for studying the *in vivo* processes underlying respiratory

diseases. In addition, comparing primary cells from a donor with a specific disease to those from a healthy donor represents a straightforward and robust way to identify therapeutic targets for lung diseases.

Models of respiratory diseases

There are several respiratory diseases that vary in their etiology (i.e., smoking, air pollution, mutations, viral infections, bacterial infections), pathophysiology (e.g., inflammatory disease, effusion, neoplasia, obstructive pulmonary disease), and duration (i.e., acute or chronic).

Establishing *in vivo* models that accurately mimic these complex human diseases is extremely challenging and time-consuming. In contrast, isolating primary cells from donors with specific disease states, environmental exposures (e.g., smoker versus non-smoker), and HLA types allows for the generation of versatile, functional, and physiologically relevant human

airway models. Challenges associated with the establishment of animal models of human respiratory diseases are exacerbated in the case of emerging, fast-spreading, life-threatening infectious respiratory diseases, such as COVID-19. Despite extensive efforts, progress in the development of animal models of COVID-19 remains limited.

Animal models require laborious and time-consuming genetic manipulations to render non-human organisms susceptible to human viruses. In contrast, primary human cells can be easily infected *in vitro* to study infectious lung diseases and identify therapeutic targets.

Models of human airways: Airway organoids, air-liquid interface, and lung-on-chip systems

Human airways are complex and consist of multiple cell types that interact with each other. The spatial relationship and interplay between different cells in the airway can affect the development and progression of respiratory diseases. Airway organoids from primary human cells accurately resemble the 3D environment of human lungs. Consequently, human airway organoids are useful *in vitro* models for studying the role of the complex microenvironment and 3D architecture of airways in respiratory diseases, as well as for identifying and testing drug candidates.

In contrast to cell lines used as *in vitro* airway models, air-liquid interface (ALI) culture using primary cells more accurately resembles the *in vivo* environment of human airways and enables the characterization of changes in ALI and epithelial barriers in respiratory disease.

Lung-on-chip models are also valuable models of human airways. They provide dynamic, microfluidic platforms that can better recapitulate the mechanical forces and physiological conditions present in human lungs.

Models of respiratory infections

Challenges associated with the establishment of animal models of human respiratory diseases are exacerbated in the case of emerging, fast-spreading, life-threatening infectious respiratory diseases, such as COVID-19. Despite extensive efforts, progress in the development of animal models of COVID-19 is limited.

In contrast to animal models that require laborious and time-consuming genetic manipulations to render non-human organisms susceptible to human viruses, primary human cells can be easily infected *in vitro* to study infectious lung diseases and identify therapeutic targets.

Air-liquid interface culture

Our Air-Liquid Interface (ALI) culture system enables you to generate a reproducible and physiologically relevant 3D airway model with primary human bronchial epithelial cells (HBEpCs) and functional epithelial barriers (Fig. 1). Our cell culture medium, Air-Liquid Interface Medium (ALI-Airway), is free of serum and bovine pituitary extract (BPE) and contains a low level of growth factors,

helping you maintain a non-proliferating cell population over a period of 28 days. Cells are grown on a semipermeable porous membrane that mimics the respiratory basement membrane.

We have demonstrated successful ALI culture generation using our primary human bronchial (HBEpC), nasal (HNEpC), and tracheal (HTEpC) epithelial cells, as detailed in our

application notes. The functionality of HBEpCs can be monitored through the fast development of an epithelial barrier formed by tight junctions. In addition to the ALI-Airway medium, we provide HBEpCs that are prescreened for proper barrier function in our ALI medium. We also offer HBEpC from HLA-typed donors and donors with chronic obstructive pulmonary disease (COPD) or asthma.

Key advantages of our ALI culture system:

- Minimizes lot-to-lot variation using optimized BPE-free and serum-free formulation
- Offers stable and uniform transepithelial electrical resistance (TEER) values of $> 500 \Omega \text{ cm}^2$ for up to 4 weeks of culture
- Enables culturing human differentiated bronchial epithelial cells with cell viability $> 70\%$



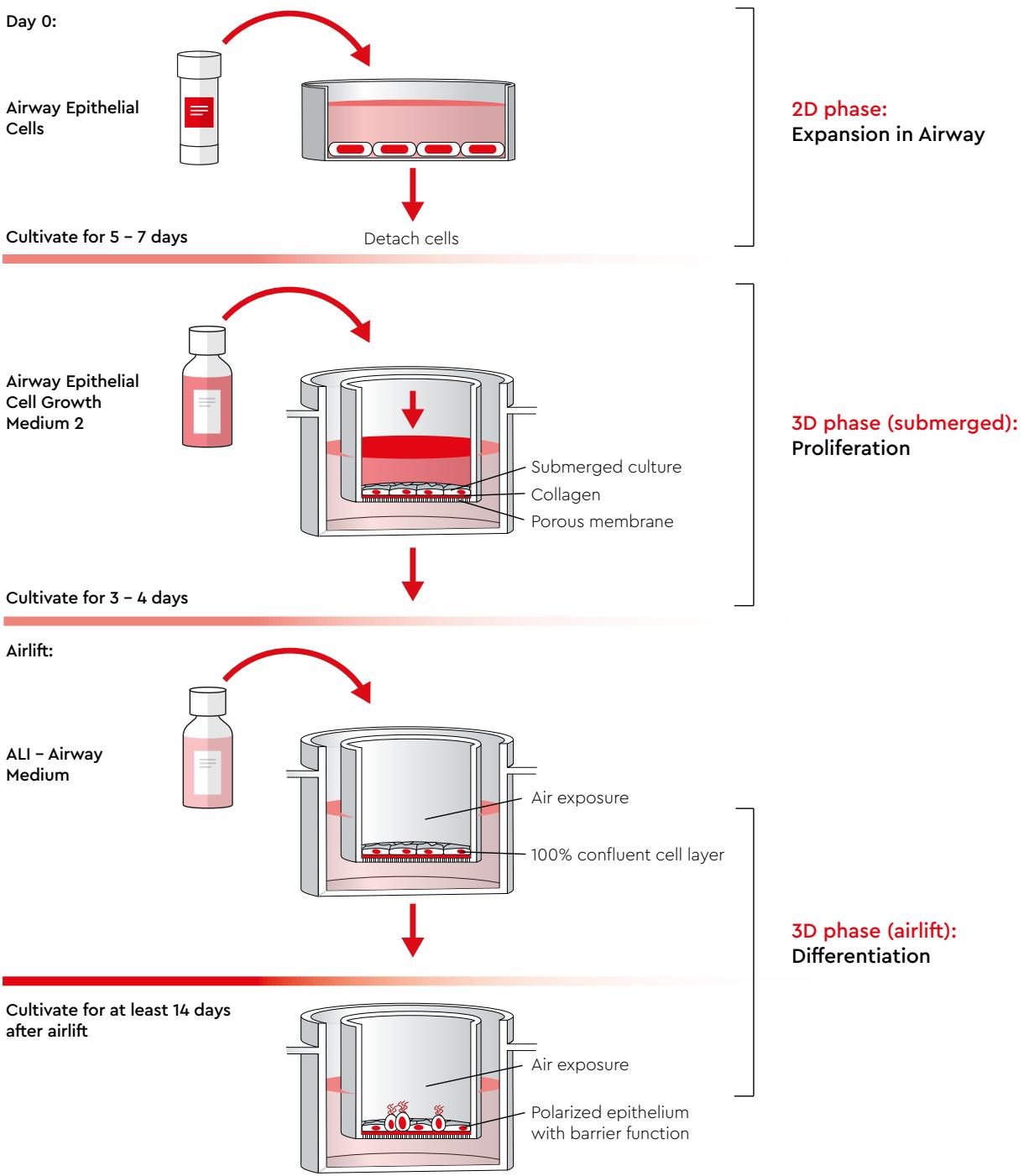


Fig. 1: Overview of our Air-Liquid Interface (ALI) culture system. Our streamlined workflow combines 2D expansion and 3D differentiation phases. This workflow allows for optimal differentiation and function of ALI-prescreened HBEpCs. For detailed protocols and methodology, please refer to our comprehensive application note: [Air-Liquid Interface Culture System for Standardized Respiratory Research](#).

Our ALI-prescreened cells maintain TEER values $>500 \Omega \text{ cm}^2$ up to 4 weeks of culture for HBEpCs (Fig. 2) in our ALI culture system. Similarly, TEER values $>500 \Omega \text{ cm}^2$ can be obtained in up to 14 days of culture for HNEpC and HTEpC when you follow the instructions described in our application note: [Air-Liquid Interface Culture System for Standardized Respiratory Research](#).

Influence of ALI-media composition of epithelial barrier integrity

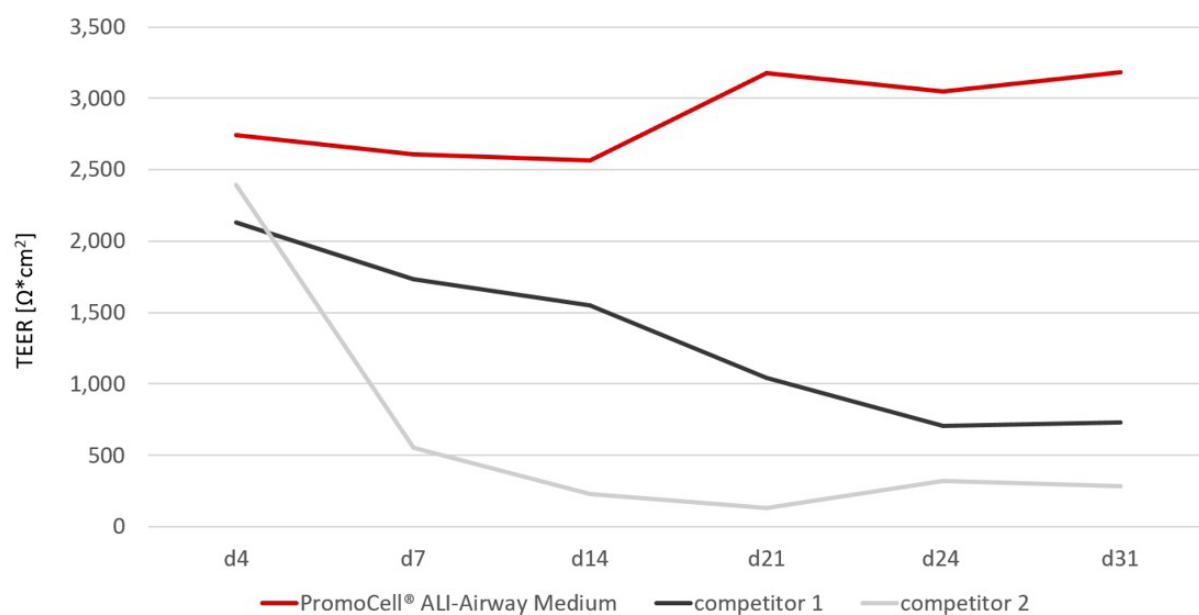


Fig. 2: TEER values of barrier-forming HBEPcs over a culture period of 28 days post-airlift. Our standardized ALI system consistently delivers superior barrier formation (higher TEER values) compared to competitor media. Our ALI-Airway medium ensures stable TEER values $>500 \Omega \text{ cm}^2$ over different time points. For detailed protocols and methodology, see our application note.

Our ALI-prescreened HBEPcs maintain TEER values $>500 \Omega \text{ cm}^2$ for at least 14 days of culture in our ALI culture system (Fig. 2) when you follow the instructions described in our application note: [Air-Liquid Interface Culture System for Standardized Respiratory Research](#).

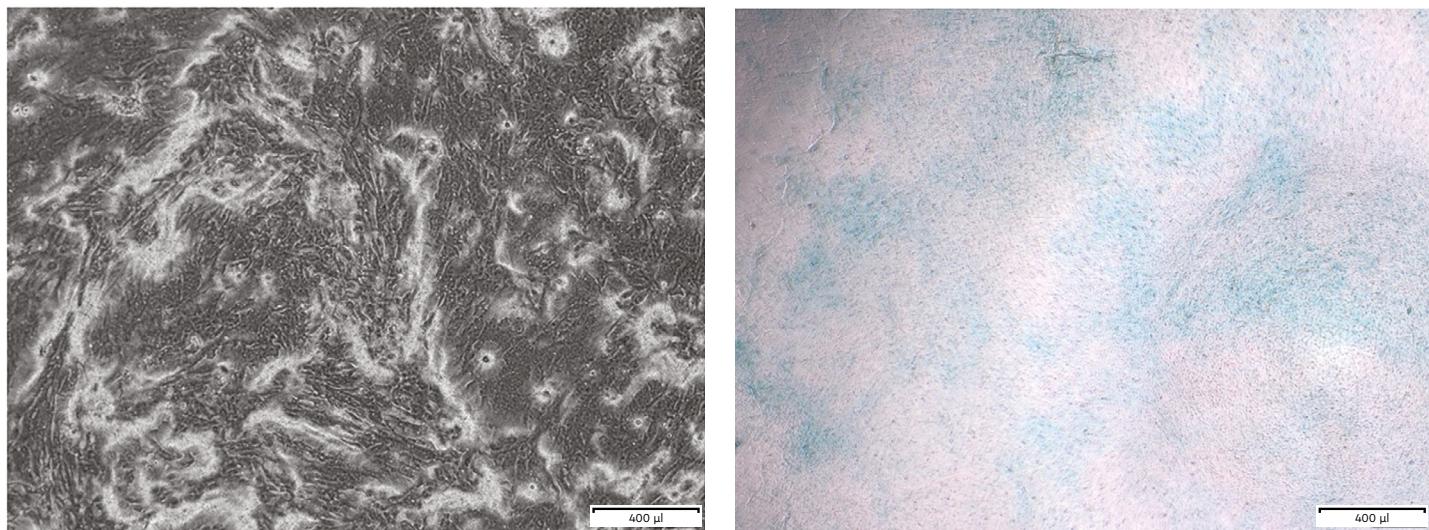


Fig. 3: Mucin production indicates differentiation of HBEPC cultured under air-liquid conditions. **Left:** Microscopy of unfixed HBEPC after culturing in ALI-Airway medium for 33 days. **Right:** Alcian Blue staining of fixed cells after 33 days in culture. Alcian Blue stains sulfated and carboxylated and acid mucopolysaccharides and sulfated and carboxylated sialomucins (glycoproteins), which are indicated in blue (4x magnification, scale bar 400 µm).

Airway organoid model

Our ALI-Airway medium can be used to generate 3D bronchospheres, which are organoid-like structures formed by HBEPs when cultured in basal membrane extract (BME). Bronchospheres can be used for

high-throughput drug screening. Fully differentiated organoids have a cell-free center lumen surrounded by a polarized epithelial cell layer (Fig. 4). Depending on their polarity along the apical-basal axis, multiciliated cells

can face the apical membrane or the lumen of the airway organoid. Exposure of HBEPs to the Rho-associated protein kinase inhibitor (ROCKi) Y-27632 increases their colony-forming efficiency by around 20%.

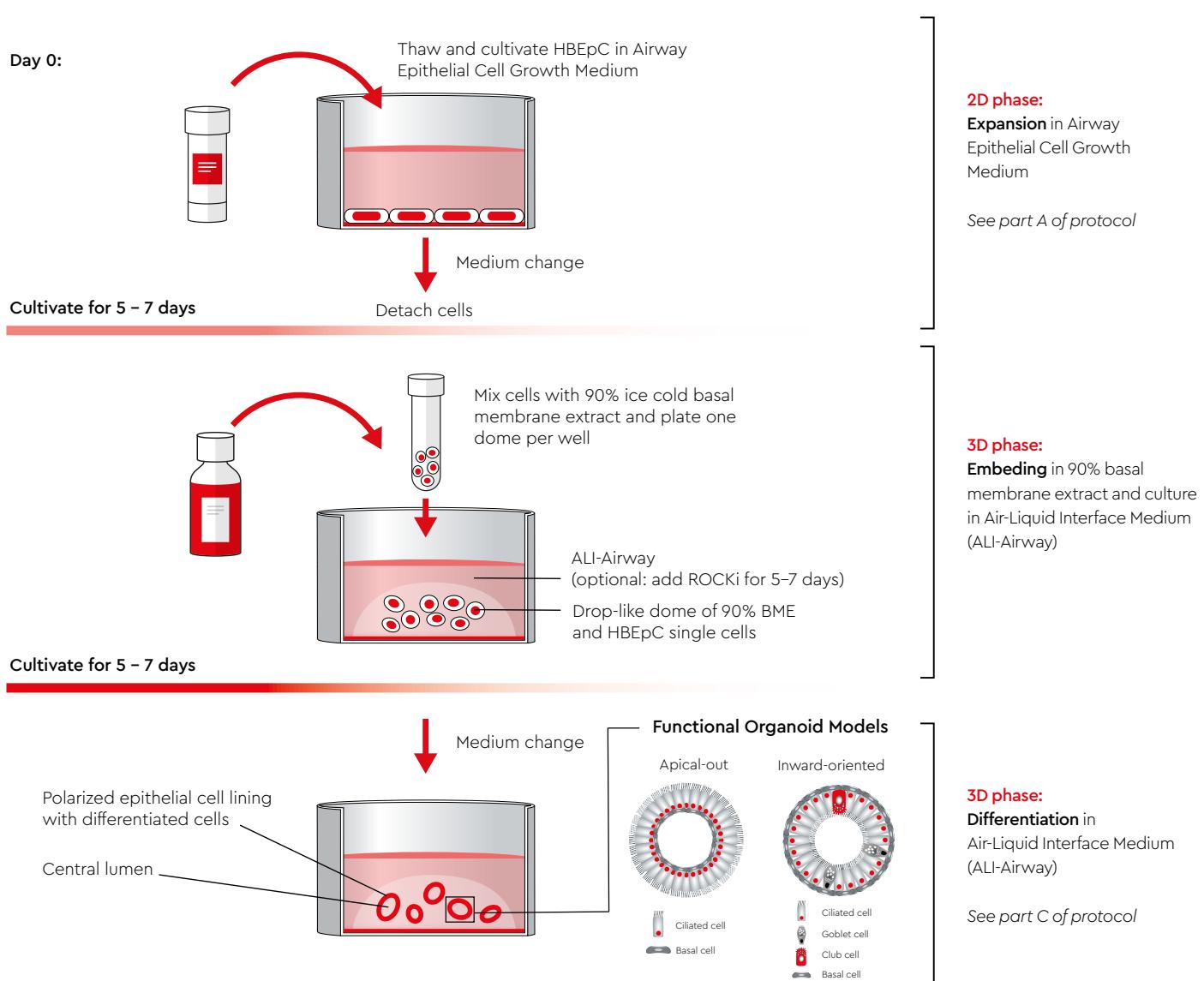


Fig. 4: Overview of our airway organoid model. Our protocol enables efficient generation of functional 3D bronchospheres with consistent differentiation. The differentiation process is completed after 4 weeks of culture in ALI-Airway medium. For detailed methodology, please see our application note [Generation of human airway organoids from primary cells](#).

Bronchospheres cultured in our ALI-Airway medium show the first signs of differentiation after 2 weeks of culture when ciliated cells become visible. Ciliated cells maintain their high

self-renewal potential for over 4 weeks. The second stage of differentiation is the formation of the airway organoid lumen, which can be visualized by microscopy (Fig. 5). Organoids

with apical-out ciliated cells tend to rotate in the matrix because of synchronous ciliary beating. Differentiation is completed after 4 weeks of 3D culture.

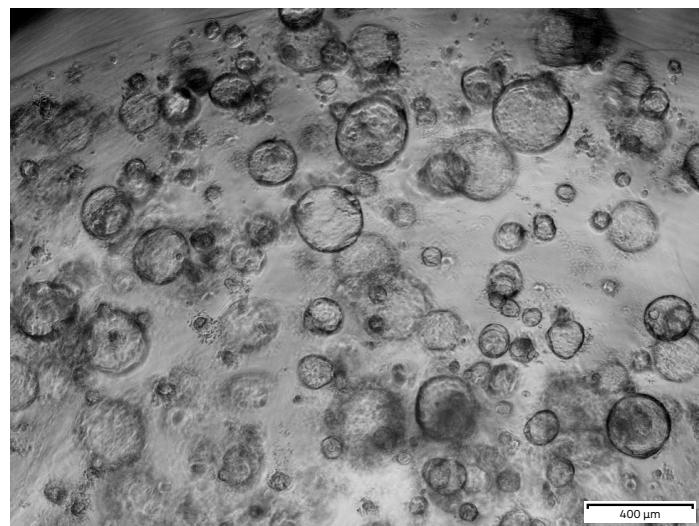
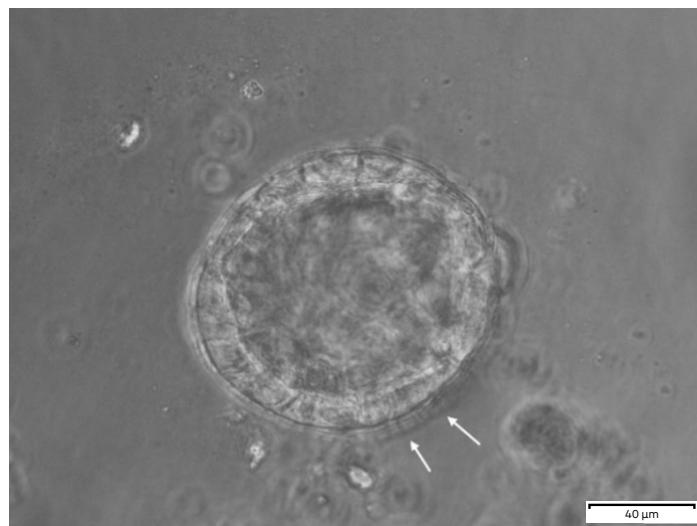


Fig. 5: Differentiated airway organoids (3D bronchospheres) demonstrate long-term viability. **Left:** Bright-field microscopy images showing differentiation of HBEPs after 16 days of 3D culture in ALI-Airway medium. **Right:** Long-term culture of airway organoids, which maintain their self-renewal potential. HBEPs were seeded in multiwell plates (10.000 cells/well). Our system supports sustained organoid development with polarization and functional characteristics. Detailed protocols are available in our application note: [Generation of human airway organoids from primary cells](#).

Models for chronic respiratory diseases

Respiratory diseases represent a significant healthcare and economic burden and are a leading cause of death. Common diseases of the respiratory system include chronic obstructive pulmonary disease (COPD),

asthma, lung cancer, multidrug-resistant bacterial infections (e.g., tuberculosis), and infectious diseases (e.g., influenza and SARS).

Our wide range of primary cells from patient donors can help you identify new

therapeutic targets and evaluate the efficacy and safety of potential drug candidates for the treatment of acute and chronic respiratory diseases.

What we offer

Donors with COPD Type I-IV

- Bronchial Epithelial Cells (HBEPs)
- Small Airway Epithelial Cells (HSAEPs)
- Bronchial Smooth Muscle Cells (HBSMC)
- Pulmonary Fibroblasts (HPF)

Donors with asthma

- Bronchial Epithelial Cells (HBEPs)
- Small Airway Epithelial Cells (HSAEPs)
- Bronchial Smooth Muscle Cells (HBSMC)
- Pulmonary Fibroblasts (HPF)

Donors tested negative for COVID-19

- Nasal Epithelial Cells (HNEPs)
- Pulmonary Microvascular Endothelial Cells (HPMEC)

Our primary cells allow for the establishment of respiratory disease models with different smoker statuses (i.e., never-smoker, non-smoker, and smoker) and HLA types. Our streamlined tissue procurement with established partners enables us to source tissues on a regular basis and provide you with a wide variety of cell models and lot sizes that align with your experimental needs.

Advantages of our primary cells from patient donors

- Verified cell identity through cell marker expression and morphology
- Verified cell growth performance and cell morphology for several passages for each lot
- Wide variety of model characteristics and lot sizes

All cell types from patient donors can be cultured in our cell culture media to ensure optimal growth performance for your experiments.

Choose a suitable culture medium for respiratory research

The cell culture formulation you need among available options depends on your specific research objectives, and selecting the right media formulation is key for experimental success. Different formulations offer distinct

advantages for various applications, from basic cell expansion to specialized differentiation protocols. Beyond supporting optimal cell growth, media formulation plays a critical role in experimental planning and

risk assessment processes. Our diverse range of cell culture media formulations ensures consistent excellence for cell expansion and maintenance across various research needs.

Determine which media formulation is most appropriate for your research application.

Media	Serum-free	BPE-free	Xeno-free	Phenol red-free
 Airway Epithelial Cell Growth Medium	✓			
 Airway Epithelial Cell Basal Medium (prf)	✓			✓
 Airway Epithelial Cell Growth Medium 2	✓	✓		
 Airway Epithelial Cell Growth Medium XF (prf)	✓	✓	✓	✓
 Small Airway Epithelial Cell Growth Medium	✓			

Media	Serum-free	BPE-free	Xeno-free	Phenol red-free
 Small Airway Epithelial Cell Basal Medium (prf)	✓			✓
 Air-Liquid Interface Medium (ALI-Airway)	✓	✓		



Our solutions for respiratory research

Human primary cells

Cell type	Description	Cat. n°.	Marker	Recommended culture media*
Epithelial cells	Human Nasal Epithelial Cells (HNEpC)	C-12620	Cytokeratin ⁺	C-21040 C-21050 C-21060
	Human Tracheal Epithelial Cells (HTEpC)	C-12644	Cytokeratin ⁺	C-21040 C-21050 C-21060
	Human Bronchial Epithelial Cells (HBEpC)	C-12640	Cytokeratin ⁺	C-21040 C-21050 C-21060
	Human Small Airway Epithelial Cells (HSAEpC)	C-12642	Cytokeratin ⁺	C-21070
Fibroblasts	Human Pulmonary Fibroblasts (HPF)	C-12360	CD90 ⁺	C-23020
Endothelial cells	Human Pulmonary Artery Endothelial Cells (HPAEC)	C-12241	CD31 ⁺ Dil-Ac-LDL uptake ⁺	C-22010 C-22011
	Human Pulmonary Microvascular Endothelial Cells (HPMEC)	C-12281	CD31 ⁺ Dil-Ac-LDL uptake ⁺	C-22020 C-22022
Smooth muscle cells	Human Pulmonary Artery Smooth Muscle Cells (HPASMC)	C-12521	Smooth muscle α -actin ⁺ CD31 ⁻	C-22062
	Human Tracheal Smooth Muscle Cells (HTSMC)	C-12565	Smooth muscle α -actin ⁺ CD31 ⁻	C-22062
	Human Bronchial Smooth Muscle Cells (HBSMC)	C-12561	Smooth muscle α -actin ⁺ CD31 ⁻	C-22062

*The catalog numbers in this table are for media in ready-to-use packaging.

Cell culture media for expansion and differentiation

Cell type	Product	Size	Catalog Number
Epithelial cells	Airway Epithelial Cell Growth Medium	500 ml	C-21060 / Kit C-21160
	Airway Epithelial Cell Growth Medium 2	500 ml	C-21040
	Airway Epithelial Cell Growth Medium XF (prf)	500 ml	C-21050
	Small Airway Epithelial Cell Growth Medium	500 ml	C-21070 / Kit C-21170
	Air-Liquid Interface Medium (ALI-Airway)	500 ml	C-21080
Fibroblasts	Fibroblast Growth Medium 2	500 ml	C-23020 / Kit C-23120
Endothelial cells	Endothelial Cell Growth Medium	500 ml	C-22010 / Kit C-22110
	Endothelial Cell Growth Medium 2	500 ml	C-22011 / Kit C-22111
	Endothelial Cell Growth Medium MV	500 ml	C-22020 / Kit C-22120
	Endothelial Cell Growth Medium MV 2	500 ml	C-22022 / Kit C-22121
Smooth muscle cells	Smooth Muscle Cell Growth Medium 2	500 ml	C-22062 / Kit C-22162

*The catalog numbers listed in this table refer to media in ready-to-use packaging, unless stated otherwise.

Reagents for dissociation and cryopreservation

Product	Size	Catalog number
DetachKit	30 ml	C-41200
	125 ml	C-41210
	250 ml	C-41220
Cryo-SFM Plus	30 ml	C-29920
	125 ml	C-29922
Cryo-SFM Plus (prf)	30 ml	C-29930
	125 ml	C-29932

Get expert guidance

Choosing the right cell types and culture media can make the difference between successful experiments and frustrating setbacks. Our respiratory research specialists understand the unique challenges of your work and are ready to help you select the optimal combination of cells and media for your specific research goals.

Contact our experts today to discuss your respiratory research needs: scientific.support@promocell.com

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