Correlation of Nasal Morphology to Air-Conditioning and Clearance 1 Function. 2 3 David E. White^a, Ahmed M. Al-Jumaily^a, James Bartley^b, Jun Lu^a 4 5 ^a Institute of Biomedical Technologies, Auckland University of Technology, Auckland, New 6 Zealand. 7 ^b Department of Surgery, University of Auckland, Auckland, New Zealand. 8 9 10 11 **Corresponding Author:** Professor Ahmed M. Al-Jumaily 12 Institute of Biomedical Technologies, 13 Auckland University of Technology, 14 Private Bag 92006, 15 Auckland, 1142, 16 New Zealand. 17 E-mail: ahmed.aljumaily@aut.ac.nz 18

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- 3 Nasal morphology plays an important functional role in the maintenance of upper airway
- 4 health. Identification of functional regions, based on morphological attributes, assists in
- 5 correlating location to primary purpose. The effects of morphological variation on heat and
- 6 water mass transport in congested and patent nasal airways were investigated by examining
- 7 nasal cross-sectional MRI images from 8 healthy subjects. This research confirms the
- 8 previous identification of functional air-conditioning regions within the nose. The first is the
- 9 anterior region where the morphology prevents over-stressing of tissue heat and fluid supply
- 10 near the nares. The second is the mid region where low flow velocity favours olfaction and
- particle deposition. The third is the posterior region which demonstrates an increase in heat
- and water mass flux coefficients to compensate for rising air humidity and temperature.
- 13 Factors identified within the congested airway that favour enhanced mucocillary clearance
- were also identified.

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KEYWORDS

17 nasal morphology, convection coefficient, hydraulic diameter

1. INTRODUCTION

- 3 The mucociliary blanket in the nasal cavity provides the human airways with the first line of
- 4 defence against infection, particles and airborne pollutants through entrapment in sticky
- 5 mucus (Bossi et al., 2004; Cone, 2009; Widdicombe, 2002). Within the nose, this defensive
- 6 layer is continuously transported towards the pharynx, where it is cleared by swallowing
- 7 (Antunes et al., 2009) or expectoration. Mucocillary transport velocity (MTV) is
- 8 significantly influenced by airway fluid hydration state (Button and Boucher, 2008) and
- 9 normally ranges from 3-25 mm/min (Boek et al., 2002; Mygind and Dahl, 1998).

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- 11 The interior nasal mucosa also provides heat and moisture to condition inhaled air and to
- prevent lower airway dryness (Doorly et al., 2008a; Rouadi et al., 1999). This air-
- conditioning function is achieved through the combined action of anatomical features, which
- include complex and narrow internal geometry (Croce et al., 2006; Doorly et al., 2008b;
- 15 Segal et al., 2008; Wolf et al., 2004), and vascular/cellular regulatory systems (White et al.,
- 16 2011).

- 18 *1.1. Nasal morphology*
- 19 Modern imaging techniques, such as computed axial tomography (CT) and magnetic
- 20 resonance imaging (MRI), have enabled the capture of accurate *in-vivo* nasal geometric
- 21 information, which goes beyond that previously obtained by acoustic rhinometry (AR).
- 22 Earlier investigations comparing CT and AR cross-sectional data has a good correlation
- within the relatively open anterior spaces of the nose (Cakmak et al., 2003; Hilberg et al.,
- 24 1989). However, AR tends to underestimate cross-sectional area (CSA) beyond the complex
- and sometimes obstructed turbinate region (Cakmak et al., 2005). Post-processing of CT data

- also provides additional morphological information unavailable through AR techniques, such
- 2 as airway perimeter and total airway volume.
- 3 Previous analysis of flow trajectories has identified three functional regions within the two
- 4 enantiomorphic parts of the nasal airways where nasal morphology is correlated to respiratory
- 5 function (Mlynski et al., 2001):
- 1. The anterior tract, consisting of the vestibule, isthmus and anterior cavum, which act
- as a curved nozzle-diffuser that stabilizes and redirects airflow across the turbinates.
- 8 2. The mid functional tract, consisting of a slit-like space that presents a large mucosal
- 9 surface area to the airflow.

- 3. The posterior tract, consisting of the posterior cavum, choanae and epipharynx, which
- act as a curved nozzle to stabilize and redirect airflow to the lower airways.
- Many morphological studies have been undertaken to improve the functional understanding
- of geometry as well as the complex dynamic behaviour within this part of the human airway.
- 15 These studies have resulted in many numerical models of heat and water mass transfer.
- Analysis techniques vary, but can involve direct importation of CT morphology data into
- computational fluid dynamics (CFD) software to provide airflow predictions within complex
- 3-D domains (Chen et al., 2010; Croce et al., 2006; Garcia et al., 2007; Lindemann et al.,
- 19 2004; Pless et al., 2004). Other methods use mathematical models derived from continuity
- and transport equations, which are applied to simplified anatomic based geometry to predict
- 21 airway heat and water mass transfer (Daviskas et al., 1990; Hanna and Scherer, 1986; Naftali,
- 22 1998; Tawhai, 2003; Tsu et al., 1988). In this case, the internal forced-convection
- coefficients for heat and water mass transfer are predicted by the non-dimensional Nusselt
- 24 (N_u) and Sherwood (S_h) numbers, determined from two components:

- Momentum to thermal diffusivity ratio to yield a Prandtl number (P_r) for heat transfer
 or momentum to mass diffusivity to yield a Schmidt number (S_c) for mass transfer.
- 3 2. Airflow character, such as laminar or turbulent, predicted by Reynolds number (R_e)
- 4 (Cengel, 2006).
- 5 Under conditions of constant fluid properties P_r and S_c remain unchanged meaning airflow
- 6 character effectively governs the heat and water mass transfer coefficients. Airflow regime
- 7 state is normally identified by the magnitude of the Re, which expresses the ratio of inertia to
- 8 viscous forces as:

$$R_e = VD/v \tag{1}$$

- where V and D represent the mean fluid velocity and pipe diameter respectively, and ν
- 11 represents the fluid kinematic viscosity. When considering non-circular ducts, such as those
- found within the nose, D is replaced with a hydraulic diameter (D_h) to account for interaction
- between the moving fluid and boundary surfaces (Fox and McDonald, 1994; Sabersky et al.,
- 14 1999). This parameter describes the ratio of flow sectional properties, expressed
- 15 mathematically as:

16
$$D_h=4xCSA/Perimeter$$
 (2)

18 Through use of the continuity equation, air volume flow rate (Q) can be written as:

$$Q=V \times CSA \tag{3}$$

- 20 In reality, the inhaled air physical properties do not significantly change during passage
- 21 through the nose. Given this, variation in CSA and D_b correspond to geometrical changes,
- 22 which directly influence the magnitude of R_e and hence the heat and water mass transport.

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1.2. Nasal Cycle

- 1 It is well known that 20 40 % of the population (Davis and Eccles, 2004; Hanif et al., 2000)
- 2 experience periodic congestion/decongestion of the erectile tissue in each side of the nose
- 3 (Druce, 1988; Lindemann et al., 2003). This 'nasal cycle' results in alternating patent and
- 4 congested passages within the two enantiomorphic parts of the airway for periods ranging
- from 1-7 hours (Kennedy et al., 1988). This time span is made up from combinations of
- 6 discrete ultradian periods spanning 1-1½ hours (Atanasov et al., 2003). The nasal cycle
- 7 usually goes unnoticed since the total nasal airflow resistance remains unaffected (Eccles,
- 8 1982; Wolf et al., 2004).

- 10 In this study, the nasal morphology of eight individuals was investigated using MRI
- techniques. Unlike previous work, morphological analysis was used to identify functional
- regions within the nose. To the best of our knowledge, no one has considered the distribution
- of CSA and D_h throughout the two enantiomorphic parts of the nasal airway in terms of its
- physiological significance to local heat and water flux. The difference in airflow between
- patent and congested airways was also examined with regards to the nasal cycle.

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2. MATERIALS AND METHODS

- 18 2.1. Morphology acquisition
- 19 Four male and four female healthy non-smoking participants, of mixed ethnicity and ages
- ranging from 21 51 years and with no prior history of upper airway disease, volunteered for
- 21 the study. Ethical approval was granted by The Auckland University of Technology Ethics
- 22 Committee, application number 10/121, with each participant giving informed consent. Each
- 23 participant also undertook a visual nasal internal examination to ensure the absence of
- 24 morphological abnormalities such as significantly displaced septum, mucosal inflammation
- or other nasal pathological conditions. All participants had not received any form of

- 1 medicine and had not consumed food for 2 hours, or alcohol for 24 hours, prior to
- 2 undertaking the MRI scan. They also undertook a maximal voluntary nasal ventilation test,
- 3 measured simultaneously from the nares, to determine the status of their nasal cycle
- 4 immediately prior to scanning. This test was repeated directly upon conclusion of scanning
- 5 to ascertain if changes in nasal cycle had occurred during testing. Participants were studied
- 6 in the supine position within a 3-Tesla MRI scanner (Siemens Magnetom Skyra) using a head
- 7 array coil. The region of interest within the nose spanned from the anterior isthmus to the
- 8 posterior choanae. Morphologic image acquisition used T1-weighted sagittal slices of
- 9 repetition time (TR) 700 ms, and echo time (TE) of 12.0 ms. Slice thickness was 0.78 mm,
- echo spacing 4.05 ms and duration of each scan was approximately 5 minutes.
- 12 2.2. Airway measurement

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- Post processing of acquired DICOM format MRI images was undertaken using 3 D-Doctor
- 14 TM (Able Software Corp) to define airway boundaries using 0.78 mm coronal slices orientated
- perpendicular to the floor of the nose. Nasal airway geometric properties for each participant
- were compiled and plotted using MatLab TM (MathWorks, Natick, MA, USA).

18 3. RESULTS

- 19 *3.1 Geometry comparisons*
- 20 Figure 1 shows the distribution of morphological CSA and perimeter data along the nasal
- 21 airway vs. non-dimensional airway position (X/L), where X is the distance from the vestibule
- and L is the total distance from the vestibule to posterior region of interest. Figure 1 presents
- 23 data for both patent and congested airways and demonstrates magnitudes and distribution
- within the normal physiological range (Hilberg et al., 1989; Lang et al., 2003; Yokley, 2009).
- Non-dimensional airway position is used to account for inter-participant variation in nasal

- 1 passage length. The solid curve in each of these figures represents the line of best fit for
- 2 each parameter recorded and trends change in nasal morphology along the nasal passages.
- 3 The data in Figure 1 assumes uncorrelated residuals. While this is a questionable assumption,
- 4 and indeed a lag plot shows appreciable correlation between consecutive residuals, a formal
- 5 analysis using a Gaussian process (Hankin, 2005) shows that the effect of assuming
- 6 uncorrelated residuals is likely to be negligible. There is no information to support gender as
- 7 being a significant factor in the distribution of this data.

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4. DISCUSSION AND ANALYSIS

- 10 4.1 Cross sectional area distribution
- 11 Comparison of morphological properties between patent and congested airways (Figs. 1A to
- 12 1D) shows CSA as the only parameter that varies. The distribution of CSA (Figs. 1a & 1b)
- correlates well with other data obtained by both AR and MRI techniques (Çakmak et al.,
- 2003; Cheng et al., 1996; Hilberg et al., 1989; Philip and Renato, 1996; Subramaniam et al.,
- 15 1998; Wen et al., 2008) and demonstrates the same variation occurring between patent and
- 16 congested airways (Lang et al., 2003). One participant demonstrated uncharacteristically low
- 17 CSA and perimeter results within their congested airway.

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- 4.2. Hydraulic diameter distribution
- Figure 2 shows D_h vs. X/L and indicates that D_h in both human patent and congested airways
- 21 follows a similar trend; however, the former has relatively higher values than the latter. Both
- of these compare poorly to patent and congested airways, measured using the AR techniques
- 23 (Lang et al., 2003), which ranged from 30-60 mm measured at the anterior head of the
- inferior turbinate (not shown in Figure 2). Unlike imaging techniques used in this research,
- 25 rhinoresistometry measures flow and pressure in order to calculate D_h which may account for

- this poor correlation. For comparison purposes, results previously obtained from human
- 2 cadaver mouldings (Hanna, 1983) are shown in Figure 2. These demonstrate slightly higher
- 3 values than the data obtained in this research which is probably due to unavoidable tissue
- 4 shrinkage and the absence of perfusion within cadaveric erectile tissue. D_h values measured
- 5 using MRI in live canine nasal airways (Craven et al., 2007) (also shown in Figure 2) over
- 6 the same region demonstrate similar results, both in terms of magnitude and distribution.
- 7 These close results may be attributed to both human and canine airways having similar
- 8 heating and humidifying functional requirements using similar tidal volumes.

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- 4.3. Functional regions
- Examination of the D_h distribution along the non-dimensional airway (Figure 2) suggests the
- presence of three distinct function regions (Mlynski et al., 2001). The first 20% (X/L=0.2) of
- the anterior region of the airway is characterized by a reduction in which corresponds to a
- progressive narrowing of CSA. The mid functional portion, covering a further 60% of the
- airway is characterized by a constant minimum D_h value. Over this region, fluctuation in
- 16 CSA (Figs. 1 A & 1B) is counterbalanced by change in perimeter (Figs. 1 C & D). The
- posterior portion, commencing approximately 80 % (X/L=0.8) into the airway, is
- characterized by a rapid increase in D_h.

- 20 4.4 Heat and water flux
- 21 Assuming constant air physical properties identifies R_e as playing a major role in regulating
- 22 the heat and mass transfer processes within the nose. Figure 3A shows the distribution of R_e
- 23 vs. X/L. There is a significant reduction in R_e magnitude within the mid functional region of
- both patent and congested airways. This indicates a reduction in flow velocity in this region

- 1 which favours olfaction function and particulate deposition as well as an increase in the
- 2 duration of time air is exposed to the mucosal surfaces.
- 3 Figure 3B shows the net heat and water flux (N-Flux $_{qw}$) vs. X/L. This variable, derived from
- 4 R_e and airway surface area, effectively quantifies the total heat and water flux distribution
- 5 along the nasal airway. Cool and dry air potentially provides the greatest air-conditioning
- 6 challenges to the nasal mucosa by placing a disproportionate heat and water mass flux burden
- 7 on the anterior airway. To overcome this 'shock' effect, it appears that the initial anterior
- 8 region, covering the transition between the skin at the nares and middle airway epithelia
- 9 (X/L=0 0.20), has reduced N-Flux_{qw}. During tidal breathing, cyclic exchanges of heat and
- water need to be distributed evenly along the mucosal surface to balance out tissue loadings.
- 11 A constant N-Flux $_{qw}$ value within the mid functional region of the patent airway must
- contribute to achieving this under conditions of alternating airflow. This region covers the
- majority of patent airway length (X/L=0.20-0.80) and results in progressive heating and
- 14 humidifying of inhaled air as it traverses the length of the nasal passage. Of note are the
- 15 fluctuating suppressed N-Flux_{qw} values for congested airways. The posterior regions in both
- patent and congested airways are characterised by an increase in N-Flux_{qw} which would
- ensure further heat and water flux occurs within this locale.
- 18 4.5 Mucocillary clearance
- 19 Periodic congestion/decongestion of the nasal airways has been shown previously to assist in
- 20 purging contaminants entrapped within the mucus layer of the congested airway (Soane et al.,
- 21 2001). In our investigation, reduction in N-Flux_{qw} occurred at two locations within the mid
- functional portion of the congested airway (Fig 3B). Although the purpose of the nasal cycle
- 23 is currently not fully understood, it is thought to play a role in balancing mucosal heat and
- 24 water fluxes (Elad et al., 2008), as well as allowing the cells on the congested side to rest and
- recharge (Eccles, 1982). This reduction in N-Flux_{qw} coincides with reduced air flow in the

- 1 congested airway and should result in no net change in air-conditioning of the combined
- 2 airways. Earlier research has found that the degree of water saturation did not correlate to
- any position along the airway or time during the nasal cycle (Lindemann et al., 2003).
- 4 Reduced heat and water demand within the congested airway leads to replenishment of
- 5 airway fluid levels and additional mucus hydration, both of which have been shown to
- 6 improve MTV (Kilgour et al., 2004). Further, diminished airflow within the congested
- 7 airway also leads to a reduction in particulate and other airborne pollutant deposition rates
- 8 into the sticky mucus layer, further aiding mucocillary clearance. Since MTV is
- 9 predominantly regulated by purinergic mechanisms (Braiman et al., 2000; Bucheimer and
- Linden, 2004; Button and Boucher, 2008), mucus transport acts independently of the nasal
- 11 cycle.

13

CONCLUSIONS

- 14 This investigation has identified three functional regions occurring within the nasal airways.
- 15 The anterior functional region of both patent and congested airways demonstrates a reduction
- in N-Flux_{aw} that prevents local stressing of underlying tissue heat and water supply. The mid
- functional regions of the two enantiomorphic nasal airways demonstrate a reduction in R_e
- which favours olfactory function and particulate deposition by increasing the time of air is
- 19 exposed to the mucosal surfaces. This functional region of the patent airway also experiences
- 20 constant N-Flux_{qw} which balances tissue heat and water supply during the period of greatest
- 21 demand. The posterior region of both airways experience increases in N-Flux_{qw} in order to
- 22 achieve further air-conditioning despite reduced driving gradients. Finally, reduced N-Flux_{aw}
- 23 occurs within the congested airway which supports earlier findings of enhanced mucus
- 24 clearance occurring within this passage.

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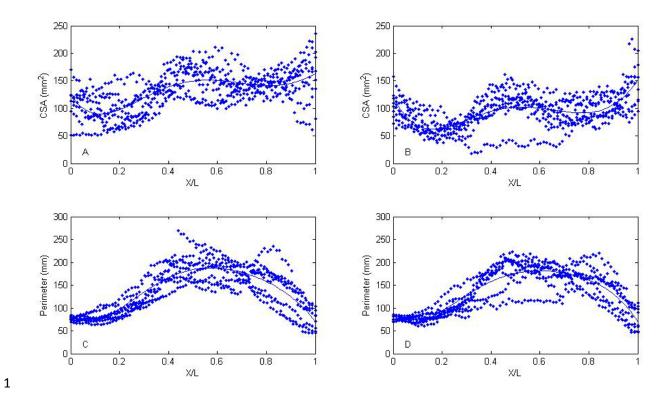
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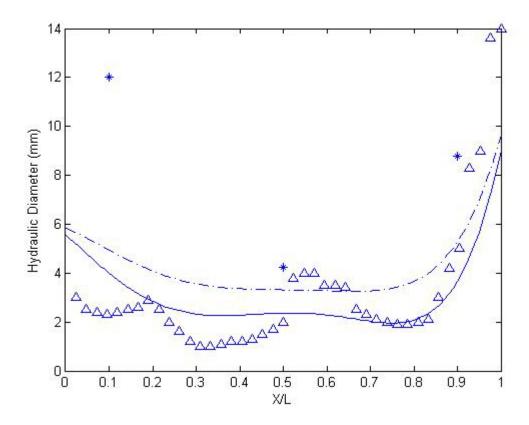
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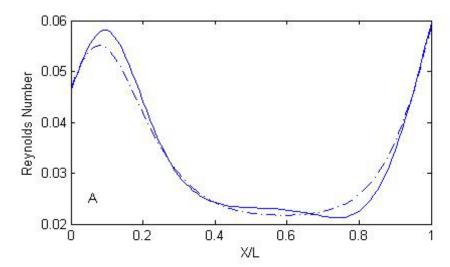
2 Figure 1: Distribution of participant morphological parameters. (A) Patent airway cross-

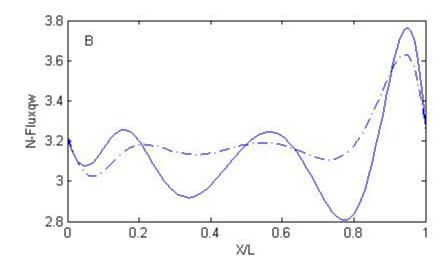
- 3 sectional area. (B) Congested airway cross-sectional area. (C) Patent airway perimeter. (D)
- 4 Congested airway perimeter.



2 Figure 2: Human and canine airway hydraulic diameter distribution: — — human patent

- 3 airway (MRI), human congested airway (MRI), △ canine airway (Craven et al. (2007),
- 4 * human cadaver (Hanna, 1983).





4 Figure 3: Distribution of airway heat and mass transfer components. (A) Reynolds number.

(B) Net heat and water flux coefficients. — — human patent airway, ——— human congested

6 airway.